

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 --> 00:00:08.724 So thanks for waiting.

NOTE Confidence: 0.920264657142857

00:00:08.724 --> 00:00:12.775 And so I'll you know shorten this a

NOTE Confidence: 0.920264657142857

00:00:12.775 --> 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 --> 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 --> 00:00:28.360 a little time for questions so.

NOTE Confidence: 0.9402536

00:00:35.160 --> 00:00:36.880 Let's see. No,

NOTE Confidence: 0.9301902

00:00:36.880 --> 00:00:38.175 you have to do it on the
NOTE Confidence: 0.9301902

00:00:38.175 --> 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 --> 00:00:53.580 so we're gonna blaze through the historical.
NOTE Confidence: 0.919390781818182

00:00:53.580 --> 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 --> 00:01:02.030 majority of the time then presenting
NOTE Confidence: 0.919390781818182

00:01:02.030 --> 00:01:05.480 some hopefully interesting secondary
NOTE Confidence: 0.919390781818182

00:01:05.480 --> 00:01:08.172 analyses we've done with the data
NOTE Confidence: 0.919390781818182

00:01:08.172 --> 00:01:12.060 from the completed randomized trial.
NOTE Confidence: 0.919390781818182

00:01:12.060 --> 00:01:17.338 So just this is the sort of kube
NOTE Confidence: 0.919390781818182

00:01:17.338 --> 00:01:19.820 bolckoff model of alcohol use disorder
NOTE Confidence: 0.919390781818182

00:01:19.820 --> 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 --> 00:01:27.698 we can argue about whether these

NOTE Confidence: 0.919390781818182
00:01:27.698 --> 00:01:29.455 are really separate domains,
NOTE Confidence: 0.919390781818182
00:01:29.455 --> 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 --> 00:01:39.963 negative negative emotionality,
NOTE Confidence: 0.919390781818182
00:01:39.963 --> 00:01:43.847 and executive function problems.
NOTE Confidence: 0.919390781818182
00:01:43.850 --> 00:01:48.794 And we'll come back to this later looking
NOTE Confidence: 0.919390781818182
00:01:48.794 --> 00:01:52.046 at the data I was going to for for.
NOTE Confidence: 0.919390781818182
00:01:52.050 --> 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 --> 00:01:59.310 I was going to say a little
NOTE Confidence: 0.919390781818182
00:01:59.310 --> 00:02:00.610 bit about the how the
NOTE Confidence: 0.950317
00:02:02.970 --> 00:02:04.470 ideal paradigm of translational science
NOTE Confidence: 0.950317
00:02:04.470 --> 00:02:07.530 is not really what we've been doing here.
NOTE Confidence: 0.950317
00:02:07.530 --> 00:02:10.122 We've been working with things that
NOTE Confidence: 0.950317
00:02:10.122 --> 00:02:12.203 were serendipitously discovered and kind
NOTE Confidence: 0.950317

00:02:12.203 --> 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 --> 00:02:17.430 worth spending much time on.

NOTE Confidence: 0.950317

00:02:17.430 --> 00:02:19.662 And I think everybody probably also

NOTE Confidence: 0.950317

00:02:19.662 --> 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 --> 00:02:24.630 kinds of psychedelics and so forth.

NOTE Confidence: 0.950317

00:02:24.630 --> 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 --> 00:02:32.790 but other relatively similar,

NOTE Confidence: 0.950317

00:02:32.790 --> 00:02:35.750 mostly 5HT2A agonist or partial

NOTE Confidence: 0.950317

00:02:35.750 --> 00:02:38.790 agonist type drugs in the in this talk

NOTE Confidence: 0.91525645

00:02:41.870 --> 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 --> 00:02:49.976 You know, there was a lot of research and

NOTE Confidence: 0.91525645

00:02:49.976 --> 00:02:52.720 a lot of interest in classic psychedelics,

NOTE Confidence: 0.91525645

00:02:52.720 --> 00:02:55.555 both as a model of psychosis and

NOTE Confidence: 0.91525645

00:02:55.555 --> 00:02:57.722 then quickly people who became

NOTE Confidence: 0.91525645

00:02:57.722 --> 00:03:01.694 interested in the potential therapeutic

NOTE Confidence: 0.91525645

00:03:01.694 --> 00:03:04.830 uses of classic psychedelics.

NOTE Confidence: 0.91525645

00:03:04.830 --> 00:03:07.915 And then that really shot

NOTE Confidence: 0.91525645

00:03:07.915 --> 00:03:10.973 shut down around 1970.

NOTE Confidence: 0.91525645

00:03:10.973 --> 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 --> 00:03:22.618 use disorder and six randomized trials

NOTE Confidence: 0.91525645

00:03:22.618 --> 00:03:25.764 which was which I'll I'll get back to

NOTE Confidence: 0.91525645

00:03:25.764 --> 00:03:28.400 it in a moment showing you some the

NOTE Confidence: 0.91525645

00:03:28.400 --> 00:03:30.140 meta analysis that was done of those.

NOTE Confidence: 0.91525645

00:03:30.140 --> 00:03:33.900 But so now we're in the second wave

NOTE Confidence: 0.91525645

00:03:33.900 --> 00:03:37.260 and this is this goes up to 2020,

NOTE Confidence: 0.91525645

00:03:37.260 --> 00:03:40.106 we'd probably be up to the ceiling.

NOTE Confidence: 0.91525645

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 --> 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 --> 00:03:54.840 relative paucity of real useful data
NOTE Confidence: 0.91525645

00:03:54.840 --> 00:03:57.128 their clinical or mechanistic although
NOTE Confidence: 0.91525645

00:03:57.128 --> 00:03:58.836 that is that is starting to change.
NOTE Confidence: 0.91525645

00:03:58.840 --> 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 --> 00:04:10.590 disorder and other addictions to some extent.
NOTE Confidence: 0.91525645

00:04:10.590 --> 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 --> 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 --> 00:04:23.949 LSD in the treatment of alcohol use disorder.
NOTE Confidence: 0.91525645

00:04:23.950 --> 00:04:28.120 And these were they were all

NOTE Confidence: 0.91525645

00:04:28.120 --> 00:04:30.070 double-blind up to Drug Administration.

NOTE Confidence: 0.91525645

00:04:30.070 --> 00:04:34.798 And then sometimes people were knew that

NOTE Confidence: 0.91525645

00:04:34.798 --> 00:04:39.810 they were getting no drugs for example, but.

NOTE Confidence: 0.91525645

00:04:39.810 --> 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 --> 00:04:47.870 high doses of LSD,

NOTE Confidence: 0.91525645

00:04:47.870 --> 00:04:49.290 up to 800 micrograms,

NOTE Confidence: 0.91525645

00:04:49.290 --> 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 --> 00:04:57.130 inactive control conditions.

NOTE Confidence: 0.91525645

00:04:57.130 --> 00:05:00.460 And the psychotherapeutic models range

NOTE Confidence: 0.91525645

00:05:00.460 --> 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 --> 00:05:09.420 psychotherapy afterwards as well.

NOTE Confidence: 0.91525645

00:05:09.420 --> 00:05:12.185 So the the main finding of this
NOTE Confidence: 0.91525645

00:05:12.185 --> 00:05:15.448 was that there was a,
NOTE Confidence: 0.91525645

00:05:15.448 --> 00:05:18.780 you know a robust and very consistent
NOTE Confidence: 0.91525645

00:05:18.780 --> 00:05:20.600 effect across these trials.
NOTE Confidence: 0.91525645

00:05:20.600 --> 00:05:23.330 It was really very little evidence
NOTE Confidence: 0.91525645

00:05:23.409 --> 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 --> 00:05:31.047 spite of all the variability and the
NOTE Confidence: 0.91525645

00:05:31.047 --> 00:05:35.240 overall 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 --> 00:05:41.160 first available follow up time point.
NOTE Confidence: 0.91525645

00:05:41.160 --> 00:05:43.190 And those effects appeared to
NOTE Confidence: 0.91525645

00:05:43.190 --> 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 --> 00:05:50.063 but every single one of those

NOTE Confidence: 0.947441742857143

00:05:50.063 --> 00:05:50.999 trials technically failed.

NOTE Confidence: 0.947441742857143

00:05:51.000 --> 00:05:52.918 Yes one of them was like on

NOTE Confidence: 0.94629164

00:05:52.920 --> 00:05:54.800 the border, but that's right.

NOTE Confidence: 0.94629164

00:05:54.800 --> 00:05:57.604 And they were all under powered and they all,

NOTE Confidence: 0.94629164

00:05:57.604 --> 00:05:58.878 but they all look exactly the same.

NOTE Confidence: 0.94629164

00:05:58.880 --> 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 --> 00:06:06.595 It's just. This is a good lesson

NOTE Confidence: 0.94629164

00:06:06.595 --> 00:06:08.500 in not doing underpowered trials

NOTE Confidence: 0.94629164

00:06:08.572 --> 00:06:11.026 unless you want to convince people

NOTE Confidence: 0.94629164

00:06:11.026 --> 00:06:13.090 of something that isn't true.

NOTE Confidence: 0.94629164

00:06:13.090 --> 00:06:15.334 So there was one controlled trial

NOTE Confidence: 0.94629164

00:06:15.334 --> 00:06:18.005 that maybe is less well known of

NOTE Confidence: 0.94629164

00:06:18.005 --> 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 --> 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 --> 00:06:32.460 very large and these people,
NOTE Confidence: 0.94629164

00:06:32.460 --> 00:06:33.990 these were all.
NOTE Confidence: 0.94629164

00:06:33.990 --> 00:06:35.350 People who were on,
NOTE Confidence: 0.773826372857143

00:06:37.510 --> 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 --> 00:06:44.649 they would either go out and be on
NOTE Confidence: 0.773826372857143

00:06:44.649 --> 00:06:47.368 parole and or they would stay in the
NOTE Confidence: 0.773826372857143

00:06:47.368 --> 00:06:49.485 hospital and they'd get this preparation
NOTE Confidence: 0.773826372857143

00:06:49.485 --> 00:06:51.550 and then a high dose LSD session
NOTE Confidence: 0.773826372857143

00:06:51.550 --> 00:06:53.708 and then go to the aftercare clinic.
NOTE Confidence: 0.773826372857143

00:06:53.710 --> 00:06:57.238 And then both groups were followed for a year
NOTE Confidence: 0.773826372857143

00:06:57.238 --> 00:07:01.296 and part of their parole was daily urine.
NOTE Confidence: 0.773826372857143

00:07:01.296 --> 00:07:02.620 Drug monitoring.
NOTE Confidence: 0.773826372857143

00:07:02.620 --> 00:07:04.720 So if the outcome data is is

NOTE Confidence: 0.773826372857143

00:07:04.720 --> 00:07:06.580 is certainly of high quality.

NOTE Confidence: 0.773826372857143

00:07:06.580 --> 00:07:13.017 And so the the very kind of low

NOTE Confidence: 0.773826372857143

00:07:13.017 --> 00:07:15.530 tech figure on the right there from

NOTE Confidence: 0.773826372857143

00:07:15.608 --> 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 --> 00:07:26.482 rates at 369 and 12 months and you

NOTE Confidence: 0.773826372857143

00:07:26.482 --> 00:07:28.886 can see there's a a real pretty.

NOTE Confidence: 0.773826372857143

00:07:28.886 --> 00:07:30.816 Robust separation starting as early

NOTE Confidence: 0.773826372857143

00:07:30.816 --> 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 --> 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 --> 00:07:44.225 probably what you would have expected.

NOTE Confidence: 0.773826372857143

00:07:44.230 --> 00:07:47.006 The 5% that is in people with you

NOTE Confidence: 0.773826372857143

00:07:47.006 --> 00:07:49.890 know this was they weren't getting
NOTE Confidence: 0.773826372857143

00:07:49.890 --> 00:07:52.365 methadone and they were just out in the
NOTE Confidence: 0.773826372857143

00:07:52.365 --> 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 --> 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 --> 00:08:03.766 you otherwise would have seen.
NOTE Confidence: 0.773826372857143

00:08:03.770 --> 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 --> 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 --> 00:08:15.335 first study that was published
NOTE Confidence: 0.773826372857143

00:08:15.335 --> 00:08:19.760 of Psychedelic for Addiction in
NOTE Confidence: 0.773826372857143

00:08:19.760 --> 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 --> 00:08:27.790 open label study, 15 participants.

NOTE Confidence: 0.773826372857143

00:08:27.790 --> 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 --> 00:08:37.942 remember exactly the dose,

NOTE Confidence: 0.939247154

00:08:37.942 --> 00:08:39.790 but it was a high dose,

NOTE Confidence: 0.939247154

00:08:39.790 --> 00:08:46.270 25 milligrams or so. And the people,

NOTE Confidence: 0.939247154

00:08:46.270 --> 00:08:48.223 in fact some of them got a third dose.

NOTE Confidence: 0.939247154

00:08:48.230 --> 00:08:49.838 And So what you see here

NOTE Confidence: 0.939247154

00:08:49.838 --> 00:08:50.910 is cigarettes per day.

NOTE Confidence: 0.939247154

00:08:50.910 --> 00:08:54.778 So this is a continuous outcome, very large.

NOTE Confidence: 0.939247154

00:08:54.778 --> 00:08:59.040 Effect at the six months and the

NOTE Confidence: 0.939247154

00:08:59.040 --> 00:09:01.160 majority of the participants,

NOTE Confidence: 0.939247154

00:09:01.160 --> 00:09:03.278 80% of them in fact were

NOTE Confidence: 0.939247154

00:09:03.280 --> 00:09:06.320 abstinent at that point.

NOTE Confidence: 0.939247154

00:09:06.320 --> 00:09:09.071 They did some longer term follow-ups up

NOTE Confidence: 0.939247154

00:09:09.071 --> 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 --> 00:09:19.332 And found that nine out of 15 of them

NOTE Confidence: 0.939247154

00:09:19.332 --> 00:09:21.515 still at 60% were still abstinent.

NOTE Confidence: 0.939247154

00:09:21.515 --> 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 --> 00:09:27.372 small open label study.

NOTE Confidence: 0.939247154

00:09:27.372 --> 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 --> 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 --> 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 --> 00:09:50.160 open label RCT and the results look

NOTE Confidence: 0.939247154

00:09:50.160 --> 00:09:52.380 not quite as striking as this,

NOTE Confidence: 0.939247154

00:09:52.380 --> 00:09:55.080 but still highly significant in the

NOTE Confidence: 0.939247154

00:09:55.080 --> 00:09:57.300 interim analyses that they've done.

NOTE Confidence: 0.939247154

00:09:57.300 --> 00:10:01.660 And we're that's that's at Johns Hopkins,

NOTE Confidence: 0.939247154

00:10:01.660 --> 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 --> 00:10:15.593 Matt Johnson and NYU are doing a

NOTE Confidence: 0.939247154

00:10:15.593 --> 00:10:18.429 three site night of funded study now,

NOTE Confidence: 0.939247154

00:10:18.430 --> 00:10:22.576 which is a placebo control for

NOTE Confidence: 0.939247154

00:10:22.576 --> 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 --> 00:10:28.750 so that'll be the first double-blind money.

NOTE Confidence: 0.9402536

00:10:30.870 --> 00:10:33.588 So that's that. So alcohol now.

NOTE Confidence: 0.9402536

00:10:33.590 --> 00:10:35.550 So this was published a year later.

NOTE Confidence: 0.9402536

00:10:35.550 --> 00:10:38.076 This is the pilot study that.

NOTE Confidence: 0.9402536

00:10:38.080 --> 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,

NOTE Confidence: 0.9402536

00:10:42.760 --> 00:10:43.699 so very similar,
NOTE Confidence: 0.9402536
00:10:43.699 --> 00:10:46.320 kind of designed to the the smoking study.
NOTE Confidence: 0.9402536
00:10:46.320 --> 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 --> 00:10:55.648 but we did look at the drinking
NOTE Confidence: 0.9402536
00:10:55.648 --> 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 --> 00:11:01.560 So for anybody that's not
NOTE Confidence: 0.9402536
00:11:01.560 --> 00:11:02.712 in the alcohol field,
NOTE Confidence: 0.9402536
00:11:02.720 --> 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 --> 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 --> 00:11:15.215 roughly it's .6 of an ounce of
NOTE Confidence: 0.9402536
00:11:15.215 --> 00:11:16.865 absolute ethanol and it's about

NOTE Confidence: 0.9402536

00:11:16.865 --> 00:11:19.728 a beer or a glass of wine or a

NOTE Confidence: 0.9402536

00:11:19.728 --> 00:11:23.453 shot of 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 --> 00:11:31.272 sessions happened at weeks four

NOTE Confidence: 0.9402536

00:11:31.272 --> 00:11:34.296 and eight after the second and

NOTE Confidence: 0.9402536

00:11:34.296 --> 00:11:36.760 third time point on the graph.

NOTE Confidence: 0.9402536

00:11:36.760 --> 00:11:39.364 So you can see there's some

NOTE Confidence: 0.9402536

00:11:39.364 --> 00:11:41.168 improvement between baseline with

NOTE Confidence: 0.9402536

00:11:41.168 --> 00:11:44.360 the first time point in week four.

NOTE Confidence: 0.9402536

00:11:44.360 --> 00:11:45.932 That's while people were

NOTE Confidence: 0.9402536

00:11:45.932 --> 00:11:47.277 receiving some psychotherapy,

NOTE Confidence: 0.9402536

00:11:47.277 --> 00:11:50.619 which was, part of which was.

NOTE Confidence: 0.9402536

00:11:50.620 --> 00:11:53.204 Motivational interviewing and and

NOTE Confidence: 0.9402536

00:11:53.204 --> 00:11:55.148 cognitive behavioral skills training.

NOTE Confidence: 0.9402536

00:11:55.148 --> 00:11:56.768 And so they were getting
NOTE Confidence: 0.9402536

00:11:56.768 --> 00:11:57.740 some active treatment.
NOTE Confidence: 0.9402536

00:11:57.740 --> 00:11:59.472 There was some improvement,
NOTE Confidence: 0.9402536

00:11:59.472 --> 00:12:00.338 not surprisingly,
NOTE Confidence: 0.9402536

00:12:00.340 --> 00:12:03.380 but after week four,
NOTE Confidence: 0.9402536

00:12:03.380 --> 00:12:05.180 the second month after the
NOTE Confidence: 0.9402536

00:12:05.180 --> 00:12:06.260 first psilocybin session,
NOTE Confidence: 0.9402536

00:12:06.260 --> 00:12:08.836 there was a marked reduction in
NOTE Confidence: 0.9402536

00:12:08.836 --> 00:12:11.688 heavy drinking days and the second
NOTE Confidence: 0.9402536

00:12:11.688 --> 00:12:13.458 psilocybin session at week eight.
NOTE Confidence: 0.9402536

00:12:13.460 --> 00:12:15.860 And those games were basically
NOTE Confidence: 0.9402536

00:12:15.860 --> 00:12:20.369 maintained out to week 36, which was.
NOTE Confidence: 0.9402536

00:12:20.370 --> 00:12:24.409 28 weeks after the second psilocybin session.
NOTE Confidence: 0.9402536

00:12:24.410 --> 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 --> 00:12:31.495 baseline and the the week four,

NOTE Confidence: 0.9402536

00:12:31.495 --> 00:12:34.689 which is sort of the real baseline because

NOTE Confidence: 0.9402536

00:12:34.689 --> 00:12:37.410 that's the immediate pre psilocybin

NOTE Confidence: 0.886931566666667

00:12:39.490 --> 00:12:43.570 baseline and the effect sizes are

NOTE Confidence: 0.886931566666667

00:12:43.570 --> 00:12:46.218 you know on the range of .8 to.

NOTE Confidence: 0.886931566666667

00:12:46.220 --> 00:12:48.215 To 1.0 or a little bit more.

NOTE Confidence: 0.886931566666667

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

NOTE Confidence: 0.886931566666667

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

NOTE Confidence: 0.94312879

00:12:51.140 --> 00:12:53.100 dose was after that third data point.

NOTE Confidence: 0.94312879

00:12:53.100 --> 00:12:55.252 So there's no further, there is no further

NOTE Confidence: 0.94312879

00:12:55.252 --> 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 --> 00:13:02.260 yeah. And you know and it might have,

NOTE Confidence: 0.93522184625

00:13:02.260 --> 00:13:03.835 you know the effect might have not

NOTE Confidence: 0.93522184625

00:13:03.835 --> 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,

NOTE Confidence: 0.93522184625

00:13:07.660 --> 00:13:09.080 but that's a big question
NOTE Confidence: 0.93522184625

00:13:09.080 --> 00:13:10.500 too is how many sessions.
NOTE Confidence: 0.93522184625

00:13:10.500 --> 00:13:12.116 So this is the design of the completed
NOTE Confidence: 0.93522184625

00:13:12.116 --> 00:13:13.579 trial and this has been published.
NOTE Confidence: 0.93522184625

00:13:13.580 --> 00:13:15.362 I don't want to spend a lot of time.
NOTE Confidence: 0.93522184625

00:13:15.370 --> 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 --> 00:13:21.186 the way a lot of these
NOTE Confidence: 0.93522184625

00:13:21.186 --> 00:13:21.970 studies have been done,
NOTE Confidence: 0.93522184625

00:13:21.970 --> 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: 0.9519942333333333

00:13:27.930 --> 00:13:29.930 manualized psychotherapy before the first
NOTE Confidence: 0.9519942333333333

00:13:29.930 --> 00:13:32.929 one in between the two and afterwards.
NOTE Confidence: 0.9519942333333333

00:13:32.930 --> 00:13:37.670 And these are a combination of content
NOTE Confidence: 0.9519942333333333

00:13:37.670 --> 00:13:41.004 that's intended to help people
NOTE Confidence: 0.9519942333333333

00:13:41.004 --> 00:13:44.706 get prepared for the sessions and.

NOTE Confidence: 0.9519942333333333
00:13:44.710 --> 00:13:48.278 Be able to manage the intense subjective
NOTE Confidence: 0.9519942333333333
00:13:48.278 --> 00:13:52.550 effects and to hopefully figure out
NOTE Confidence: 0.9519942333333333
00:13:52.550 --> 00:13:56.110 how to make use of the experience.
NOTE Confidence: 0.9519942333333333
00:13:56.110 --> 00:13:58.952 And so that's the sort of psychedelic
NOTE Confidence: 0.9519942333333333
00:13:58.952 --> 00:14:01.970 part of the treatment and the other
NOTE Confidence: 0.9519942333333333
00:14:01.970 --> 00:14:04.850 part is evidence based treatment for
NOTE Confidence: 0.9519942333333333
00:14:04.850 --> 00:14:07.880 alcohol use disorder to help them.
NOTE Confidence: 0.9519942333333333
00:14:07.880 --> 00:14:09.680 Actually make some behavior changes.
NOTE Confidence: 0.9519942333333333
00:14:09.680 --> 00:14:12.784 So in this case again it was combination
NOTE Confidence: 0.9519942333333333
00:14:12.784 --> 00:14:15.855 of of motivational enhancement therapy
NOTE Confidence: 0.9519942333333333
00:14:15.855 --> 00:14:20.700 and some very brief cognitive behavioral
NOTE Confidence: 0.9519942333333333
00:14:20.700 --> 00:14:24.908 skills based therapy and so the.
NOTE Confidence: 0.9519942333333333
00:14:24.908 --> 00:14:28.550 We followed them in the double-blind
NOTE Confidence: 0.9519942333333333
00:14:28.663 --> 00:14:30.287 out to week 36.
NOTE Confidence: 0.9519942333333333
00:14:30.290 --> 00:14:32.222 There was an open label extension phase
NOTE Confidence: 0.9519942333333333

00:14:32.222 --> 00:14:34.167 that I'm not going to talk about,
NOTE Confidence: 0.9519942333333333

00:14:34.170 --> 00:14:36.200 but here's the primary outcome from the
NOTE Confidence: 0.9519942333333333

00:14:36.200 --> 00:14:38.529 papers which was percent heavy drinking day.
NOTE Confidence: 0.9519942333333333

00:14:38.530 --> 00:14:41.122 So same thing we were looking at in the
NOTE Confidence: 0.9519942333333333

00:14:41.122 --> 00:14:45.169 pilot and you can see very similar curves.
NOTE Confidence: 0.9519942333333333

00:14:45.170 --> 00:14:47.735 The psilocybin is going to be in red and
NOTE Confidence: 0.9519942333333333

00:14:47.735 --> 00:14:50.678 the control in blue and all of these slides.
NOTE Confidence: 0.9519942333333333

00:14:50.678 --> 00:14:53.726 So the again there's a big.
NOTE Confidence: 0.9519942333333333

00:14:53.730 --> 00:14:55.030 Decrease during those first
NOTE Confidence: 0.9519942333333333

00:14:55.030 --> 00:14:56.330 four weeks of therapy,
NOTE Confidence: 0.9519942333333333

00:14:56.330 --> 00:15:00.160 but then another large increase in the well,
NOTE Confidence: 0.9519942333333333

00:15:00.160 --> 00:15:02.740 decrease in drinking in the psilocybin
NOTE Confidence: 0.9519942333333333

00:15:02.740 --> 00:15:05.890 group after the first medication session,
NOTE Confidence: 0.9519942333333333

00:15:05.890 --> 00:15:08.930 which then is for the most part maintained.
NOTE Confidence: 0.9519942333333333

00:15:08.930 --> 00:15:12.636 I mean we can you know it looks just that,
NOTE Confidence: 0.9519942333333333

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

NOTE Confidence: 0.9519942333333333

00:15:13.488 --> 00:15:16.210 there may be some attenuation of the effect.

NOTE Confidence: 0.9519942333333333

00:15:16.210 --> 00:15:20.718 There really wasn't a time effect.

NOTE Confidence: 0.9519942333333333

00:15:20.720 --> 00:15:22.309 You know once you take the baseline

NOTE Confidence: 0.9519942333333333

00:15:22.309 --> 00:15:25.220 out of this that that was significant

NOTE Confidence: 0.9519942333333333

00:15:25.220 --> 00:15:27.596 but or a group by time interaction.

NOTE Confidence: 0.9519942333333333

00:15:27.600 --> 00:15:31.660 So they weren't the the separation wasn't

NOTE Confidence: 0.9519942333333333

00:15:31.660 --> 00:15:35.400 diminishing to a significant extent.

NOTE Confidence: 0.9519942333333333

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

NOTE Confidence: 0.9519942333333333

00:15:37.375 --> 00:15:39.085 you know maybe the treatment effects

NOTE Confidence: 0.9519942333333333

00:15:39.085 --> 00:15:40.919 wearing off a little bit but maybe not.

NOTE Confidence: 0.9519942333333333

00:15:40.920 --> 00:15:44.580 Yeah, Hedges G for this outcome was .52.

NOTE Confidence: 0.9519942333333333

00:15:44.580 --> 00:15:46.960 So it's a solid medium sized effect

NOTE Confidence: 0.9519942333333333

00:15:46.960 --> 00:15:49.519 which is much better than you would

NOTE Confidence: 0.9519942333333333

00:15:49.520 --> 00:15:52.599 ever see with you know something

NOTE Confidence: 0.9519942333333333

00:15:52.599 --> 00:15:55.008 like naltrexone or a camper say so.

NOTE Confidence: 0.9519942333333333

00:15:55.008 --> 00:15:56.640 So that was encouraging.
NOTE Confidence: 0.9519942333333333

00:15:56.640 --> 00:15:59.000 Drinks per day is another.
NOTE Confidence: 0.9519942333333333

00:15:59.000 --> 00:16:02.200 This is just total quantity of
NOTE Confidence: 0.9519942333333333

00:16:02.200 --> 00:16:05.320 alcohol divided by the number of
NOTE Confidence: 0.9519942333333333

00:16:05.320 --> 00:16:08.686 days and so it looks very similar.
NOTE Confidence: 0.9519942333333333

00:16:08.690 --> 00:16:11.514 And you know and that was also significant
NOTE Confidence: 0.9519942333333333

00:16:11.514 --> 00:16:13.887 over the entire follow up period
NOTE Confidence: 0.9335446666666667

00:16:16.170 --> 00:16:18.360 and these are some of the dichotomous
NOTE Confidence: 0.9335446666666667

00:16:18.360 --> 00:16:19.970 outcomes we looked at and I don't
NOTE Confidence: 0.9335446666666667

00:16:19.970 --> 00:16:21.686 want to spend too much time on this,
NOTE Confidence: 0.9335446666666667

00:16:21.690 --> 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 --> 00:16:27.970 this is the last week of follow up.
NOTE Confidence: 0.9335446666666667

00:16:27.970 --> 00:16:28.810 So that would be
NOTE Confidence: 0.9402536

00:16:31.170 --> 00:16:35.660 7 to 8. Well, seven months after
NOTE Confidence: 0.9402536

00:16:35.660 --> 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

NOTE Confidence: 0.9402536

00:16:41.865 --> 00:16:45.080 of the psilocybin patients versus 24% of

NOTE Confidence: 0.9402536

00:16:45.080 --> 00:16:47.595 the control participants were abstinent,

NOTE Confidence: 0.9402536

00:16:47.595 --> 00:16:50.120 completely abstinent for that month.

NOTE Confidence: 0.9402536

00:16:50.120 --> 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 --> 00:16:58.650 which is you know, it doesn't doesn't

NOTE Confidence: 0.9402536

00:16:58.650 --> 00:16:59.800 mean they're in full remission,

NOTE Confidence: 0.9402536

00:16:59.800 --> 00:17:03.328 but it's you know they're they're not having.

NOTE Confidence: 0.9402536

00:17:03.330 --> 00:17:05.290 They're not doing binge drinking at least.

NOTE Confidence: 0.9402536

00:17:05.290 --> 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 --> 00:17:13.570 and problems related to alcohol.

NOTE Confidence: 0.9402536

00:17:13.570 --> 00:17:15.058 This is the SIP short inventory

NOTE Confidence: 0.9402536

00:17:15.058 --> 00:17:16.644 of problems that you know covers

NOTE Confidence: 0.9402536

00:17:16.644 --> 00:17:18.004 a number of different areas.

NOTE Confidence: 0.9402536

00:17:18.010 --> 00:17:20.215 So that was just showing that there

NOTE Confidence: 0.9402536

00:17:20.215 --> 00:17:21.476 was functional significance to

NOTE Confidence: 0.9402536

00:17:21.476 --> 00:17:24.178 these changes as well much larger

NOTE Confidence: 0.9402536

00:17:24.178 --> 00:17:28.076 decrease in the psilocybin group so.

NOTE Confidence: 0.9402536

00:17:28.076 --> 00:17:30.204 Here's one thing that is not in the

NOTE Confidence: 0.9402536

00:17:30.204 --> 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 --> 00:17:36.558 It's just in terms of outcomes.

NOTE Confidence: 0.9402536

00:17:36.560 --> 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 --> 00:17:47.297 did it matter whether people had already

NOTE Confidence: 0.9402536

00:17:47.297 --> 00:17:49.741 stopped or cut down on their drinking

NOTE Confidence: 0.9402536

00:17:49.741 --> 00:17:51.913 before they received the sill assignment?

NOTE Confidence: 0.9402536

00:17:51.920 --> 00:17:53.390 Did you know in terms of what

NOTE Confidence: 0.9402536

00:17:53.390 --> 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 --> 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 --> 00:18:06.935 so that would be WHO risk level

NOTE Confidence: 0.9402536

00:18:06.935 --> 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 --> 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 --> 00:18:20.544 basically within relatively safe

NOTE Confidence: 0.9402536

00:18:20.544 --> 00:18:22.320 guidelines which are comparable,

NOTE Confidence: 0.9402536

00:18:22.320 --> 00:18:23.712 not exactly the same,

NOTE Confidence: 0.9402536

00:18:23.712 --> 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 --> 00:18:31.682 And so the ones in this with the

NOTE Confidence: 0.9402536

00:18:31.682 --> 00:18:34.646 solid lines are the ones who have
NOTE Confidence: 0.9402536

00:18:34.650 --> 00:18:38.526 who are still drinking at a,
NOTE Confidence: 0.9402536

00:18:38.530 --> 00:18:41.374 you know at a clinically significant
NOTE Confidence: 0.9402536

00:18:41.374 --> 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 --> 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 --> 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 --> 00:19:04.164 both groups maintain that almost it's
NOTE Confidence: 0.9402536

00:19:04.164 --> 00:19:06.782 it's just a flatline for both groups
NOTE Confidence: 0.9402536

00:19:06.782 --> 00:19:09.096 there's it's really no evidence that
NOTE Confidence: 0.9402536

00:19:09.096 --> 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 --> 00:19:18.173 but the it's a different story with

NOTE Confidence: 0.9402536

00:19:18.173 --> 00:19:20.501 the people who are still symptomatic.

NOTE Confidence: 0.9402536

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

NOTE Confidence: 0.9402536

00:19:21.802 --> 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 --> 00:19:27.626 this is, you know,

NOTE Confidence: 0.9402536

00:19:27.626 --> 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 --> 00:19:30.590 But.

NOTE Confidence: 0.9368991333333333

00:19:32.900 --> 00:19:35.820 42 total in the in the higher risk

NOTE Confidence: 0.9368991333333333

00:19:35.820 --> 00:19:38.115 group but still the you know it's

NOTE Confidence: 0.9368991333333333

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

NOTE Confidence: 0.9368991333333333

00:19:39.887 --> 00:19:41.543 looks like the groups are diverging

NOTE Confidence: 0.9368991333333333

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

NOTE Confidence: 0.9368991333333333

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

NOTE Confidence: 0.9368991333333333

00:19:44.900 --> 00:19:46.764 So it didn't look exactly like that for

NOTE Confidence: 0.9368991333333333

00:19:46.764 --> 00:19:48.780 all the outcomes for total abstinence,
NOTE Confidence: 0.9368991333333333

00:19:48.780 --> 00:19:51.300 there was some of the people who were
NOTE Confidence: 0.9368991333333333

00:19:51.300 --> 00:19:53.456 abstinent, you know or close to abstinent,
NOTE Confidence: 0.9368991333333333

00:19:53.460 --> 00:19:56.300 they still they benefited some
NOTE Confidence: 0.9368991333333333

00:19:56.300 --> 00:19:59.180 but for these continuous measures
NOTE Confidence: 0.9368991333333333

00:19:59.180 --> 00:20:00.950 it it looks like they're it.
NOTE Confidence: 0.9368991333333333

00:20:00.950 --> 00:20:03.438 You really get a larger effect for the
NOTE Confidence: 0.9368991333333333

00:20:03.438 --> 00:20:06.241 people who are still symptomatic and so non.
NOTE Confidence: 0.9368991333333333

00:20:06.241 --> 00:20:08.227 So that has implications both for
NOTE Confidence: 0.9368991333333333

00:20:08.227 --> 00:20:10.349 treatment and for study design in terms
NOTE Confidence: 0.9368991333333333

00:20:10.349 --> 00:20:12.748 of who you want to have in the trial.
NOTE Confidence: 0.88056946

00:20:15.310 --> 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 --> 00:20:24.629 real most of this in the last few years.
NOTE Confidence: 0.88056946

00:20:24.630 --> 00:20:26.758 There are quite a few studies now looking
NOTE Confidence: 0.88056946

00:20:26.758 --> 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 --> 00:20:33.501 disorder and it's really only three

NOTE Confidence: 0.88056946

00:20:33.501 --> 00:20:35.775 of those actually are with psilocybin.

NOTE Confidence: 0.88056946

00:20:35.780 --> 00:20:39.938 There's a number of other compounds

NOTE Confidence: 0.88056946

00:20:39.940 --> 00:20:44.512 including psilocybin and variety of

NOTE Confidence: 0.88056946

00:20:44.512 --> 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 --> 00:20:52.855 voluntary consumption preference,

NOTE Confidence: 0.88056946

00:20:52.855 --> 00:20:53.530 selfadministration,

NOTE Confidence: 0.88056946

00:20:53.530 --> 00:20:55.555 behavioral sensitization and

NOTE Confidence: 0.88056946

00:20:55.555 --> 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 --> 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 --> 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 --> 00:21:19.666 and there are a few negative studies.
NOTE Confidence: 0.88056946

00:21:19.670 --> 00:21:22.094 One of them was by the same group
NOTE Confidence: 0.88056946

00:21:22.094 --> 00:21:23.984 that published this the second study
NOTE Confidence: 0.88056946

00:21:23.984 --> 00:21:26.180 that that that I that's in the
NOTE Confidence: 0.88056946

00:21:26.180 --> 00:21:27.710 bottom of the slide here.
NOTE Confidence: 0.88056946

00:21:27.710 --> 00:21:31.620 They using a different model in this study,
NOTE Confidence: 0.88056946

00:21:31.620 --> 00:21:34.341 they were able to demonstrate that
NOTE Confidence: 0.88056946

00:21:34.341 --> 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 --> 00:21:47.201 and it also restored the M blue
NOTE Confidence: 0.88056946

00:21:47.201 --> 00:21:50.024 2 receptor gene expression in the
NOTE Confidence: 0.88056946

00:21:50.024 --> 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 --> 00:21:57.932 They also demonstrated in a different
NOTE Confidence: 0.88056946

00:21:57.932 --> 00:22:00.373 cohort of animals that knocked down

NOTE Confidence: 0.88056946

00:22:00.373 --> 00:22:03.710 of this of the M blue tube receptor

NOTE Confidence: 0.88056946

00:22:03.710 --> 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 --> 00:22:11.110 Q induced reinstatement and so.

NOTE Confidence: 0.88056946

00:22:11.110 --> 00:22:13.598 So the story is that this

NOTE Confidence: 0.88056946

00:22:13.598 --> 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 --> 00:22:19.790 right now probably.

NOTE Confidence: 0.88056946

00:22:19.790 --> 00:22:22.454 But they haven't you know shown that this

NOTE Confidence: 0.88056946

00:22:22.454 --> 00:22:25.368 is what happens when psilocybin treated.

NOTE Confidence: 0.88056946

00:22:25.370 --> 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 --> 00:22:28.530 it's a hypothesis and a testable 1.

NOTE Confidence: 0.88056946

00:22:28.530 --> 00:22:31.127 So we'll see how that pans out.

NOTE Confidence: 0.88056946

00:22:31.130 --> 00:22:31.490 But
NOTE Confidence: 0.94377579

00:22:34.370 --> 00:22:36.827 that's, you know, one of the more
NOTE Confidence: 0.94377579

00:22:36.827 --> 00:22:38.955 sophisticated efforts to really come up
NOTE Confidence: 0.94377579

00:22:38.955 --> 00:22:41.321 with an explanation on a molecular level.
NOTE Confidence: 0.943128857142857

00:22:41.330 --> 00:22:42.646 Do you know where that knockdown was?
NOTE Confidence: 0.943128857142857

00:22:42.650 --> 00:22:45.330 Was that medial Pfc or was it global or was I
NOTE Confidence: 0.86817964

00:22:48.530 --> 00:22:49.350 think it was global?
NOTE Confidence: 0.86817964

00:22:49.350 --> 00:22:50.930 I think it was. I think it were.
NOTE Confidence: 0.9402536

00:22:53.010 --> 00:22:54.610 Yeah, I think I think so.
NOTE Confidence: 0.9191548025

00:23:01.330 --> 00:23:04.132 Okay. There's been more done in
NOTE Confidence: 0.9191548025

00:23:04.132 --> 00:23:06.610 animal models of depression and
NOTE Confidence: 0.9191548025

00:23:06.610 --> 00:23:09.250 anxiety and stress related disorders
NOTE Confidence: 0.9191548025

00:23:09.250 --> 00:23:11.806 and there have been, you know,
NOTE Confidence: 0.9191548025

00:23:11.810 --> 00:23:14.090 persisting effects demonstrated
NOTE Confidence: 0.9191548025

00:23:14.090 --> 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 --> 00:23:21.610 including for swim test,
NOTE Confidence: 0.9191548025
00:23:21.610 --> 00:23:22.888 aversive foot shock
NOTE Confidence: 0.950317
00:23:25.290 --> 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 --> 00:23:35.898 it's the, it's it's a hedonic response.
NOTE Confidence: 0.950317
00:23:35.900 --> 00:23:38.616 So it's they recover their interest in
NOTE Confidence: 0.950317
00:23:38.620 --> 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 --> 00:23:50.580 anxiolytic and antidepressant effects and
NOTE Confidence: 0.950317
00:23:50.580 --> 00:23:56.220 also the facilitation of fear extinction.
NOTE Confidence: 0.950317
00:23:56.220 --> 00:23:59.804 So, so this all looks promising and
NOTE Confidence: 0.950317
00:23:59.804 --> 00:24:02.252 and fairly consistent across you know
NOTE Confidence: 0.950317
00:24:02.252 --> 00:24:04.380 a number of of of different systems
NOTE Confidence: 0.943175857142857
00:24:06.460 --> 00:24:09.248 neuroplastic effects you know
NOTE Confidence: 0.943175857142857

00:24:09.248 --> 00:24:11.850 been clearly demonstrated including
NOTE Confidence: 0.943175857142857

00:24:11.850 --> 00:24:13.890 spinogenesis and eptogenesis and
NOTE Confidence: 0.943175857142857

00:24:13.890 --> 00:24:16.440 and some extent neurogenesis that's
NOTE Confidence: 0.943175857142857

00:24:16.513 --> 00:24:18.739 you know the significance of that.
NOTE Confidence: 0.943175857142857

00:24:18.740 --> 00:24:20.228 Well the significance of all of
NOTE Confidence: 0.943175857142857

00:24:20.228 --> 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 --> 00:24:28.830 so it doesn't mean that this is all
NOTE Confidence: 0.943175857142857

00:24:28.830 --> 00:24:31.504 great and causing anything but it you
NOTE Confidence: 0.943175857142857

00:24:31.504 --> 00:24:34.632 know this does this is something that
NOTE Confidence: 0.943175857142857

00:24:34.632 --> 00:24:36.815 reliably happens and it you know that's
NOTE Confidence: 0.943175857142857

00:24:36.815 --> 00:24:38.510 the case with ketamine as well so.
NOTE Confidence: 0.922931007692308

00:24:41.000 --> 00:24:43.664 But we'll see and there's also
NOTE Confidence: 0.922931007692308

00:24:43.664 --> 00:24:44.996 these antiinflammatory effects
NOTE Confidence: 0.922931007692308

00:24:44.996 --> 00:24:47.609 of classic psychedelics which you

NOTE Confidence: 0.922931007692308

00:24:47.609 --> 00:24:49.343 know really it's not clear what

NOTE Confidence: 0.922931007692308

00:24:49.343 --> 00:24:50.999 the significance is at this point,

NOTE Confidence: 0.922931007692308

00:24:51.000 --> 00:24:51.732 but they're pretty,

NOTE Confidence: 0.922931007692308

00:24:51.732 --> 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 --> 00:24:59.200 sub sub experiential doses that.

NOTE Confidence: 0.922931007692308

00:24:59.200 --> 00:25:03.536 So it's whole other possible mechanism

NOTE Confidence: 0.922931007692308

00:25:03.536 --> 00:25:06.440 of action at least for some,

NOTE Confidence: 0.922931007692308

00:25:06.440 --> 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 --> 00:25:14.669 about these drugs AS5HD2A agonists or

NOTE Confidence: 0.941259872

00:25:14.669 --> 00:25:17.387 partial agonists that that is true.

NOTE Confidence: 0.941259872

00:25:17.390 --> 00:25:19.308 And a lot of the effects definitely

NOTE Confidence: 0.941259872

00:25:19.308 --> 00:25:23.229 seem to be mediated by that receptor

NOTE Confidence: 0.941259872

00:25:23.230 --> 00:25:25.522 and you know, we can attenuate

NOTE Confidence: 0.941259872

00:25:25.522 --> 00:25:27.549 those effectors by by blocking it
NOTE Confidence: 0.9377376925

00:25:29.630 --> 00:25:32.178 or knocking it out.
NOTE Confidence: 0.9377376925

00:25:32.180 --> 00:25:34.882 But there are a couple of studies
NOTE Confidence: 0.9377376925

00:25:34.882 --> 00:25:36.947 demonstrating that in animal models
NOTE Confidence: 0.9377376925

00:25:36.947 --> 00:25:39.521 that you can get the neurotrophic
NOTE Confidence: 0.9377376925

00:25:39.521 --> 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 --> 00:25:52.044 at least to the extent that you don't
NOTE Confidence: 0.932202801

00:25:52.044 --> 00:25:55.046 get the head twitch which is considered
NOTE Confidence: 0.932202801

00:25:55.046 --> 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 --> 00:25:59.945 I wouldn't say it's the
NOTE Confidence: 0.932202801

00:25:59.945 --> 00:26:01.354 animal equivalent, but it's.
NOTE Confidence: 0.932202801

00:26:01.354 --> 00:26:05.250 If it's the kind of dose that would,
NOTE Confidence: 0.932202801

00:26:05.250 --> 00:26:07.357 it's the animal counterpart to the to
NOTE Confidence: 0.932202801

00:26:07.357 --> 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

NOTE Confidence: 0.932202801

00:26:10.919 --> 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 --> 00:26:15.767 human it's going to be a psychedelic.

NOTE Confidence: 0.932202801

00:26:15.770 --> 00:26:19.370 So that's so the point of this is

NOTE Confidence: 0.932202801

00:26:19.370 --> 00:26:21.770 you know not that to a agonism isn't

NOTE Confidence: 0.932202801

00:26:21.770 --> 00:26:23.760 important but that it may be possible

NOTE Confidence: 0.932202801

00:26:23.760 --> 00:26:25.810 to get some of these therapeutic

NOTE Confidence: 0.932202801

00:26:25.810 --> 00:26:29.540 benefits without.

NOTE Confidence: 0.932202801

00:26:29.540 --> 00:26:31.969 The mind altering effects or at least

NOTE Confidence: 0.932202801

00:26:31.969 --> 00:26:34.617 without you know the same extent of them.

NOTE Confidence: 0.932202801

00:26:34.620 --> 00:26:37.460 So that remains to be demonstrated in people,

NOTE Confidence: 0.932202801

00:26:37.460 --> 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 --> 00:26:46.613 we can say about it again that you

NOTE Confidence: 0.932202801

00:26:46.613 --> 00:26:48.404 know the trial that we completed

NOTE Confidence: 0.932202801

00:26:48.404 --> 00:26:50.094 wasn't wasn't a mechanistic study,
NOTE Confidence: 0.932202801

00:26:50.100 --> 00:26:53.316 but we had a lot of self report
NOTE Confidence: 0.932202801

00:26:53.316 --> 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 --> 00:27:00.924 I'll show you some of the results
NOTE Confidence: 0.932202801

00:27:00.924 --> 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 --> 00:27:07.189 it's very small but it's you know somewhat
NOTE Confidence: 0.94276945

00:27:09.430 --> 00:27:12.370 interesting I think.
NOTE Confidence: 0.94276945

00:27:12.370 --> 00:27:15.770 So starting with some of the
NOTE Confidence: 0.94276945

00:27:15.770 --> 00:27:17.706 obvious self report measures,
NOTE Confidence: 0.94276945

00:27:17.710 --> 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 --> 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

NOTE Confidence: 0.94276945

00:27:32.641 --> 00:27:35.486 pretty much over the the entire period.

NOTE Confidence: 0.94276945

00:27:35.486 --> 00:27:39.014 So these just quick or quickly Orient you to

NOTE Confidence: 0.94276945

00:27:39.014 --> 00:27:41.870 these slides because I'll show four of them.

NOTE Confidence: 0.94276945

00:27:41.870 --> 00:27:46.350 I think the where it says week four,

NOTE Confidence: 0.94276945

00:27:46.350 --> 00:27:48.943 that's really the baseline in this MMRM.

NOTE Confidence: 0.94276945

00:27:48.943 --> 00:27:49.882 So that's not.

NOTE Confidence: 0.94276945

00:27:49.882 --> 00:27:52.319 Part of the repeated measure that's that's a,

NOTE Confidence: 0.94276945

00:27:52.320 --> 00:27:53.740 that's the baseline covariate

NOTE Confidence: 0.94276945

00:27:53.740 --> 00:27:55.515 and where it says baseline,

NOTE Confidence: 0.94276945

00:27:55.520 --> 00:27:56.815 that's the beginning of the

NOTE Confidence: 0.94276945

00:27:56.815 --> 00:27:58.597 study and that's in there just to

NOTE Confidence: 0.94276945

00:27:58.597 --> 00:28:00.116 kind of show you what they were,

NOTE Confidence: 0.94276945

00:28:00.120 --> 00:28:01.116 where they were at the beginning.

NOTE Confidence: 0.94276945

00:28:01.120 --> 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

NOTE Confidence: 0.94276945

00:28:04.980 --> 00:28:05.800 connected the dots, but
NOTE Confidence: 0.931427584615385

00:28:08.240 --> 00:28:10.438 so then all of the points after
NOTE Confidence: 0.931427584615385

00:28:10.438 --> 00:28:12.320 that five weeks 5 through 36,
NOTE Confidence: 0.931427584615385

00:28:12.320 --> 00:28:14.580 those are the repeated measure.
NOTE Confidence: 0.931427584615385

00:28:14.580 --> 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 --> 00:28:21.082 called the treatment effect and it
NOTE Confidence: 0.931427584615385

00:28:21.145 --> 00:28:23.419 was highly significant and the time
NOTE Confidence: 0.931427584615385

00:28:23.419 --> 00:28:25.774 effect is called assessment and that's
NOTE Confidence: 0.931427584615385

00:28:25.774 --> 00:28:27.859 also highly significant is they're,
NOTE Confidence: 0.931427584615385

00:28:27.860 --> 00:28:29.690 they're both groups are going
NOTE Confidence: 0.931427584615385

00:28:29.690 --> 00:28:31.990 downhill and the time by treatment
NOTE Confidence: 0.931427584615385

00:28:31.990 --> 00:28:34.310 assessment is you know whether the
NOTE Confidence: 0.931427584615385

00:28:34.310 --> 00:28:36.686 slopes are the same after people
NOTE Confidence: 0.931427584615385

00:28:36.686 --> 00:28:38.898 after receiving the psilocybin.
NOTE Confidence: 0.931427584615385

00:28:38.900 --> 00:28:40.292 And so they weren't

NOTE Confidence: 0.931427584615385
00:28:40.292 --> 00:28:41.336 significantly different here,
NOTE Confidence: 0.931427584615385
00:28:41.340 --> 00:28:43.220 but you know maybe close.
NOTE Confidence: 0.916508510333333
00:28:44.140 --> 00:28:45.700 It is interesting that the cravings
NOTE Confidence: 0.916508510333333
00:28:45.700 --> 00:28:47.276 continues to angle down throughout most
NOTE Confidence: 0.916508510333333
00:28:47.276 --> 00:28:49.244 of the followup as opposed to your other
NOTE Confidence: 0.916508510333333
00:28:49.296 --> 00:28:50.880 outcome measures where you got your
NOTE Confidence: 0.916508510333333
00:28:50.880 --> 00:28:52.536 your strongest effect at week five,
NOTE Confidence: 0.916508510333333
00:28:52.536 --> 00:28:54.920 the five to eight time point and then
NOTE Confidence: 0.916508510333333
00:28:54.984 --> 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: 0.933544666666667
00:28:59.620 --> 00:29:01.897 Well, I mean I think it's a real effect,
NOTE Confidence: 0.933544666666667
00:29:01.900 --> 00:29:04.388 it's the, the best way to decrease craving
NOTE Confidence: 0.933544666666667
00:29:04.388 --> 00:29:06.897 is to get people to stop drinking.
NOTE Confidence: 0.933544666666667
00:29:06.900 --> 00:29:08.988 So you know it always goes
NOTE Confidence: 0.933544666666667
00:29:08.988 --> 00:29:10.620 down if people were asked.
NOTE Confidence: 0.933544666666667

00:29:10.620 --> 00:29:13.168 So you know downstream this, you know this.
NOTE Confidence: 0.933544666666667

00:29:13.168 --> 00:29:15.718 You know, like we will maybe try to spend,
NOTE Confidence: 0.933544666666667

00:29:15.720 --> 00:29:17.380 those people are not drinking
NOTE Confidence: 0.933544666666667

00:29:17.380 --> 00:29:19.040 because we decreased their craving.
NOTE Confidence: 0.933544666666667

00:29:19.040 --> 00:29:20.140 But after week five,
NOTE Confidence: 0.933544666666667

00:29:20.140 --> 00:29:21.240 if they stop drinking,
NOTE Confidence: 0.933544666666667

00:29:21.240 --> 00:29:22.758 then their craving is also going
NOTE Confidence: 0.933544666666667

00:29:22.758 --> 00:29:24.240 down because they stop drinking.
NOTE Confidence: 0.933544666666667

00:29:24.240 --> 00:29:28.720 So it it's it's not a very strong causal
NOTE Confidence: 0.933544666666667

00:29:28.720 --> 00:29:30.239 even suggestion that we can make here,
NOTE Confidence: 0.933544666666667

00:29:30.240 --> 00:29:31.200 but it's, you know,
NOTE Confidence: 0.933544666666667

00:29:31.200 --> 00:29:32.188 it's a possibility, right?
NOTE Confidence: 0.933544666666667

00:29:32.188 --> 00:29:33.528 If we don't know the
NOTE Confidence: 0.933544666666667

00:29:33.528 --> 00:29:34.600 direction of the causality,
NOTE Confidence: 0.941930833333333

00:29:37.200 --> 00:29:41.196 this is something kind of like.
NOTE Confidence: 0.941930833333333

00:29:41.200 --> 00:29:44.920 Craving it's temptation from the Alcohol

NOTE Confidence: 0.9419308333333333
00:29:44.920 --> 00:29:46.360 Abstinent Selfefficacy questionnaire.
NOTE Confidence: 0.9419308333333333
00:29:46.360 --> 00:29:49.240 So this is hypothetical craving really.
NOTE Confidence: 0.9419308333333333
00:29:49.240 --> 00:29:51.416 If you were you know in a walking
NOTE Confidence: 0.9419308333333333
00:29:51.416 --> 00:29:53.380 down the street and you passed
NOTE Confidence: 0.9419308333333333
00:29:53.380 --> 00:29:55.480 your favorite bar and your buddy
NOTE Confidence: 0.9419308333333333
00:29:55.480 --> 00:29:57.836 said come on in don't be a square,
NOTE Confidence: 0.9419308333333333
00:29:57.840 --> 00:30:00.836 how tempted would you be to drink
NOTE Confidence: 0.9419308333333333
00:30:00.840 --> 00:30:03.144 and and you rate a bunch of 20
NOTE Confidence: 0.9419308333333333
00:30:03.144 --> 00:30:07.110 different things like that and so.
NOTE Confidence: 0.9419308333333333
00:30:07.110 --> 00:30:09.182 We see in this case the separation
NOTE Confidence: 0.9419308333333333
00:30:09.182 --> 00:30:10.881 didn't really happen until after
NOTE Confidence: 0.9419308333333333
00:30:10.881 --> 00:30:12.389 the second psilocybin session,
NOTE Confidence: 0.9419308333333333
00:30:12.390 --> 00:30:13.314 which is you know,
NOTE Confidence: 0.9419308333333333
00:30:13.314 --> 00:30:15.934 so that we saw that in a couple of these
NOTE Confidence: 0.9419308333333333
00:30:15.934 --> 00:30:18.630 measures and then it is maintained.
NOTE Confidence: 0.9419308333333333

00:30:18.630 --> 00:30:22.650 So in this case we we do have a treatment
NOTE Confidence: 0.9419308333333333

00:30:22.650 --> 00:30:24.832 by assessment interaction and yeah,
NOTE Confidence: 0.9419308333333333

00:30:24.832 --> 00:30:26.837 I don't know about that.
NOTE Confidence: 0.9419308333333333

00:30:26.840 --> 00:30:28.884 Delay, but it it does look different
NOTE Confidence: 0.9419308333333333

00:30:28.884 --> 00:30:30.839 from some of the other curves.
NOTE Confidence: 0.956354982

00:30:30.960 --> 00:30:33.120 How much does this measure correlate
NOTE Confidence: 0.956354982

00:30:33.120 --> 00:30:34.560 with other clinical outcomes?
NOTE Confidence: 0.956354982

00:30:34.560 --> 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 --> 00:30:44.158 think it's, I mean people often
NOTE Confidence: 0.942083229090909

00:30:44.158 --> 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 --> 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 --> 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 --> 00:31:07.577 I can stop that it means something.
NOTE Confidence: 0.942083229090909
00:31:07.580 --> 00:31:14.328 And so you can see here there was yeah,
NOTE Confidence: 0.942083229090909
00:31:14.328 --> 00:31:16.912 this is 1 where there's a there
NOTE Confidence: 0.942083229090909
00:31:16.912 --> 00:31:18.172 isn't a treatment by assessment
NOTE Confidence: 0.942083229090909
00:31:18.172 --> 00:31:19.421 interaction that's going to sound
NOTE Confidence: 0.942083229090909
00:31:19.421 --> 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 --> 00:31:25.314 including the very baseline,
NOTE Confidence: 0.940253533333333
00:31:25.320 --> 00:31:26.762 yeah, but it's it still looks like
NOTE Confidence: 0.940253533333333
00:31:26.762 --> 00:31:27.961 the slopes would be different
NOTE Confidence: 0.940253533333333

00:31:27.961 --> 00:31:29.037 but apparently they're not.

NOTE Confidence: 0.9402535333333333

00:31:29.040 --> 00:31:31.392 But there is this you know pretty

NOTE Confidence: 0.9402535333333333

00:31:31.392 --> 00:31:33.224 strong treatment effect and it's

NOTE Confidence: 0.9402535333333333

00:31:33.224 --> 00:31:36.144 it's persists overtime and so

NOTE Confidence: 0.9402535333333333

00:31:36.144 --> 00:31:39.335 that's that's good and that is

NOTE Confidence: 0.9402535333333333

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

NOTE Confidence: 0.9402535333333333

00:31:41.345 --> 00:31:43.564 outcomes and then self compassion

NOTE Confidence: 0.9402535333333333

00:31:43.564 --> 00:31:46.262 is kind of a complicated construct,

NOTE Confidence: 0.9402535333333333

00:31:46.262 --> 00:31:48.656 but I think it's it's highly relevant.

NOTE Confidence: 0.9402535333333333

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

NOTE Confidence: 0.9402535333333333

00:31:49.260 --> 00:31:50.856 the effects are really this is I

NOTE Confidence: 0.9402535333333333

00:31:50.856 --> 00:31:52.317 think the largest effect that we

NOTE Confidence: 0.9402535333333333

00:31:52.317 --> 00:31:53.775 saw in any of the questionnaires.

NOTE Confidence: 0.941691338571429

00:31:55.900 --> 00:31:58.616 And there are two subscales to this,

NOTE Confidence: 0.941691338571429

00:31:58.620 --> 00:32:02.624 and one of them is sort of the

NOTE Confidence: 0.941691338571429

00:32:02.624 --> 00:32:03.696 opposite of self compassion.

NOTE Confidence: 0.941691338571429

00:32:03.700 --> 00:32:05.140 It's it's self criticism.

NOTE Confidence: 0.941691338571429

00:32:05.140 --> 00:32:07.762 It's the extent to which you, you know,

NOTE Confidence: 0.941691338571429

00:32:07.762 --> 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 --> 00:32:17.488 You know that kind of negative

NOTE Confidence: 0.915843438571429

00:32:17.488 --> 00:32:19.739 self talk is a big part of it.

NOTE Confidence: 0.915843438571429

00:32:19.740 --> 00:32:22.218 And the other half is self compassion,

NOTE Confidence: 0.915843438571429

00:32:22.220 --> 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 --> 00:32:27.180 well put it in perspective,

NOTE Confidence: 0.915843438571429

00:32:27.180 --> 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 --> 00:32:33.052 You know you have good qualities or

NOTE Confidence: 0.915843438571429

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 --> 00:32:36.335 or what those those sorts of things.
NOTE Confidence: 0.915843438571429

00:32:36.340 --> 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 --> 00:32:43.332 so if we want to you know in
NOTE Confidence: 0.915843438571429

00:32:43.332 --> 00:32:44.828 terms of the the three domains,
NOTE Confidence: 0.915843438571429

00:32:44.830 --> 00:32:47.901 is this executive control or or you
NOTE Confidence: 0.915843438571429

00:32:47.901 --> 00:32:49.798 know negative or you know less negative
NOTE Confidence: 0.915843438571429

00:32:49.798 --> 00:32:51.548 affect it's probably both but I it
NOTE Confidence: 0.915843438571429

00:32:51.548 --> 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 --> 00:32:58.238 in in capturing something that
NOTE Confidence: 0.915843438571429

00:32:58.238 --> 00:32:59.878 changes you know both cognitively
NOTE Confidence: 0.915843438571429

00:32:59.878 --> 00:33:01.190 and effectively for people.
NOTE Confidence: 0.916288475

00:33:03.830 --> 00:33:05.720 So that's interesting and we you know
NOTE Confidence: 0.916288475

00:33:05.720 --> 00:33:07.965 we're working on a paper where I mean it

NOTE Confidence: 0.916288475

00:33:07.965 --> 00:33:09.890 looks like they're you know there is some.

NOTE Confidence: 0.916288475

00:33:09.890 --> 00:33:13.122 It it does predict some amount of the

NOTE Confidence: 0.916288475

00:33:13.122 --> 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: 0.934123811111111

00:33:17.930 --> 00:33:19.040 The self compassion is the

NOTE Confidence: 0.934123811111111

00:33:19.040 --> 00:33:19.928 biggest of the individual.

NOTE Confidence: 0.934123811111111

00:33:19.930 --> 00:33:20.958 Well that's the biggest

NOTE Confidence: 0.934123811111111

00:33:20.958 --> 00:33:22.243 effect just in between group.

NOTE Confidence: 0.934123811111111

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

NOTE Confidence: 0.934123811111111

00:33:23.670 --> 00:33:25.090 of like the self report,

NOTE Confidence: 0.934123811111111

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

NOTE Confidence: 0.934123811111111

00:33:27.310 --> 00:33:29.290 the self report itself. Have

NOTE Confidence: 0.933544666666667

00:33:29.290 --> 00:33:31.920 you done any multivariate analysis

NOTE Confidence: 0.933544666666667

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

NOTE Confidence: 0.94226628

00:33:38.340 --> 00:33:39.260 I'm blanking on the analysis

NOTE Confidence: 0.94226628

00:33:39.260 --> 00:33:40.180 I thought you could do,
NOTE Confidence: 0.94226628

00:33:40.180 --> 00:33:42.378 but something that looks at the relationship
NOTE Confidence: 0.94226628

00:33:42.378 --> 00:33:44.339 between these two measures over time,
NOTE Confidence: 0.94226628

00:33:44.340 --> 00:33:47.956 because the curves as you've pointed as
NOTE Confidence: 0.94226628

00:33:47.956 --> 00:33:50.058 we've gone through the curves and the
NOTE Confidence: 0.94226628

00:33:50.058 --> 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 --> 00:33:54.098 which could be noise in the data,
NOTE Confidence: 0.94226628

00:33:54.100 --> 00:33:56.704 but you might wonder if you could
NOTE Confidence: 0.94226628

00:33:56.704 --> 00:33:59.059 make some inferences across about the.
NOTE Confidence: 0.94226628

00:33:59.060 --> 00:34:00.370 You know, the unfolding of
NOTE Confidence: 0.94226628

00:34:00.370 --> 00:34:01.900 different phases of the effect of
NOTE Confidence: 0.931966094117647

00:34:01.900 --> 00:34:03.826 no, I would love to and I mean that
NOTE Confidence: 0.931966094117647

00:34:03.826 --> 00:34:05.896 is something we can do and you know
NOTE Confidence: 0.931966094117647

00:34:05.900 --> 00:34:07.124 the drinking outcomes we have in
NOTE Confidence: 0.931966094117647

00:34:07.124 --> 00:34:08.577 these one month, one month bins.

NOTE Confidence: 0.931966094117647
00:34:08.577 --> 00:34:11.687 So we can we can tease that out and we could
NOTE Confidence: 0.931966094117647
00:34:11.687 --> 00:34:14.262 do cross like kind of analysis like that.
NOTE Confidence: 0.931966094117647
00:34:14.262 --> 00:34:15.588 I mean the problem is there's
NOTE Confidence: 0.931966094117647
00:34:15.588 --> 00:34:16.857 just so many we could do,
NOTE Confidence: 0.931966094117647
00:34:16.860 --> 00:34:19.292 you know so you don't want to take
NOTE Confidence: 0.931966094117647
00:34:19.292 --> 00:34:21.957 too many bites of the apple here but.
NOTE Confidence: 0.931966094117647
00:34:21.960 --> 00:34:23.358 And it's not a huge sample,
NOTE Confidence: 0.931966094117647
00:34:23.360 --> 00:34:25.480 so I don't want to over analyze this,
NOTE Confidence: 0.931966094117647
00:34:25.480 --> 00:34:26.944 but I think you know those
NOTE Confidence: 0.931966094117647
00:34:26.944 --> 00:34:27.920 are really good questions.
NOTE Confidence: 0.931966094117647
00:34:27.920 --> 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 --> 00:34:41.110 responders and we can do that
NOTE Confidence: 0.931966094117647
00:34:41.110 --> 00:34:42.594 based on baseline characteristics.
NOTE Confidence: 0.931966094117647

00:34:42.600 --> 00:34:45.936 We could also do that based on the
NOTE Confidence: 0.931966094117647

00:34:45.936 --> 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 --> 00:34:53.099 some of these self-reports or you know,
NOTE Confidence: 0.931966094117647

00:34:53.100 --> 00:34:55.164 so you can include you look at the
NOTE Confidence: 0.931966094117647

00:34:55.164 --> 00:34:56.632 pretreatment things but then you could
NOTE Confidence: 0.931966094117647

00:34:56.632 --> 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 --> 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 --> 00:35:12.420 So how about how about the nature of the.
NOTE Confidence: 0.925158445

00:35:12.420 --> 00:35:13.646 Intoxication effect. Yeah.
NOTE Confidence: 0.925158445

00:35:13.646 --> 00:35:14.776 So we'll get to that.
NOTE Confidence: 0.925158445

00:35:14.780 --> 00:35:16.327 That's So that's the other that's the
NOTE Confidence: 0.925158445

00:35:16.327 --> 00:35:18.017 new piece that I want to show you too.
NOTE Confidence: 0.925158445

00:35:18.020 --> 00:35:20.218 We have taken some look at that.

NOTE Confidence: 0.925158445

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

NOTE Confidence: 0.925158445

00:35:24.140 --> 00:35:27.085 known that psilocybin tends to

NOTE Confidence: 0.925158445

00:35:27.085 --> 00:35:28.695 produce some personality changes.

NOTE Confidence: 0.925158445

00:35:28.695 --> 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 --> 00:35:35.578 decrease in neuroticism was that it was

NOTE Confidence: 0.925158445

00:35:35.580 --> 00:35:37.818 had been demonstrated before as well.

NOTE Confidence: 0.925158445

00:35:37.820 --> 00:35:39.938 So we saw.

NOTE Confidence: 0.925158445

00:35:39.940 --> 00:35:41.840 Yeah between group differences in

NOTE Confidence: 0.925158445

00:35:41.840 --> 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 --> 00:35:48.937 and within the psilocybin group,

NOTE Confidence: 0.925158445

00:35:48.940 --> 00:35:50.700 conscientiousness also increased but you

NOTE Confidence: 0.925158445

00:35:50.700 --> 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,

NOTE Confidence: 0.925158445

00:35:56.916 --> 00:36:00.244 I don't want to over reify these personality

NOTE Confidence: 0.925158445

00:36:00.244 --> 00:36:02.196 dimensions either because you know,

NOTE Confidence: 0.925158445

00:36:02.196 --> 00:36:04.020 I mean we we like to think about

NOTE Confidence: 0.925158445

00:36:04.074 --> 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 --> 00:36:08.758 you know you're going to act

NOTE Confidence: 0.925158445

00:36:08.758 --> 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 --> 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 --> 00:36:22.260 somebody 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 --> 00:36:25.780 there is some truth to that.

NOTE Confidence: 0.925158445

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

NOTE Confidence: 0.925158445

00:36:27.372 --> 00:36:29.362 these were pretty robust differences

NOTE Confidence: 0.925158445

00:36:29.362 --> 00:36:31.819 and it's particularly interesting

NOTE Confidence: 0.925158445

00:36:31.820 --> 00:36:34.564 if you look at the facets that

NOTE Confidence: 0.925158445

00:36:34.564 --> 00:36:36.247 actually changed because there's

NOTE Confidence: 0.925158445

00:36:36.247 --> 00:36:38.387 a lot inside of neuroticism.

NOTE Confidence: 0.925158445

00:36:38.390 --> 00:36:40.945 And what actually changed significantly

NOTE Confidence: 0.925158445

00:36:40.945 --> 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 --> 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 --> 00:36:51.410 it wasn't that they wanted to

NOTE Confidence: 0.925158445

00:36:51.410 --> 00:36:53.586 go out and party but not drink.

NOTE Confidence: 0.925158445

00:36:53.590 --> 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 --> 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 --> 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 --> 00:37:16.830 when we talk about cognitive flexibility,
NOTE Confidence: 0.925158445

00:37:16.830 --> 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 --> 00:37:19.507 of different things.
NOTE Confidence: 0.925158445

00:37:19.510 --> 00:37:22.366 But this is one kind of flexibility perhaps
NOTE Confidence: 0.925158445

00:37:22.366 --> 00:37:25.470 that that we're saying that could be useful.

NOTE Confidence: 0.925158445
00:37:25.470 --> 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 --> 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 --> 00:37:32.590 does this mean they've they had,
NOTE Confidence: 0.925158445
00:37:32.590 --> 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 --> 00:37:43.845 so that's suggestive but don't
NOTE Confidence: 0.939723947368421
00:37:43.845 --> 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 --> 00:37:50.306 So, so here's the really the
NOTE Confidence: 0.939723947368421
00:37:50.306 --> 00:37:51.850 really new part that these next
NOTE Confidence: 0.939723947368421
00:37:51.850 --> 00:37:53.416 two things I have not presented
NOTE Confidence: 0.939723947368421

00:37:53.416 --> 00:37:55.088 before and be interested to see
NOTE Confidence: 0.939723947368421

00:37:55.088 --> 00:37:56.458 what people think about them.
NOTE Confidence: 0.939723947368421

00:37:56.460 --> 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: 0.928512966666667

00:38:00.550 --> 00:38:04.330 time is at the time is. It's a construct.
NOTE Confidence: 0.928512966666667

00:38:04.330 --> 00:38:06.682 I don't want to over reapply the
NOTE Confidence: 0.928512966666667

00:38:06.682 --> 00:38:09.109 construct and I shouldn't take it instead
NOTE Confidence: 0.9301902

00:38:09.110 --> 00:38:09.990 of the Uber, but
NOTE Confidence: 0.94226628

00:38:12.830 --> 00:38:17.430 so we used four questionnaires
NOTE Confidence: 0.94226628

00:38:17.430 --> 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 --> 00:38:23.854 And I'll tell you more about
NOTE Confidence: 0.94226628

00:38:23.854 --> 00:38:25.180 what those were shortly.
NOTE Confidence: 0.94226628

00:38:25.180 --> 00:38:26.998 There are other questionnaires that are
NOTE Confidence: 0.94226628

00:38:26.998 --> 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 --> 00:38:32.216 I kind of wish we did.

NOTE Confidence: 0.94226628

00:38:32.220 --> 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 --> 00:38:39.908 There's one called the that specifically

NOTE Confidence: 0.94226628

00:38:39.908 --> 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 --> 00:38:48.960 I think there's even a couple more now, but.

NOTE Confidence: 0.94226628

00:38:48.960 --> 00:38:52.537 And I decided that it it's a

NOTE Confidence: 0.94226628

00:38:52.537 --> 00:38:54.490 little bit complicated because not

NOTE Confidence: 0.94226628

00:38:54.490 --> 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 --> 00:38:58.713 but there are people who didn't

NOTE Confidence: 0.94226628

00:38:58.720 --> 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 --> 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 --> 00:39:09.080 psychotherapy, but they
NOTE Confidence: 0.96641844

00:39:09.080 --> 00:39:09.880 declined psychotherapy
NOTE Confidence: 0.87031739625

00:39:09.880 --> 00:39:12.280 in the follow up, but just just said,
NOTE Confidence: 0.87031739625

00:39:12.280 --> 00:39:13.636 you know, that was too much
NOTE Confidence: 0.87031739625

00:39:13.636 --> 00:39:15.079 for me or boring for me.
NOTE Confidence: 0.87031739625

00:39:15.079 --> 00:39:17.190 So and we we we wanted to be
NOTE Confidence: 0.87031739625

00:39:17.190 --> 00:39:19.460 able to let people do that. So
NOTE Confidence: 0.9452853

00:39:20.340 --> 00:39:21.912 and the rent of course people
NOTE Confidence: 0.9452853

00:39:21.912 --> 00:39:23.347 getting the same drug diphenhydromin
NOTE Confidence: 0.9452853

00:39:23.347 --> 00:39:24.977 or 07 in both sessions.
NOTE Confidence: 0.9452853

00:39:24.980 --> 00:39:27.780 So if they are pretty sure they
NOTE Confidence: 0.9452853

00:39:27.780 --> 00:39:29.424 got placebo the first time they're
NOTE Confidence: 0.9452853

00:39:29.424 --> 00:39:33.540 going to then they know yeah so
NOTE Confidence: 0.93514248

00:39:33.540 --> 00:39:36.012 so there wasn't a significantly different
NOTE Confidence: 0.93514248

00:39:36.012 --> 00:39:39.174 number of of dropouts in the in the but

NOTE Confidence: 0.93514248

00:39:39.174 --> 00:39:41.245 there were more people who declined

NOTE Confidence: 0.93514248

00:39:41.245 --> 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 --> 00:39:46.682 for different reasons and

NOTE Confidence: 0.95031694375

00:39:46.682 --> 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 --> 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 --> 00:39:58.142 be missing the boat because it just,

NOTE Confidence: 0.938097092857143

00:39:58.142 --> 00:39:59.638 you know, clinically people

NOTE Confidence: 0.938097092857143

00:39:59.638 --> 00:40:01.202 could have a terrible, you know,

NOTE Confidence: 0.938097092857143

00:40:01.202 --> 00:40:03.204 a miserable experience one time in a.

NOTE Confidence: 0.938097092857143

00:40:03.210 --> 00:40:05.262 Be a tivic experience the next time and and

NOTE Confidence: 0.938097092857143

00:40:05.262 --> 00:40:07.247 and we you know we definitely saw that.

NOTE Confidence: 0.938097092857143

00:40:07.250 --> 00:40:08.789 So I, you know I think we'd lose a

NOTE Confidence: 0.938097092857143

00:40:08.789 --> 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 --> 00:40:15.888 to add consider it sort of a dose,
NOTE Confidence: 0.938097092857143

00:40:15.890 --> 00:40:17.370 you know dose effect,
NOTE Confidence: 0.938097092857143

00:40:17.370 --> 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 --> 00:40:24.572 you get and I mean I don't know
NOTE Confidence: 0.938097092857143

00:40:24.572 --> 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 --> 00:40:29.978 So if they didn't have a second
NOTE Confidence: 0.938097092857143

00:40:29.978 --> 00:40:32.419 session they just get 0 for that one.
NOTE Confidence: 0.938097092857143

00:40:32.420 --> 00:40:33.484 And if they and and if they
NOTE Confidence: 0.938097092857143

00:40:33.484 --> 00:40:34.580 did have a second session,
NOTE Confidence: 0.938097092857143

00:40:34.580 --> 00:40:36.380 then the two are added together
NOTE Confidence: 0.938097092857143

00:40:36.380 --> 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 --> 00:40:41.416 look at and a lot of you know,

NOTE Confidence: 0.938097092857143

00:40:41.420 --> 00:40:44.096 other outcomes you could look at.

NOTE Confidence: 0.938097092857143

00:40:44.100 --> 00:40:46.540 So to keep it simple,

NOTE Confidence: 0.938097092857143

00:40:46.540 --> 00:40:49.124 what we're going to look at here is

NOTE Confidence: 0.938097092857143

00:40:49.124 --> 00:40:50.900 the correlations between you know,

NOTE Confidence: 0.938097092857143

00:40:50.900 --> 00:40:54.300 all these different Christira,

NOTE Confidence: 0.938097092857143

00:40:54.300 --> 00:40:54.460 sorry.

NOTE Confidence: 0.948639664166667

00:40:57.340 --> 00:40:59.195 So we're going to look at the

NOTE Confidence: 0.948639664166667

00:40:59.195 --> 00:41:00.433 correlation between these these

NOTE Confidence: 0.948639664166667

00:41:00.433 --> 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 --> 00:41:10.100 for the whole subsequent seven

NOTE Confidence: 0.948639664166667

00:41:10.100 --> 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 --> 00:41:18.303 after it's the first month after the

NOTE Confidence: 0.948639664166667

00:41:18.303 --> 00:41:20.319 second psilocybin session all the way
NOTE Confidence: 0.948639664166667

00:41:20.376 --> 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 --> 00:41:26.400 this for the whole sample.
NOTE Confidence: 0.948639664166667

00:41:26.400 --> 00:41:28.220 I don't think the whole sample is
NOTE Confidence: 0.948639664166667

00:41:28.220 --> 00:41:29.334 necessarily all that informative
NOTE Confidence: 0.948639664166667

00:41:29.334 --> 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 --> 00:41:33.836 if we looked at blood pressure
NOTE Confidence: 0.948639664166667

00:41:33.836 --> 00:41:35.916 as a mediator or you know this,
NOTE Confidence: 0.948639664166667

00:41:35.920 --> 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 --> 00:41:42.296 the psilocybin people have much higher blood
NOTE Confidence: 0.948639664166667

00:41:42.296 --> 00:41:44.398 pressure increase in and what do you know,
NOTE Confidence: 0.948639664166667

00:41:44.400 --> 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 --> 00:41:49.836 if there's a relationship there.
NOTE Confidence: 0.948639664166667
00:41:49.840 --> 00:41:50.820 And we could demonstrate that
NOTE Confidence: 0.948639664166667
00:41:50.820 --> 00:41:53.100 and that would be dumb, right?
NOTE Confidence: 0.948639664166667
00:41:53.100 --> 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 --> 00:41:59.744 It's a lot more convincing if we can
NOTE Confidence: 0.948639664166667
00:41:59.744 --> 00:42:01.070 demonstrate a relationship within
NOTE Confidence: 0.948639664166667
00:42:01.070 --> 00:42:03.050 the psilocybin group that like when
NOTE Confidence: 0.948639664166667
00:42:03.050 --> 00:42:05.516 they got this particular kind of experience,
NOTE Confidence: 0.948639664166667
00:42:05.520 --> 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 --> 00:42:11.060 with the effect being larger and
NOTE Confidence: 0.948639664166667
00:42:11.060 --> 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 --> 00:42:22.840 the people who were.
NOTE Confidence: 0.94654315875

00:42:22.840 --> 00:42:24.478 At a level of base of 0,
NOTE Confidence: 0.94654315875

00:42:24.480 --> 00:42:26.285 meaning abstinent people who were
NOTE Confidence: 0.94654315875

00:42:26.285 --> 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 --> 00:42:35.155 and then people who were at the
NOTE Confidence: 0.94654315875

00:42:35.155 --> 00:42:37.920 low level And then so I remember
NOTE Confidence: 0.94654315875

00:42:37.920 --> 00:42:38.880 first the abstinent people,
NOTE Confidence: 0.94654315875

00:42:38.880 --> 00:42:40.470 then the the low level drinkers
NOTE Confidence: 0.94654315875

00:42:40.470 --> 00:42:42.040 and then the moderate drinkers.
NOTE Confidence: 0.94654315875

00:42:42.040 --> 00:42:45.330 So you get smaller and smaller groups
NOTE Confidence: 0.94654315875

00:42:45.330 --> 00:42:46.916 so you know less and less power,
NOTE Confidence: 0.94654315875

00:42:46.920 --> 00:42:49.720 but you can still look at what the
NOTE Confidence: 0.94654315875

00:42:49.720 --> 00:42:51.660 correlations are and if they're.
NOTE Confidence: 0.94654315875

00:42:51.660 --> 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 --> 00:43:01.898 you may not know these scales

NOTE Confidence: 0.94654315875

00:43:01.898 --> 00:43:03.820 so I'll describe them briefly.

NOTE Confidence: 0.94654315875

00:43:03.820 --> 00:43:07.380 This is the the five D5 dimensional

NOTE Confidence: 0.94654315875

00:43:07.380 --> 00:43:09.580 altered states of consciousness scale.

NOTE Confidence: 0.94654315875

00:43:09.580 --> 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 --> 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 --> 00:43:29.960 you know medium size samples.

NOTE Confidence: 0.94654315875

00:43:29.960 --> 00:43:31.836 And so it has these factors as

NOTE Confidence: 0.94654315875

00:43:31.836 --> 00:43:33.801 five but these are the the first

NOTE Confidence: 0.94654315875

00:43:33.801 --> 00:43:35.904 three here are the main ones that

NOTE Confidence: 0.94654315875

00:43:35.904 --> 00:43:37.480 people generally care about.

NOTE Confidence: 0.94654315875

00:43:37.480 --> 00:43:40.273 First one is is what they call
NOTE Confidence: 0.94654315875

00:43:40.273 --> 00:43:42.006 oceanic boundlessness because they
NOTE Confidence: 0.94654315875

00:43:42.006 --> 00:43:43.870 think mystical experiences is.
NOTE Confidence: 0.94654315875

00:43:43.870 --> 00:43:44.454 Hocus pocus.
NOTE Confidence: 0.94654315875

00:43:44.454 --> 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 --> 00:43:50.990 It probably sounds better in German, but
NOTE Confidence: 0.944027375

00:43:53.510 --> 00:43:56.750 it's anyway it's really this this you know,
NOTE Confidence: 0.944027375

00:43:56.750 --> 00:43:59.710 oneness, 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 --> 00:44:05.624 pretty tightly on to what the other
NOTE Confidence: 0.944027375

00:44:05.624 --> 00:44:07.624 groups have called mystical experience.
NOTE Confidence: 0.944027375

00:44:07.630 --> 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 --> 00:44:15.014 and we use the scientific one it's

NOTE Confidence: 0.944027375

00:44:15.014 --> 00:44:16.149 and they're measuring the same thing,

NOTE Confidence: 0.944027375

00:44:16.150 --> 00:44:17.910 but they've got a a,

NOTE Confidence: 0.944027375

00:44:17.910 --> 00:44:21.422 a better name for it I think so And

NOTE Confidence: 0.944027375

00:44:21.422 --> 00:44:23.630 then visionary restructuralization

NOTE Confidence: 0.944027375

00:44:23.630 --> 00:44:28.172 that means really the the perceptual

NOTE Confidence: 0.944027375

00:44:28.172 --> 00:44:32.482 effects you know alters altered

NOTE Confidence: 0.944027375

00:44:32.482 --> 00:44:36.070 perception and then dread of ego.

NOTE Confidence: 0.944027375

00:44:36.070 --> 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 --> 00:44:46.989 or annihilation or whatever.

NOTE Confidence: 0.944027375

00:44:46.990 --> 00:44:48.625 And then the general scores

NOTE Confidence: 0.944027375

00:44:48.625 --> 00:44:50.750 is the sum of those three.

NOTE Confidence: 0.944027375

00:44:50.750 --> 00:44:54.030 So and then looking across the the columns,

NOTE Confidence: 0.944027375

00:44:54.030 --> 00:44:56.670 the 1st is the whole sample.

NOTE Confidence: 0.944027375

00:44:56.670 --> 00:44:59.225 Then we have the whole psilocybin group,

NOTE Confidence: 0.944027375

00:44:59.230 --> 00:45:02.366 44 people who were not abstinent before
NOTE Confidence: 0.944027375

00:45:02.366 --> 00:45:05.838 they got the the the psilocybin now.
NOTE Confidence: 0.944027375

00:45:05.840 --> 00:45:10.000 21 people who were drinking at least at the
NOTE Confidence: 0.8928256632

00:45:12.480 --> 00:45:14.034 medium risk level and then 14
NOTE Confidence: 0.8928256632

00:45:14.034 --> 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 --> 00:45:19.596 gotten the four weeks of therapy.
NOTE Confidence: 0.8928256632

00:45:19.600 --> 00:45:21.850 So because people had improved quite
NOTE Confidence: 0.8928256632

00:45:21.850 --> 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 --> 00:45:29.514 be pulling most of the weight here.
NOTE Confidence: 0.8928256632

00:45:29.520 --> 00:45:31.164 So that the the.
NOTE Confidence: 0.8928256632

00:45:31.164 --> 00:45:33.630 The the correlate you know the
NOTE Confidence: 0.8928256632

00:45:33.720 --> 00:45:35.504 the significant, you know,
NOTE Confidence: 0.8928256632

00:45:35.504 --> 00:45:36.464 I don't know what significance
NOTE Confidence: 0.8928256632

00:45:36.464 --> 00:45:37.360 means in this context,

NOTE Confidence: 0.8928256632

00:45:37.360 --> 00:45:40.665 but the 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 --> 00:45:45.110 bold and red and the ones that

NOTE Confidence: 0.8928256632

00:45:45.110 --> 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 --> 00:45:50.869 them but you could look at the the

NOTE Confidence: 0.8928256632

00:45:50.869 --> 00:45:54.163 correlations just as well and they're

NOTE Confidence: 0.8928256632

00:45:54.163 --> 00:45:56.640 probably more interesting actually,

NOTE Confidence: 0.8928256632

00:45:56.640 --> 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 --> 00:46:02.290 the oceanic boundlessness factor

NOTE Confidence: 0.8928256632

00:46:02.290 --> 00:46:06.010 is correlated with drinks per day

NOTE Confidence: 0.9301902

00:46:08.250 --> 00:46:10.976 at each at each level,

NOTE Confidence: 0.9301902

00:46:10.976 --> 00:46:12.386 and the correlation gets higher

NOTE Confidence: 0.9301902

00:46:12.386 --> 00:46:14.303 as you get to the people who

NOTE Confidence: 0.9301902

00:46:14.303 --> 00:46:15.603 are drinking more and more.
NOTE Confidence: 0.9301902

00:46:15.610 --> 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 --> 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 --> 00:46:28.390 more before you got the psilocybin.
NOTE Confidence: 0.9301902

00:46:28.390 --> 00:46:29.308 So that's so
NOTE Confidence: 0.92616476

00:46:29.310 --> 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 --> 00:46:35.263 people who aren't drinking much
NOTE Confidence: 0.92616476

00:46:35.263 --> 00:46:37.510 don't have far to go that's true.
NOTE Confidence: 0.92616476

00:46:37.510 --> 00:46:39.110 So you would expect increased
NOTE Confidence: 0.938705325384616

00:46:39.750 --> 00:46:40.818 but they could but they couldn't
NOTE Confidence: 0.938705325384616

00:46:40.818 --> 00:46:42.148 go the I mean they could they
NOTE Confidence: 0.943607995

00:46:42.150 --> 00:46:43.470 could have gone the other right.
NOTE Confidence: 0.943607995

00:46:43.470 --> 00:46:45.442 But I would expect larger I would

NOTE Confidence: 0.943607995

00:46:45.442 --> 00:46:47.822 expect larger correlations with a

NOTE Confidence: 0.943607995

00:46:47.822 --> 00:46:49.730 higher baseline for this analysis

NOTE Confidence: 0.943607995

00:46:49.730 --> 00:46:51.388 more room to more room to improve.

NOTE Confidence: 0.943607995

00:46:51.390 --> 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 --> 00:46:57.470 Well, if it's working, you would if it's

NOTE Confidence: 0.938995675

00:46:57.470 --> 00:46:59.350 something you would expect that. But

NOTE Confidence: 0.94427896

00:46:59.630 --> 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 --> 00:47:04.494 It's working better in the people who

NOTE Confidence: 0.94427896

00:47:04.494 --> 00:47:05.910 are drinking more as opposed to just.

NOTE Confidence: 0.94427896

00:47:05.910 --> 00:47:06.855 It's working right.

NOTE Confidence: 0.94427896

00:47:06.855 --> 00:47:08.430 Pretty well across the board.

NOTE Confidence: 0.94427896

00:47:08.430 --> 00:47:09.718 And you see it more in the people

NOTE Confidence: 0.94427896

00:47:09.718 --> 00:47:10.603 that are drinking, right.
NOTE Confidence: 0.94427896

00:47:10.603 --> 00:47:10.949 That's all.
NOTE Confidence: 0.9452853

00:47:11.070 --> 00:47:13.662 Well, it's, yeah, they're just more to do.
NOTE Confidence: 0.9452853

00:47:13.662 --> 00:47:14.842 I mean, there's more benefit
NOTE Confidence: 0.9452853

00:47:14.842 --> 00:47:18.430 to be had, I guess. So, yeah.
NOTE Confidence: 0.9452853

00:47:18.430 --> 00:47:20.390 And then, you know, the other.
NOTE Confidence: 0.948639706666667

00:47:20.880 --> 00:47:22.056 It is interesting that the oceanic
NOTE Confidence: 0.948639706666667

00:47:22.056 --> 00:47:23.280 boundless just jumps out as carrying.
NOTE Confidence: 0.938240864

00:47:24.120 --> 00:47:26.720 Yeah. And the other ones,
NOTE Confidence: 0.938240864

00:47:26.720 --> 00:47:28.335 the correlations don't really even
NOTE Confidence: 0.938240864

00:47:28.335 --> 00:47:30.793 go up except for the general which
NOTE Confidence: 0.938240864

00:47:30.793 --> 00:47:32.570 includes and that's it's oceanic
NOTE Confidence: 0.938240864

00:47:32.570 --> 00:47:34.400 boundless as it's pulling that one.
NOTE Confidence: 0.938240864

00:47:34.400 --> 00:47:38.630 So, so that's so that's interesting.
NOTE Confidence: 0.938240864

00:47:38.630 --> 00:47:40.390 This is the mystical
NOTE Confidence: 0.938240864

00:47:40.390 --> 00:47:41.270 experience questionnaire.

NOTE Confidence: 0.938240864

00:47:41.270 --> 00:47:43.496 So this one there are you know

NOTE Confidence: 0.938240864

00:47:43.496 --> 00:47:45.068 there's there's been factor

NOTE Confidence: 0.938240864

00:47:45.068 --> 00:47:46.709 analyses demonstrating this.

NOTE Confidence: 0.938240864

00:47:46.710 --> 00:47:47.667 This four factor,

NOTE Confidence: 0.938240864

00:47:47.667 --> 00:47:49.900 it's been done a couple of different

NOTE Confidence: 0.938240864

00:47:49.960 --> 00:47:51.822 ways but the sort of most standard

NOTE Confidence: 0.938240864

00:47:51.822 --> 00:47:53.664 one now is these four factors

NOTE Confidence: 0.938240864

00:47:53.664 --> 00:47:56.422 11 is what they which they call

NOTE Confidence: 0.938240864

00:47:56.422 --> 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 --> 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 --> 00:48:11.303 that go separately transcendence

NOTE Confidence: 0.935222015

00:48:11.303 --> 00:48:14.030 of space and time and ineffability.

NOTE Confidence: 0.935222015

00:48:14.030 --> 00:48:15.405 So those I think ineffability
NOTE Confidence: 0.935222015

00:48:15.405 --> 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 --> 00:48:20.309 half of the items and then they're
NOTE Confidence: 0.935222015

00:48:20.309 --> 00:48:22.250 smaller numbers after that but
NOTE Confidence: 0.935222015

00:48:22.250 --> 00:48:25.177 so here you know in the whole
NOTE Confidence: 0.935222015

00:48:25.177 --> 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 --> 00:48:32.910 significant correlations and
NOTE Confidence: 0.935222015

00:48:32.910 --> 00:48:35.520 then as you go across. The
NOTE Confidence: 0.9452853

00:48:37.920 --> 00:48:42.164 they tend to increase to some extent.
NOTE Confidence: 0.9452853

00:48:42.164 --> 00:48:45.982 And so even you know that so that the MEQ 43,
NOTE Confidence: 0.9452853

00:48:45.982 --> 00:48:48.630 that's the old that the old version of
NOTE Confidence: 0.9452853

00:48:48.709 --> 00:48:50.808 the scale where they had and there's
NOTE Confidence: 0.9452853

00:48:50.808 --> 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 --> 00:48:58.020 put those all together, it's all the

NOTE Confidence: 0.9452853

00:48:58.020 --> 00:48:59.920 aspects of the mystical experience.

NOTE Confidence: 0.9452853

00:48:59.920 --> 00:49:02.080 Yeah, you can see those correlations

NOTE Confidence: 0.9452853

00:49:02.080 --> 00:49:03.476 going you know all the way up to.

NOTE Confidence: 0.927129504545455

00:49:05.750 --> 00:49:08.704 Over .5 in the in this smallest

NOTE Confidence: 0.927129504545455

00:49:08.704 --> 00:49:11.390 group and it's it's so it's a

NOTE Confidence: 0.927129504545455

00:49:11.390 --> 00:49:15.750 nice kind of consistent effect

NOTE Confidence: 0.927129504545455

00:49:15.750 --> 00:49:19.590 here and this is another mysticism

NOTE Confidence: 0.927129504545455

00:49:19.590 --> 00:49:22.070 scale that hood mysticism scale.

NOTE Confidence: 0.927129504545455

00:49:22.070 --> 00:49:25.846 And these are I think you know some of

NOTE Confidence: 0.927129504545455

00:49:25.846 --> 00:49:28.090 the largest correlations of all with

NOTE Confidence: 0.927129504545455

00:49:28.170 --> 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 --> 00:49:36.700 With nothing, being one with

NOTE Confidence: 0.927129504545455

00:49:36.700 --> 00:49:39.436 nothingness or kind of going inside

NOTE Confidence: 0.927129504545455

00:49:39.436 --> 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 --> 00:49:48.420 with the universe and the plants
NOTE Confidence: 0.927129504545455

00:49:48.487 --> 00:49:50.257 and the animals and everything.
NOTE Confidence: 0.927129504545455

00:49:50.260 --> 00:49:51.930 And interpretation means that it's
NOTE Confidence: 0.927129504545455

00:49:51.930 --> 00:49:53.600 sort of their religious meaning
NOTE Confidence: 0.927129504545455

00:49:53.655 --> 00:49:54.820 that people give to it.
NOTE Confidence: 0.927129504545455

00:49:54.820 --> 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 --> 00:50:03.470 the more you drank.
NOTE Confidence: 0.927129504545455

00:50:03.470 --> 00:50:06.793 The higher the correlation again and
NOTE Confidence: 0.927129504545455

00:50:06.793 --> 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 --> 00:50:15.665 and trying to so it's made by Rick
NOTE Confidence: 0.927129504545455

00:50:15.665 --> 00:50:17.836 Strassman and he actually based it

NOTE Confidence: 0.927129504545455
00:50:17.836 --> 00:50:21.030 on the the five what are called
NOTE Confidence: 0.927129504545455
00:50:21.030 --> 00:50:24.195 skandas in in Buddhist psychology
NOTE Confidence: 0.927129504545455
00:50:24.195 --> 00:50:26.870 that they're just different aspects
NOTE Confidence: 0.927129504545455
00:50:26.870 --> 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 --> 00:50:33.910 because somatic is one of them.
NOTE Confidence: 0.927129504545455
00:50:33.910 --> 00:50:35.746 So that's not even mind really.
NOTE Confidence: 0.927129504545455
00:50:35.750 --> 00:50:37.082 But so we're made-up of these
NOTE Confidence: 0.927129504545455
00:50:37.082 --> 00:50:37.748 things they say.
NOTE Confidence: 0.927129504545455
00:50:37.750 --> 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 --> 00:50:41.015 physical experience of your body
NOTE Confidence: 0.927129504545455
00:50:41.015 --> 00:50:43.793 that didn't seem to be very strongly
NOTE Confidence: 0.927129504545455
00:50:43.793 --> 00:50:47.990 related to anything affective experience.
NOTE Confidence: 0.927129504545455
00:50:47.990 --> 00:50:50.018 Emotions strongly correlated
NOTE Confidence: 0.927129504545455

00:50:50.018 --> 00:50:52.076 in this case and none of those
NOTE Confidence: 0.927129504545455

00:50:52.076 --> 00:50:54.095 other skills really get at that
NOTE Confidence: 0.927129504545455

00:50:54.095 --> 00:50:56.842 particularly except for positive mood.
NOTE Confidence: 0.927129504545455

00:50:56.842 --> 00:51:01.050 So that's so that one shows some
NOTE Confidence: 0.927129504545455

00:51:01.050 --> 00:51:02.890 strong correlations of perceptual,
NOTE Confidence: 0.927129504545455

00:51:02.890 --> 00:51:05.200 just as this is pretty much what's
NOTE Confidence: 0.927129504545455

00:51:05.200 --> 00:51:08.810 measured in the five DASC under the
NOTE Confidence: 0.927129504545455

00:51:08.810 --> 00:51:11.830 the visionary restructuralization,
NOTE Confidence: 0.927129504545455

00:51:11.830 --> 00:51:14.090 nothing there.
NOTE Confidence: 0.927129504545455

00:51:14.090 --> 00:51:16.930 And then changes in cognition,
NOTE Confidence: 0.927129504545455

00:51:16.930 --> 00:51:18.226 there's, you know,
NOTE Confidence: 0.927129504545455

00:51:18.226 --> 00:51:19.830 moderate sized correlations
NOTE Confidence: 0.927129504545455

00:51:19.830 --> 00:51:22.050 there and volition.
NOTE Confidence: 0.927129504545455

00:51:22.050 --> 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 --> 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 --> 00:51:31.130 not so much in the larger group.
NOTE Confidence: 0.927129504545455
00:51:31.130 --> 00:51:33.725 So I'm not sure that I'd be
NOTE Confidence: 0.927129504545455
00:51:33.725 --> 00:51:34.650 quite as confident in that.
NOTE Confidence: 0.927129504545455
00:51:34.650 --> 00:51:36.190 One is intensity a combination
NOTE Confidence: 0.927129504545455
00:51:36.190 --> 00:51:37.450 of the rest is a separate,
NOTE Confidence: 0.927129504545455
00:51:37.450 --> 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 --> 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 --> 00:51:48.990 it's interesting and it's and it's not
NOTE Confidence: 0.927129504545455
00:51:49.067 --> 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 --> 00:51:57.142 So anyway the conclusion from this I

NOTE Confidence: 0.927129504545455

00:51:57.142 --> 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 --> 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 --> 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 --> 00:52:13.767 if some of these are simply

NOTE Confidence: 0.91988512

00:52:13.767 --> 00:52:15.290 surrogate markers for how much

NOTE Confidence: 0.91988512

00:52:15.290 --> 00:52:17.180 the drug affected the neurons.

NOTE Confidence: 0.91988512

00:52:17.180 --> 00:52:18.380 Yeah right.

NOTE Confidence: 0.91988512

00:52:18.380 --> 00:52:19.864 But the so the specificity if you

NOTE Confidence: 0.91988512

00:52:19.864 --> 00:52:21.498 have this you know more specificity

NOTE Confidence: 0.91988512

00:52:21.500 --> 00:52:22.928 to the mysticism than the other

NOTE Confidence: 0.91988512

00:52:22.928 --> 00:52:24.108 components starts target that they're

NOTE Confidence: 0.91988512

00:52:24.108 --> 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 --> 00:52:29.262 how got in there and how tightly

NOTE Confidence: 0.91988512

00:52:29.262 --> 00:52:30.424 did it bind the receptors and how

NOTE Confidence: 0.91988512

00:52:30.424 --> 00:52:31.492 much did it affect the neurons

NOTE Confidence: 0.943608066666667

00:52:32.900 --> 00:52:35.300 and how are the preparatory sessions

NOTE Confidence: 0.943608066666667

00:52:35.300 --> 00:52:39.046 how are they how are they introduced

NOTE Confidence: 0.943608066666667

00:52:39.046 --> 00:52:44.420 to it they so we we tried to be very.

NOTE Confidence: 0.939582706666667

00:52:46.470 --> 00:52:49.270 Non directive and as far as what kind

NOTE Confidence: 0.939582706666667

00:52:49.270 --> 00:52:52.590 of an experience is a good experience

NOTE Confidence: 0.939582706666667

00:52:52.590 --> 00:52:55.758 but we did we we were willing to

NOTE Confidence: 0.939582706666667

00:52:55.758 --> 00:52:58.155 suggest that you know you you whatever

NOTE Confidence: 0.939582706666667

00:52:58.155 --> 00:53:00.329 experience you have you know that that

NOTE Confidence: 0.939582706666667

00:53:00.329 --> 00:53:02.369 might be the one you need to that

NOTE Confidence: 0.939582706666667

00:53:02.435 --> 00:53:04.571 that's the one for you and it's it's
NOTE Confidence: 0.939582706666667

00:53:04.571 --> 00:53:07.056 you know you may do do with it what
NOTE Confidence: 0.939582706666667

00:53:07.056 --> 00:53:09.324 you can and see if see if you can but
NOTE Confidence: 0.939582706666667

00:53:09.324 --> 00:53:10.683 do you think they fidelity actually
NOTE Confidence: 0.939582706666667

00:53:10.683 --> 00:53:12.891 that was true because that's what is my
NOTE Confidence: 0.939582706666667

00:53:12.945 --> 00:53:16.409 concern is you know if if it's an if then.
NOTE Confidence: 0.939582706666667

00:53:16.410 --> 00:53:18.740 Type situation where you're told
NOTE Confidence: 0.939582706666667

00:53:18.740 --> 00:53:20.290 that if you have this experience,
NOTE Confidence: 0.939582706666667

00:53:20.290 --> 00:53:21.454 you're going to get better.
NOTE Confidence: 0.939582706666667

00:53:21.454 --> 00:53:22.918 And then therefore if I didn't
NOTE Confidence: 0.939582706666667

00:53:22.918 --> 00:53:23.650 have that experience,
NOTE Confidence: 0.939582706666667

00:53:23.650 --> 00:53:25.888 I'm not going to get better.
NOTE Confidence: 0.939582706666667

00:53:25.890 --> 00:53:29.520 Well, so like I said, we, we, you know,
NOTE Confidence: 0.939582706666667

00:53:29.520 --> 00:53:31.290 I don't know what people believe.
NOTE Confidence: 0.939582706666667

00:53:31.290 --> 00:53:31.500 Yeah.
NOTE Confidence: 0.939582706666667

00:53:31.500 --> 00:53:32.970 I mean because we also said it,

NOTE Confidence: 0.939582706666667

00:53:32.970 --> 00:53:36.423 it it's also possible that you know that

NOTE Confidence: 0.939582706666667

00:53:36.423 --> 00:53:38.181 this is going to do something in your brain

NOTE Confidence: 0.939582706666667

00:53:38.181 --> 00:53:39.930 and it may have nothing to do with that.

NOTE Confidence: 0.939582706666667

00:53:39.930 --> 00:53:40.890 So that equipoise,

NOTE Confidence: 0.939582706666667

00:53:40.890 --> 00:53:43.358 I mean so that was so that was

NOTE Confidence: 0.939582706666667

00:53:43.358 --> 00:53:46.396 so that was put out there too.

NOTE Confidence: 0.939582706666667

00:53:46.400 --> 00:53:50.916 I but I think you know I think

NOTE Confidence: 0.939582706666667

00:53:50.916 --> 00:53:52.908 people have expected expectations

NOTE Confidence: 0.939582706666667

00:53:52.908 --> 00:53:55.678 sure and and therapist due to so

NOTE Confidence: 0.939582706666667

00:53:55.680 --> 00:53:57.871 but we we we you know definitely

NOTE Confidence: 0.939582706666667

00:53:57.871 --> 00:53:59.224 we're not privileging mystical

NOTE Confidence: 0.939582706666667

00:53:59.224 --> 00:54:01.360 type experience per se for example

NOTE Confidence: 0.948639696666667

00:54:06.240 --> 00:54:08.556 we probably should wrap up because

NOTE Confidence: 0.948639696666667

00:54:08.560 --> 00:54:11.360 it's getting but but MRI so

NOTE Confidence: 0.922857967

00:54:11.360 --> 00:54:12.392 I'm going to show you this

NOTE Confidence: 0.922857967

00:54:12.392 --> 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 --> 00:54:22.057 of it because this is only 11 subjects,
NOTE Confidence: 0.9402536

00:54:22.060 --> 00:54:25.980 but I think it's it's interesting pilot data.
NOTE Confidence: 0.9402536

00:54:25.980 --> 00:54:30.740 We had a task that evaluated a
NOTE Confidence: 0.9402536

00:54:30.740 --> 00:54:33.868 response to visual alcohol cues
NOTE Confidence: 0.9402536

00:54:33.868 --> 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 --> 00:54:45.061 days after first dose of the drug.
NOTE Confidence: 0.9402536

00:54:45.070 --> 00:54:47.404 And we were you know interested
NOTE Confidence: 0.9402536

00:54:47.404 --> 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 --> 00:54:56.046 the difference in change of those

NOTE Confidence: 0.9402536

00:54:56.046 --> 00:54:58.940 between the two groups and we looked

NOTE Confidence: 0.9402536

00:54:58.940 --> 00:55:03.210 at regions that we thought were most

NOTE Confidence: 0.9402536

00:55:03.339 --> 00:55:06.550 likely to be involved in you know.

NOTE Confidence: 0.965411993

00:55:09.420 --> 00:55:11.760 Craving response and reward processing

NOTE Confidence: 0.965411993

00:55:11.760 --> 00:55:14.100 and and negative emotional processing.

NOTE Confidence: 0.965411993

00:55:14.100 --> 00:55:19.530 So lateral medial Pfc and singular

NOTE Confidence: 0.965411993

00:55:19.530 --> 00:55:21.844 and ventral indoors will stray

NOTE Confidence: 0.965411993

00:55:21.844 --> 00:55:25.540 them and when we found significant

NOTE Confidence: 0.941930728333333

00:55:27.740 --> 00:55:32.186 effects we then only for those.

NOTE Confidence: 0.941930728333333

00:55:32.190 --> 00:55:34.554 Roi's we looked at functional connectivity

NOTE Confidence: 0.941930728333333

00:55:34.554 --> 00:55:37.152 between groups to see if there are

NOTE Confidence: 0.941930728333333

00:55:37.152 --> 00:55:39.150 any differences in between Roi's,

NOTE Confidence: 0.941930728333333

00:55:39.150 --> 00:55:42.470 between no between the groups. So if

NOTE Confidence: 0.98302305

00:55:42.990 --> 00:55:44.510 functional connectivity compared between

NOTE Confidence: 0.91878229125

00:55:44.670 --> 00:55:46.215 groups, yeah, yeah, for the

NOTE Confidence: 0.91878229125

00:55:46.215 --> 00:55:48.638 for the ROI that was that was
NOTE Confidence: 0.91878229125

00:55:48.638 --> 00:55:50.266 different between the groups.
NOTE Confidence: 0.9352219

00:55:53.590 --> 00:55:56.100 And then we also looked at whether
NOTE Confidence: 0.9352219

00:55:56.100 --> 00:55:59.400 these the differences that we found.
NOTE Confidence: 0.9352219

00:55:59.400 --> 00:56:01.280 Had any relationship to the
NOTE Confidence: 0.9352219

00:56:01.280 --> 00:56:03.160 change in drinks per day,
NOTE Confidence: 0.9352219

00:56:03.160 --> 00:56:06.037 so this is a just a summary
NOTE Confidence: 0.9352219

00:56:06.040 --> 00:56:07.600 in terms of the bold contrast.
NOTE Confidence: 0.9352219

00:56:07.600 --> 00:56:12.640 There were some, you know, fairly small
NOTE Confidence: 0.9654121

00:56:15.160 --> 00:56:17.744 clusters in right ventrolateral
NOTE Confidence: 0.9654121

00:56:17.744 --> 00:56:19.036 prefrontal cortex,
NOTE Confidence: 0.9654121

00:56:19.040 --> 00:56:22.008 left dorsolateral prefrontal cortex,
NOTE Confidence: 0.9654121

00:56:22.008 --> 00:56:24.234 and left caudate.
NOTE Confidence: 0.9654121

00:56:24.240 --> 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 --> 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

NOTE Confidence: 0.89145567

00:56:34.360 --> 00:56:36.980 areas in left medial prefrontal

NOTE Confidence: 0.89145567

00:56:36.980 --> 00:56:40.120 and that were also left odd eight.

NOTE Confidence: 0.945620746666667

00:56:40.240 --> 00:56:41.998 I'm sorry, is this these task

NOTE Confidence: 0.945620746666667

00:56:41.998 --> 00:56:43.639 data from the before psilocybin

NOTE Confidence: 0.945620746666667

00:56:43.639 --> 00:56:45.279 or the after psilocybin?

NOTE Confidence: 0.945620746666667

00:56:45.280 --> 00:56:48.572 It's where the after minus before.

NOTE Confidence: 0.945620746666667

00:56:48.572 --> 00:56:50.900 So it's task dependent

NOTE Confidence: 0.945620746666667

00:56:50.900 --> 00:56:52.520 activation after minus.

NOTE Confidence: 0.945620746666667

00:56:52.520 --> 00:56:54.840 Identical task, dependent activation,

NOTE Confidence: 0.945620746666667

00:56:54.840 --> 00:56:56.920 yeah between groups and then relative

NOTE Confidence: 0.957025933333333

00:56:56.920 --> 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 --> 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

NOTE Confidence: 0.78374635

00:57:12.520 --> 00:57:16.352 alcohol Q responding.
NOTE Confidence: 0.78374635

00:57:16.352 --> 00:57:18.312 Areas that that did have
NOTE Confidence: 0.78374635

00:57:18.312 --> 00:57:19.488 some connectivity changes,
NOTE Confidence: 0.78374635

00:57:19.490 --> 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 --> 00:57:24.114 again let's not get too carried
NOTE Confidence: 0.78374635

00:57:24.114 --> 00:57:26.690 away here, but this is this
NOTE Confidence: 0.945045695238095

00:57:28.770 --> 00:57:30.214 right intralateral Pfc area
NOTE Confidence: 0.945045695238095

00:57:30.214 --> 00:57:32.917 and using that as a seed then
NOTE Confidence: 0.945045695238095

00:57:32.917 --> 00:57:34.753 there was increased functional
NOTE Confidence: 0.945045695238095

00:57:34.753 --> 00:57:37.330 connectivity with this area in the
NOTE Confidence: 0.22011086

00:57:39.690 --> 00:57:43.490 IFS lateral pre central gyrus there.
NOTE Confidence: 0.936228176

00:57:45.780 --> 00:57:49.620 This is the area in the left dorsal
NOTE Confidence: 0.936228176

00:57:49.620 --> 00:57:54.980 lateral Pfc and this is the left caudate,
NOTE Confidence: 0.936228176

00:57:54.980 --> 00:57:57.140 which yeah, this was,
NOTE Confidence: 0.936228176

00:57:57.140 --> 00:57:59.156 this was a bit of a surprise and

NOTE Confidence: 0.936228176

00:57:59.156 --> 00:58:01.270 you know we didn't really expect to

NOTE Confidence: 0.936228176

00:58:01.270 --> 00:58:03.944 see greater reactivity in dorsal

NOTE Confidence: 0.936228176

00:58:03.944 --> 00:58:06.308 strayatum as being associated

NOTE Confidence: 0.936228176

00:58:06.308 --> 00:58:08.100 with therapeutic response.

NOTE Confidence: 0.936228176

00:58:08.100 --> 00:58:12.260 So you know, I'm not sure what to make of it.

NOTE Confidence: 0.936228176

00:58:12.260 --> 00:58:13.580 It also showed greater

NOTE Confidence: 0.936228176

00:58:13.580 --> 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 --> 00:58:22.540 there was for that one.

NOTE Confidence: 0.8639468

00:58:22.540 --> 00:58:25.140 There was a significant relationship

NOTE Confidence: 0.8639468

00:58:25.140 --> 00:58:30.300 between that that effect and the the

NOTE Confidence: 0.8639468

00:58:30.300 --> 00:58:33.500 functional connectivity now and decreased

NOTE Confidence: 0.8639468

00:58:33.500 --> 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 --> 00:58:45.659 Yeah, the the left medial prefrontal area

NOTE Confidence: 0.8639468

00:58:45.659 --> 00:58:48.953 and which this increase in activation
NOTE Confidence: 0.8639468

00:58:48.953 --> 00:58:51.909 there was associated with decreased
NOTE Confidence: 0.9805072

00:58:54.270 --> 00:58:58.310 drinking and then this is the left
NOTE Confidence: 0.9805072

00:58:58.310 --> 00:59:02.075 caudate again, which was no functional
NOTE Confidence: 0.9805072

00:59:02.075 --> 00:59:04.338 connectivity changes in this case,
NOTE Confidence: 0.9805072

00:59:04.338 --> 00:59:06.720 but it was associated also with
NOTE Confidence: 0.9805072

00:59:06.797 --> 00:59:10.310 the decrease in drinking, so.
NOTE Confidence: 0.9805072

00:59:10.310 --> 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 --> 00:59:17.150 from the dosing to the scanning,
NOTE Confidence: 0.9805072

00:59:17.150 --> 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 --> 00:59:25.632 these are pretty large effects given
NOTE Confidence: 0.9805072

00:59:25.632 --> 00:59:27.714 you know that we see that we're seeing
NOTE Confidence: 0.9805072

00:59:27.714 --> 00:59:29.280 anything with with samples of this

NOTE Confidence: 0.9805072

00:59:29.330 --> 00:59:30.870 size and they kind of make sense.

NOTE Confidence: 0.9805072

00:59:30.870 --> 00:59:32.590 I mean they're in regions that make sense,

NOTE Confidence: 0.9805072

00:59:32.590 --> 00:59:34.762 they weren't in you know just

NOTE Confidence: 0.9805072

00:59:34.762 --> 00:59:37.319 kind of bunch of random spots.

NOTE Confidence: 0.9805072

00:59:37.320 --> 00:59:39.270 Some of them were accompanied by

NOTE Confidence: 0.9805072

00:59:39.270 --> 00:59:40.245 functional connectivity changes

NOTE Confidence: 0.9805072

00:59:40.245 --> 00:59:42.280 and some of them were correlated

NOTE Confidence: 0.9805072

00:59:42.280 --> 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 --> 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 --> 00:59:59.531 a a prime candidate for that being

NOTE Confidence: 0.941371755555556

00:59:59.531 --> 01:00:02.250 very small and you know the the

NOTE Confidence: 0.941371755555556

01:00:02.250 --> 01:00:04.102 directionality was not exactly.

NOTE Confidence: 0.941371755555556

01:00:04.102 --> 01:00:07.819 Entirely in some cases it was with
NOTE Confidence: 0.941371755555556

01:00:07.819 --> 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 --> 01:00:17.343 and the and the dorsal straight
NOTE Confidence: 0.941371755555556

01:00:17.343 --> 01:00:19.954 and we didn't really expect to see
NOTE Confidence: 0.941371755555556

01:00:19.954 --> 01:00:22.410 increases in the in this especially
NOTE Confidence: 0.941371755555556

01:00:22.410 --> 01:00:24.490 for the alcohol accused so,
NOTE Confidence: 0.941371755555556

01:00:24.490 --> 01:00:28.004 so we'll see but you know executive
NOTE Confidence: 0.941371755555556

01:00:28.010 --> 01:00:30.100 functioning is not all about.
NOTE Confidence: 0.941371755555556

01:00:30.100 --> 01:00:32.011 Just you know inhibition there might be
NOTE Confidence: 0.941371755555556

01:00:32.011 --> 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 --> 01:00:38.611 we'll we'll follow up on it and we're
NOTE Confidence: 0.941371755555556

01:00:38.611 --> 01:00:43.572 really hoping very soon to be able to
NOTE Confidence: 0.941371755555556

01:00:43.572 --> 01:00:47.092 do a much more you know much larger
NOTE Confidence: 0.941371755555556

01:00:47.092 --> 01:00:49.540 and more sophisticated version of this

NOTE Confidence: 0.941371755555556
01:00:49.540 --> 01:00:52.746 with Regina's help and NA AAA funding
NOTE Confidence: 0.941371755555556
01:00:52.746 --> 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 --> 01:01:03.480 So overall you know I think looking good
NOTE Confidence: 0.941371755555556
01:01:03.480 --> 01:01:06.680 for efficacy we need to do more there
NOTE Confidence: 0.941371755555556
01:01:06.760 --> 01:01:08.964 obviously and there's some evidence
NOTE Confidence: 0.941371755555556
01:01:08.964 --> 01:01:11.589 that these drugs are acting across all
NOTE Confidence: 0.941371755555556
01:01:11.589 --> 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 --> 01:01:22.530 this is really why I like it is because.
NOTE Confidence: 0.941371755555556
01:01:22.530 --> 01:01:24.706 I mean you know we do want to
NOTE Confidence: 0.941371755555556
01:01:24.706 --> 01:01:26.975 be able to reduce things to you
NOTE Confidence: 0.941371755555556
01:01:26.975 --> 01:01:28.445 know the level of the receptor.
NOTE Confidence: 0.941371755555556

01:01:28.450 --> 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 --> 01:01:34.210 we don't do biology,
NOTE Confidence: 0.941371755555556

01:01:34.210 --> 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 --> 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 --> 01:01:41.810 and it still should be scientific
NOTE Confidence: 0.941371755555556

01:01:41.810 --> 01:01:43.610 and it still should need to
NOTE Confidence: 0.941371755555556

01:01:43.610 --> 01:01:44.810 be rigorous and makes sense.
NOTE Confidence: 0.941371755555556

01:01:44.810 --> 01:01:47.033 But we need to find ways to you know
NOTE Confidence: 0.941371755555556

01:01:47.033 --> 01:01:49.673 looking at 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 --> 01:01:53.087 to really make sense of this so.
NOTE Confidence: 0.941371755555556

01:01:53.090 --> 01:01:57.690 So that's that thank you wonderful.
NOTE Confidence: 0.941371755555556

01:01:57.690 --> 01:01:59.328 It's really nice to see the new,

NOTE Confidence: 0.941371755555556
01:01:59.330 --> 01:02:00.050 the new stuff.
NOTE Confidence: 0.941371755555556
01:02:00.050 --> 01:02:00.290 So
NOTE Confidence: 0.855730071428571
01:02:01.130 --> 01:02:02.682 worth worth worth staying
NOTE Confidence: 0.855730071428571
01:02:02.682 --> 01:02:04.490 later enough today afternoon.
NOTE Confidence: 0.957025933333333
01:02:06.530 --> 01:02:09.514 Anybody still there? Well that's what I
NOTE Confidence: 0.957025933333333
01:02:09.514 --> 01:02:12.370 was checking out and the answer was yes.
NOTE Confidence: 0.938815971428571
01:02:15.890 --> 01:02:17.698 So do we have you know question or
NOTE Confidence: 0.938815971428571
01:02:17.698 --> 01:02:19.622 two or any comments for for Michael
NOTE Confidence: 0.938815971428571
01:02:19.622 --> 01:02:21.530 before we we call that tonight?
NOTE Confidence: 0.9301902
01:02:24.300 --> 01:02:25.220 Hello. Hi, I have
NOTE Confidence: 0.957025933333333
01:02:25.220 --> 01:02:26.180 two quick questions.
NOTE Confidence: 0.941691228571429
01:02:29.220 --> 01:02:30.588 Thank you so much for the
NOTE Confidence: 0.941691228571429
01:02:30.588 --> 01:02:31.660 presentation and the great work.
NOTE Confidence: 0.867295068
01:02:33.060 --> 01:02:34.860 I have two quick questions. One is that
NOTE Confidence: 0.867295068
01:02:34.860 --> 01:02:37.058 do you have did you collect any data
NOTE Confidence: 0.853676106666667

01:02:37.220 --> 01:02:39.824 after the like between those one and
NOTE Confidence: 0.853676106666667

01:02:39.824 --> 01:02:43.107 two after the sign in and like how
NOTE Confidence: 0.853676106666667

01:02:43.107 --> 01:02:45.403 long can you comment on how long after
NOTE Confidence: 0.853676106666667

01:02:45.403 --> 01:02:47.700 the first dose you could see any
NOTE Confidence: 0.853676106666667

01:02:47.700 --> 01:02:50.920 changes in the outcome measures and?
NOTE Confidence: 0.853676106666667

01:02:50.920 --> 01:02:54.776 If like how long do they last after
NOTE Confidence: 0.853676106666667

01:02:54.776 --> 01:02:57.240 the first psilocybin administration?
NOTE Confidence: 0.853676106666667

01:02:57.240 --> 01:02:59.812 And the second question is that why
NOTE Confidence: 0.853676106666667

01:02:59.812 --> 01:03:01.856 the second dose of the psilocybin has
NOTE Confidence: 0.853676106666667

01:03:01.856 --> 01:03:03.720 higher dose compared to first one.
NOTE Confidence: 0.849624371428572

01:03:04.320 --> 01:03:07.078 OK, yeah. So that's a good question.
NOTE Confidence: 0.849624371428572

01:03:07.080 --> 01:03:08.136 So for the first,
NOTE Confidence: 0.849624371428572

01:03:08.136 --> 01:03:11.920 as far as the first question, we there's a,
NOTE Confidence: 0.849624371428572

01:03:11.920 --> 01:03:14.506 there was a follow up at one week
NOTE Confidence: 0.849624371428572

01:03:14.506 --> 01:03:16.502 after the first psilocybin dose,
NOTE Confidence: 0.849624371428572

01:03:16.502 --> 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 --> 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 --> 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 --> 01:03:29.632 you're less depressed.
NOTE Confidence: 0.849624371428572
01:03:29.632 --> 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 --> 01:03:35.104 you know, I don't really care if
NOTE Confidence: 0.849624371428572
01:03:35.104 --> 01:03:36.966 somebody's got less craving 8
NOTE Confidence: 0.849624371428572
01:03:36.966 --> 01:03:38.696 hours after they took psilocybin.
NOTE Confidence: 0.849624371428572
01:03:38.700 --> 01:03:40.924 So maybe we should have done it the next
NOTE Confidence: 0.849624371428572
01:03:40.924 --> 01:03:42.924 day and then I am going to do that in
NOTE Confidence: 0.849624371428572
01:03:42.924 --> 01:03:44.936 the next study and just just just to see,
NOTE Confidence: 0.849624371428572
01:03:44.940 --> 01:03:45.866 you know,
NOTE Confidence: 0.849624371428572

01:03:45.866 --> 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 --> 01:03:53.620 know most of these self report things
NOTE Confidence: 0.849624371428572

01:03:53.620 --> 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 --> 01:04:02.064 does seem to the effects are a little
NOTE Confidence: 0.849624371428572

01:04:02.064 --> 01:04:04.660 larger far a little farther out it
NOTE Confidence: 0.849624371428572

01:04:04.660 --> 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 --> 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 --> 01:04:18.140 is why do we go up on the dose.
NOTE Confidence: 0.93019015

01:04:18.140 --> 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 --> 01:04:26.695 complicated because people got to get

NOTE Confidence: 0.93019015

01:04:26.695 --> 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 --> 01:04:33.910 to be able to titrate people especially

NOTE Confidence: 0.93019015

01:04:33.910 --> 01:04:36.506 if you think you know they need to have.

NOTE Confidence: 0.93019015

01:04:36.510 --> 01:04:38.390 A really strong experience

NOTE Confidence: 0.93019015

01:04:38.390 --> 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 --> 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 --> 01:04:46.036 you want to maximize the dose

NOTE Confidence: 0.93019015

01:04:46.036 --> 01:04:47.484 without you know hurting people

NOTE Confidence: 0.93019015

01:04:47.484 --> 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 --> 01:04:51.809 So with the doses were 25 milligrams

NOTE Confidence: 0.93019015

01:04:51.809 --> 01:04:53.870 and then it was 30 or 40 depending
NOTE Confidence: 0.93019015

01:04:53.870 --> 01:04:56.030 on how robust their response was.
NOTE Confidence: 0.93019015

01:04:56.030 --> 01:04:59.012 And so we gave about 15 people
NOTE Confidence: 0.93019015

01:04:59.012 --> 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 --> 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 --> 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 --> 01:05:13.478 somebody who walked in off the
NOTE Confidence: 0.936228132

01:05:13.478 --> 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 --> 01:05:18.408 some people are really
NOTE Confidence: 0.936228132

01:05:18.408 --> 01:05:19.693 challenged by the 25 milligrams.
NOTE Confidence: 0.936228132

01:05:19.700 --> 01:05:21.056 Is there any thought that you,

NOTE Confidence: 0.936228132

01:05:21.060 --> 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 --> 01:05:26.485 clear there is an inverted at least

NOTE Confidence: 0.936228132

01:05:26.485 --> 01:05:28.516 in rodents probably in units that you

NOTE Confidence: 0.936228132

01:05:28.516 --> 01:05:30.532 can bypass that sort of sweet spot.

NOTE Confidence: 0.936228132

01:05:30.540 --> 01:05:30.863 Yeah.

NOTE Confidence: 0.936228132

01:05:30.863 --> 01:05:34.130 I have not seen any evidence of that at all.

NOTE Confidence: 0.936228132

01:05:34.130 --> 01:05:37.410 I and yeah I couldn't but I've gone

NOTE Confidence: 0.936228132

01:05:37.410 --> 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 --> 01:05:48.566 they this is in the in America.

NOTE Confidence: 0.936228132

01:05:48.570 --> 01:05:50.250 I mean it was different in Europe where

NOTE Confidence: 0.936228132

01:05:50.250 --> 01:05:51.934 people are more sophisticated than subtle.

NOTE Confidence: 0.936228132

01:05:51.934 --> 01:05:53.886 But it was like you know we're going

NOTE Confidence: 0.936228132

01:05:53.886 --> 01:05:55.845 to just blow them out of the water.
NOTE Confidence: 0.936228132

01:05:55.850 --> 01:05:57.368 And it was like so one,
NOTE Confidence: 0.936228132

01:05:57.370 --> 01:05:59.154 one time get it you know one time
NOTE Confidence: 0.936228132

01:05:59.154 --> 01:06:00.717 get them ready and then just.
NOTE Confidence: 0.936228132

01:06:00.720 --> 01:06:03.276 You know the whole point is ego death right.
NOTE Confidence: 0.936228132

01:06:03.280 --> 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 --> 01:06:15.654 I mean that's not really true
NOTE Confidence: 0.936228132

01:06:15.654 --> 01:06:17.652 but you're you're conscious your
NOTE Confidence: 0.936228132

01:06:17.652 --> 01:06:21.600 consciousness is going to be obliterated.
NOTE Confidence: 0.936228132

01:06:21.600 --> 01:06:25.962 So but they just made that up so we don't
NOTE Confidence: 0.936228132

01:06:25.962 --> 01:06:29.240 know if that's necessary or not and so the.
NOTE Confidence: 0.936228132

01:06:29.240 --> 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 --> 01:06:36.400 better until you start hurting, right.

NOTE Confidence: 0.936228132

01:06:36.400 --> 01:06:38.344 So that's to maximize the dose

NOTE Confidence: 0.936228132

01:06:38.344 --> 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 --> 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 --> 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 --> 01:06:58.340 I had a question about

NOTE Confidence: 0.929773064285714

01:06:58.340 --> 01:06:59.730 and I may have missed it,

NOTE Confidence: 0.929773064285714

01:06:59.730 --> 01:07:02.370 When you talked about the study

NOTE Confidence: 0.929773064285714

01:07:02.370 --> 01:07:05.640 the the with the mouse model where

NOTE Confidence: 0.929773064285714

01:07:05.640 --> 01:07:07.950 they blocked with with Catanzeran.

NOTE Confidence: 0.929773064285714

01:07:07.950 --> 01:07:09.875 Did they do a dose response
NOTE Confidence: 0.929773064285714

01:07:09.875 --> 01:07:10.730 with the Catanzeran?
NOTE Confidence: 0.933544666666667

01:07:10.810 --> 01:07:14.842 Was it just a single dose they they
NOTE Confidence: 0.933544666666667

01:07:14.842 --> 01:07:17.635 found a dose I think I mean they they
NOTE Confidence: 0.933544666666667

01:07:17.635 --> 01:07:20.099 might have done some pilot work but
NOTE Confidence: 0.933544666666667

01:07:20.099 --> 01:07:24.065 they used a dose which was reliably.
NOTE Confidence: 0.933544666666667

01:07:24.065 --> 01:07:28.052 Blocking the head twitch and in at
NOTE Confidence: 0.933544666666667

01:07:28.052 --> 01:07:30.749 least one of the studies that also
NOTE Confidence: 0.933544666666667

01:07:30.749 --> 01:07:34.670 blocked the characteristic EE G
NOTE Confidence: 0.933544666666667

01:07:34.670 --> 01:07:39.480 changes which is you know decreased
NOTE Confidence: 0.933544666666667

01:07:39.480 --> 01:07:44.225 power and gamma that goes that
NOTE Confidence: 0.933544666666667

01:07:44.225 --> 01:07:46.355 that's that particularly goes down so
NOTE Confidence: 0.933544666666667

01:07:46.355 --> 01:07:48.369 they're in in humans and in animals
NOTE Confidence: 0.933544666666667

01:07:48.369 --> 01:07:50.147 that you can that it correlates
NOTE Confidence: 0.933544666666667

01:07:50.147 --> 01:07:53.892 with the subjective effect so but.
NOTE Confidence: 0.933544666666667

01:07:53.892 --> 01:07:59.040 You know we we know that people you know

NOTE Confidence: 0.907236928

01:08:01.800 --> 01:08:03.795 it's 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 --> 01:08:08.860 you know all of the 582 A receptors

NOTE Confidence: 0.907236928

01:08:08.860 --> 01:08:12.440 were were blocked obviously and people

NOTE Confidence: 0.941153033809524

01:08:14.480 --> 01:08:15.976 the serotonin receptors are

NOTE Confidence: 0.941153033809524

01:08:15.976 --> 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 --> 01:08:24.280 know like 10 milligrams right so.

NOTE Confidence: 0.941153033809524

01:08:24.280 --> 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 --> 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 --> 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 --> 01:08:49.198 I don't think anybody knows what it is.
NOTE Confidence: 0.946004114285714

01:08:49.200 --> 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 --> 01:08:54.706 The receptors weren't seeing any psilocybin.
NOTE Confidence: 0.946004114285714

01:08:54.710 --> 01:08:57.484 I mean I'm sure they were just if
NOTE Confidence: 0.946004114285714

01:08:57.484 --> 01:08:59.519 they were blocked enough blocked
NOTE Confidence: 0.946004114285714

01:08:59.519 --> 01:09:01.668 enough of them that they didn't
NOTE Confidence: 0.946004114285714

01:09:01.668 --> 01:09:03.342 get the head twitch and they
NOTE Confidence: 0.946004114285714

01:09:03.342 --> 01:09:05.110 didn't get that the EEG. So it's
NOTE Confidence: 0.919147976

01:09:06.550 --> 01:09:08.638 yeah, the reason I asked I'm I'm I'm
NOTE Confidence: 0.919147976

01:09:08.638 --> 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 --> 01:09:15.905 it's going to be very complex mix

NOTE Confidence: 0.919147976
01:09:15.905 --> 01:09:18.425 but from from some work I did back.
NOTE Confidence: 0.938576366666667
01:09:20.860 --> 01:09:21.958 Back in grad school and I
NOTE Confidence: 0.938576366666667
01:09:21.958 --> 01:09:23.220 didn't do the mouse head twitch,
NOTE Confidence: 0.938576366666667
01:09:23.220 --> 01:09:24.977 but I did the rabbit head Bob
NOTE Confidence: 0.938576366666667
01:09:24.980 --> 01:09:26.812 which is somewhat analogous.
NOTE Confidence: 0.938576366666667
01:09:26.812 --> 01:09:28.772 There are differences and how
NOTE Confidence: 0.938576366666667
01:09:28.772 --> 01:09:30.873 much of the antagonist you
NOTE Confidence: 0.938576366666667
01:09:30.873 --> 01:09:33.192 needed for the head Bob versus.
NOTE Confidence: 0.938576366666667
01:09:33.192 --> 01:09:34.900 I also looked at different
NOTE Confidence: 0.938576366666667
01:09:34.900 --> 01:09:35.828 behavioral outcome,
NOTE Confidence: 0.938576366666667
01:09:35.828 --> 01:09:37.220 also Pi hydrolysis,
NOTE Confidence: 0.938576366666667
01:09:37.220 --> 01:09:38.535 but it was different for
NOTE Confidence: 0.938576366666667
01:09:38.535 --> 01:09:39.480 the different outcomes.
NOTE Confidence: 0.938576366666667
01:09:39.480 --> 01:09:41.220 So I just wonder whether yes,
NOTE Confidence: 0.938576366666667
01:09:41.220 --> 01:09:42.500 it makes sense to like pick a dose
NOTE Confidence: 0.938576366666667

01:09:42.500 --> 01:09:43.736 that you'd think it's going to
NOTE Confidence: 0.938576366666667

01:09:43.736 --> 01:09:44.771 be big enough because otherwise
NOTE Confidence: 0.938576366666667

01:09:44.815 --> 01:09:46.480 your place can be just way too
NOTE Confidence: 0.938576366666667

01:09:46.480 --> 01:09:48.260 big and take much too long.
NOTE Confidence: 0.938576366666667

01:09:48.260 --> 01:09:51.179 But I wonder whether you know whether
NOTE Confidence: 0.938576366666667

01:09:51.180 --> 01:09:53.900 it just needed a higher dose or I
NOTE Confidence: 0.938576366666667

01:09:53.900 --> 01:09:55.780 mean there's also all these off,
NOTE Confidence: 0.938576366666667

01:09:55.780 --> 01:09:59.950 off, not off target because the
NOTE Confidence: 0.938576366666667

01:09:59.950 --> 01:10:01.700 transfer is not all that clean either
NOTE Confidence: 0.95283285

01:10:02.580 --> 01:10:03.820 maybe some other targets.
NOTE Confidence: 0.93421556

01:10:03.820 --> 01:10:07.619 But but but also this, you know raises
NOTE Confidence: 0.93421556

01:10:07.619 --> 01:10:09.660 the idea that even if it's going
NOTE Confidence: 0.94931066

01:10:09.660 --> 01:10:11.564 for the same receptor, they could be
NOTE Confidence: 0.94931066

01:10:11.564 --> 01:10:12.806 different on downstream pathways that
NOTE Confidence: 0.94931066

01:10:12.806 --> 01:10:14.297 are being activated or that are being
NOTE Confidence: 0.94931066

01:10:14.300 --> 01:10:16.340 blocked by by certain antagonists.

NOTE Confidence: 0.944566485714286
01:10:17.240 --> 01:10:19.040 And so that the therapeutic
NOTE Confidence: 0.944566485714286
01:10:19.040 --> 01:10:20.560 effects may still be coming from
NOTE Confidence: 0.944566485714286
01:10:20.560 --> 01:10:21.760 the same receptor, but maybe
NOTE Confidence: 0.9402536
01:10:21.760 --> 01:10:23.356 not not the same bonds genius
NOTE Confidence: 0.94830432
01:10:24.760 --> 01:10:26.040 signal or maybe there's also
NOTE Confidence: 0.88849622
01:10:26.960 --> 01:10:27.904 collo colloquialization
NOTE Confidence: 0.88849622
01:10:27.904 --> 01:10:29.880 of the perceptor. So it's
NOTE Confidence: 0.953192257142857
01:10:30.320 --> 01:10:32.154 obviously highly complex and you can't tell
NOTE Confidence: 0.93824092
01:10:32.160 --> 01:10:34.520 this from one now study,
NOTE Confidence: 0.93824092
01:10:34.520 --> 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 --> 01:10:38.760 you know, by that finding.
NOTE Confidence: 0.93824092
01:10:38.760 --> 01:10:41.200 But I still think that it's it is possible
NOTE Confidence: 0.93824092
01:10:41.200 --> 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 --> 01:10:51.275 I would guess that it that it is.
NOTE Confidence: 0.93824092

01:10:51.280 --> 01:10:53.383 Yeah. And at least in some of
NOTE Confidence: 0.93824092

01:10:53.383 --> 01:10:54.650 these at least in you know some
NOTE Confidence: 0.93824092

01:10:54.701 --> 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 --> 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 --> 01:11:03.870 know but and I mean we do know
NOTE Confidence: 0.93824092

01:11:03.870 --> 01:11:05.958 that different to a agonists
NOTE Confidence: 0.9855389

01:11:08.880 --> 01:11:10.160 activate different.
NOTE Confidence: 0.91051366

01:11:12.240 --> 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 --> 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 --> 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 --> 01:11:34.240 there there really could be,

NOTE Confidence: 0.91051366

01:11:34.240 --> 01:11:36.280 I mean principle these things

NOTE Confidence: 0.91051366

01:11:36.280 --> 01:11:38.480 might be separable or.

NOTE Confidence: 0.91051366

01:11:38.480 --> 01:11:39.280 But they might not break.

NOTE Confidence: 0.91051366

01:11:39.280 --> 01:11:40.950 So but it's definitely something

NOTE Confidence: 0.91051366

01:11:40.950 --> 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 --> 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 --> 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 --> 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 --> 01:12:11.652 They decide the low dose was was

NOTE Confidence: 0.91051366

01:12:11.652 --> 01:12:13.470 not good but the the one study

NOTE Confidence: 0.91051366

01:12:13.470 --> 01:12:15.227 that used the 75 or 80 milligram

NOTE Confidence: 0.91051366

01:12:15.227 --> 01:12:16.967 dose had it had better outcomes

NOTE Confidence: 0.91051366

01:12:16.967 --> 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 --> 01:12:27.910 really determined what the optimal

NOTE Confidence: 0.91051366

01:12:27.910 --> 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 --> 01:12:38.460 So I want to thank everyone who stuck it

NOTE Confidence: 0.9402536

01:12:38.460 --> 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 --> 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 --> 01:12:48.550 and but no this just

NOTE Confidence: 0.93421556

01:12:48.550 --> 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 --> 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 --> 01:12:55.914 Have a good weekend in there

NOTE Confidence: 0.93421556

01:12:55.914 --> 01:12:57.224 and have a good weekend.

NOTE Confidence: 0.93421556

01:12:57.230 --> 01:12:57.870 Thank you.