

WEBVTT

NOTE duration:"01:16:06"

NOTE recognizability:0.784

NOTE language:en-us

NOTE Confidence: 0.746227261666667

00:00:00.000 --> 00:00:01.080 And and they were both OK.

NOTE Confidence: 0.6382003

00:00:03.360 --> 00:00:05.158 All right. Hello everyone. Thank

NOTE Confidence: 0.96179443

00:00:05.160 --> 00:00:06.685 you for coming to our

NOTE Confidence: 0.96179443

00:00:06.685 --> 00:00:08.120 psychedelic seminar for December.

NOTE Confidence: 0.96179443

00:00:08.120 --> 00:00:11.352 We have our own Jerry Sanacora and

NOTE Confidence: 0.96179443

00:00:11.352 --> 00:00:14.510 Ben Kilmendi here today who ran our

NOTE Confidence: 0.96179443

00:00:14.510 --> 00:00:17.090 sites Yale sites contribution to

NOTE Confidence: 0.96179443

00:00:17.090 --> 00:00:20.970 the bona funded study of multi site

NOTE Confidence: 0.96179443

00:00:20.970 --> 00:00:23.520 study of psilocybin in the treatment

NOTE Confidence: 0.96179443

00:00:23.520 --> 00:00:25.680 of depression which was published in

NOTE Confidence: 0.96179443

00:00:25.680 --> 00:00:28.480 JAMA Psychiatry earlier this year.

NOTE Confidence: 0.96179443

00:00:28.480 --> 00:00:30.940 And so I I thought it'd be great to have

NOTE Confidence: 0.96179443

00:00:31.001 --> 00:00:33.393 Jerry and Ben tell us about that study,

NOTE Confidence: 0.96179443

00:00:33.400 --> 00:00:35.140 the design, the results and some
NOTE Confidence: 0.96179443

00:00:35.140 --> 00:00:37.154 of the the issues that come up in
NOTE Confidence: 0.96179443

00:00:37.154 --> 00:00:38.666 next steps from that program of
NOTE Confidence: 0.96179443

00:00:38.666 --> 00:00:40.724 research which is one of the sort of
NOTE Confidence: 0.96179443

00:00:40.724 --> 00:00:42.416 one of the leading programs that's
NOTE Confidence: 0.96179443

00:00:42.416 --> 00:00:44.320 moving towards the potential for
NOTE Confidence: 0.96179443

00:00:44.320 --> 00:00:46.358 approval of psychedelic treatments.
NOTE Confidence: 0.96179443

00:00:46.358 --> 00:00:48.204 So very current, very important.
NOTE Confidence: 0.96179443

00:00:48.204 --> 00:00:49.314 Thank you for being here.
NOTE Confidence: 0.466014546

00:00:49.520 --> 00:00:52.160 All right, thank you. Thanks Chris. Well,
NOTE Confidence: 0.944610498461538

00:00:52.160 --> 00:00:54.368 thank you all. Let me just make sure
NOTE Confidence: 0.944610498461538

00:00:54.368 --> 00:00:56.485 we can share our screen. Jessica,
NOTE Confidence: 0.944610498461538

00:00:56.485 --> 00:00:59.160 I don't know if there's a there we
NOTE Confidence: 0.750009934

00:00:59.360 --> 00:01:00.880 go should be able to
NOTE Confidence: 0.6428473275

00:01:01.760 --> 00:01:02.920 looks good now and.
NOTE Confidence: 0.59666913625

00:01:10.240 --> 00:01:12.080 OK, all right. That should be a Ben.

NOTE Confidence: 0.629248604285714
00:01:12.360 --> 00:01:14.397 OK yeah. I'm going to start up.
NOTE Confidence: 0.629248604285714
00:01:14.400 --> 00:01:16.764 So we're going to do something
NOTE Confidence: 0.629248604285714
00:01:16.764 --> 00:01:18.980 differently in a sense that Jerry
NOTE Confidence: 0.629248604285714
00:01:18.980 --> 00:01:21.427 and I are going to split the talk.
NOTE Confidence: 0.629248604285714
00:01:21.427 --> 00:01:23.138 I'm going to go over the first half
NOTE Confidence: 0.629248604285714
00:01:23.138 --> 00:01:24.640 of the talk, which is going to,
NOTE Confidence: 0.629248604285714
00:01:24.640 --> 00:01:26.020 it's going to cover more or
NOTE Confidence: 0.629248604285714
00:01:26.070 --> 00:01:27.516 less the same things that are
NOTE Confidence: 0.939710803333333
00:01:29.640 --> 00:01:32.316 no longer unfamiliar to this group.
NOTE Confidence: 0.939710803333333
00:01:32.320 --> 00:01:34.511 And so I'm going to cruise through
NOTE Confidence: 0.939710803333333
00:01:34.511 --> 00:01:37.239 so we can get to sort of the data.
NOTE Confidence: 0.939710803333333
00:01:37.240 --> 00:01:38.840 And the more interesting part of this, we
NOTE Confidence: 0.613679365
00:01:41.000 --> 00:01:41.960 have time for discussion
NOTE Confidence: 0.830800422
00:01:41.960 --> 00:01:42.728 and then Oh yeah,
NOTE Confidence: 0.830800422
00:01:42.728 --> 00:01:44.225 we want to leave as much time
NOTE Confidence: 0.830800422

00:01:44.225 --> 00:01:45.397 as possible for discussion.
NOTE Confidence: 0.830800422

00:01:45.400 --> 00:01:47.955 Let me just see how this works.
NOTE Confidence: 0.830800422

00:01:47.960 --> 00:01:50.640 OK, so disclosures,
NOTE Confidence: 0.830800422

00:01:50.640 --> 00:01:53.196 all of these above are Jerry's
NOTE Confidence: 0.830800422

00:01:53.200 --> 00:01:56.238 and I have this measly one liner
NOTE Confidence: 0.582320838571428

00:01:58.800 --> 00:02:01.880 skip. So psilocybin as as I said,
NOTE Confidence: 0.582320838571428

00:02:01.880 --> 00:02:04.640 I'm not going to go into detail about what
NOTE Confidence: 0.582320838571428

00:02:04.640 --> 00:02:06.600 psilocybin is and where it comes from.
NOTE Confidence: 0.582320838571428

00:02:06.600 --> 00:02:11.112 It's a natural occurring compound found
NOTE Confidence: 0.582320838571428

00:02:11.112 --> 00:02:14.800 in psilocybin mushrooms. It's found in.
NOTE Confidence: 0.691371978571429

00:02:17.120 --> 00:02:19.380 The particular extract comes
NOTE Confidence: 0.691371978571429

00:02:19.380 --> 00:02:21.075 from psilocybin kubensis,
NOTE Confidence: 0.691371978571429

00:02:21.080 --> 00:02:25.000 that the number of psychoactive
NOTE Confidence: 0.691371978571429

00:02:25.000 --> 00:02:25.744 species of mushrooms,
NOTE Confidence: 0.691371978571429

00:02:25.744 --> 00:02:27.232 at least in the United States,
NOTE Confidence: 0.691371978571429

00:02:27.240 --> 00:02:30.000 is in the range of 60.

NOTE Confidence: 0.691371978571429
00:02:30.000 --> 00:02:32.447 Briefly here I thought would
NOTE Confidence: 0.691371978571429
00:02:32.447 --> 00:02:34.781 be really useful to review the
NOTE Confidence: 0.691371978571429
00:02:34.781 --> 00:02:36.199 metabolism of psilocybin.
NOTE Confidence: 0.691371978571429
00:02:36.200 --> 00:02:39.035 As you can see psilocybin in its
NOTE Confidence: 0.691371978571429
00:02:39.040 --> 00:02:43.259 natural form it's not it's not active,
NOTE Confidence: 0.691371978571429
00:02:43.259 --> 00:02:44.798 it actually is.
NOTE Confidence: 0.691371978571429
00:02:44.800 --> 00:02:46.918 Once it's ingested that it is
NOTE Confidence: 0.29092836
00:02:51.640 --> 00:02:53.840 mute and turn off your your speaker
NOTE Confidence: 0.59558175
00:02:54.240 --> 00:02:55.359 it's so it
NOTE Confidence: 0.83852712125
00:02:57.440 --> 00:03:00.730 GI once it actually hits the stomach
NOTE Confidence: 0.83852712125
00:03:00.730 --> 00:03:03.315 it's dephosphorated as you can see
NOTE Confidence: 0.83852712125
00:03:03.315 --> 00:03:06.399 here and it becomes silos silosin or
NOTE Confidence: 0.83852712125
00:03:06.399 --> 00:03:09.069 silosin is the active ingredient and
NOTE Confidence: 0.83852712125
00:03:09.069 --> 00:03:12.536 the capsules that we use in our study
NOTE Confidence: 0.83852712125
00:03:12.536 --> 00:03:16.226 actually contain silosin rather than silos.
NOTE Confidence: 0.83852712125

00:03:16.226 --> 00:03:21.496 Cyben the much of the metabolism is actually

NOTE Confidence: 0.83852712125

00:03:21.496 --> 00:03:25.812 in the IT undergoes hepatic metabolism.

NOTE Confidence: 0.83852712125

00:03:25.812 --> 00:03:27.917 So it's glucurini,

NOTE Confidence: 0.83852712125

00:03:27.917 --> 00:03:30.702 Glucurini needed that it's broken

NOTE Confidence: 0.83852712125

00:03:30.702 --> 00:03:33.992 down into four hydroxy indole acetyl

NOTE Confidence: 0.83852712125

00:03:33.992 --> 00:03:37.240 hydrate and then the four HDP and four

NOTE Confidence: 0.3454221

00:03:39.400 --> 00:03:44.939 HAIAAR excreted urinary and it's

NOTE Confidence: 0.3454221

00:03:44.939 --> 00:03:48.120 the half life Typically this is out

NOTE Confidence: 0.3454221

00:03:48.120 --> 00:03:50.074 of your system within 24 hours.

NOTE Confidence: 0.3454221

00:03:50.074 --> 00:03:54.238 The half life is 4 to 6 hours.

NOTE Confidence: 0.3454221

00:03:54.240 --> 00:03:56.733 So just a little, I guess this is more

NOTE Confidence: 0.3454221

00:03:56.733 --> 00:03:59.108 what I said earlier that sell us you

NOTE Confidence: 0.3454221

00:03:59.108 --> 00:04:01.158 know the use of psilocybin in various

NOTE Confidence: 0.849401545

00:04:03.760 --> 00:04:06.735 cultures throughout the world goes

NOTE Confidence: 0.849401545

00:04:06.735 --> 00:04:10.440 back to a couple of 1000 years.

NOTE Confidence: 0.849401545

00:04:10.440 --> 00:04:13.666 But it's most prominent use that at

NOTE Confidence: 0.849401545

00:04:13.666 --> 00:04:15.805 least in the United States that's

NOTE Confidence: 0.849401545

00:04:15.805 --> 00:04:18.235 had an influence really comes from

NOTE Confidence: 0.849401545

00:04:18.235 --> 00:04:21.944 the indigenous use in Central

NOTE Confidence: 0.849401545

00:04:21.944 --> 00:04:24.932 America and Sierra Mastic region,

NOTE Confidence: 0.849401545

00:04:24.932 --> 00:04:26.708 particularly the Sierra Mastic

NOTE Confidence: 0.849401545

00:04:26.708 --> 00:04:29.240 region in Mexico. It came.

NOTE Confidence: 0.849401545

00:04:29.240 --> 00:04:33.584 It was first introduced to the States

NOTE Confidence: 0.849401545

00:04:33.584 --> 00:04:36.935 in 1950s and shortly thereafter

NOTE Confidence: 0.849401545

00:04:36.935 --> 00:04:40.360 studies with psilocybin were underway,

NOTE Confidence: 0.849401545

00:04:40.360 --> 00:04:43.765 particularly late 1950s and 1960s

NOTE Confidence: 0.849401545

00:04:43.765 --> 00:04:46.025 Psilocybin became a scheduled

NOTE Confidence: 0.849401545

00:04:46.025 --> 00:04:50.440 one compound in 1974.

NOTE Confidence: 0.849401545

00:04:50.440 --> 00:04:56.160 Here is just a quick Since recently with,

NOTE Confidence: 0.849401545

00:04:56.160 --> 00:04:59.280 you know, what's been dubbed as

NOTE Confidence: 0.849401545

00:04:59.280 --> 00:05:01.465 the Renaissance of psychedelics,

NOTE Confidence: 0.849401545

00:05:01.465 --> 00:05:04.746 there's been an interest in doing
NOTE Confidence: 0.849401545

00:05:04.746 --> 00:05:07.276 some more work with psilocybin.
NOTE Confidence: 0.849401545

00:05:07.280 --> 00:05:10.520 And this table summarizes the recent,
NOTE Confidence: 0.849401545

00:05:10.520 --> 00:05:12.752 most recent studies looking at the
NOTE Confidence: 0.849401545

00:05:12.752 --> 00:05:15.000 effect of psilocybin and depression.
NOTE Confidence: 0.849401545

00:05:15.000 --> 00:05:16.995 So there's been quite a few studies.
NOTE Confidence: 0.849401545

00:05:17.000 --> 00:05:19.542 This is actually not a comprehensive table.
NOTE Confidence: 0.849401545

00:05:19.542 --> 00:05:21.252 There's some studies that actually
NOTE Confidence: 0.849401545

00:05:21.252 --> 00:05:25.160 go back to 2016 that were done in
NOTE Confidence: 0.849401545

00:05:25.160 --> 00:05:28.240 patients with anxiety and depressive
NOTE Confidence: 0.849401545

00:05:28.240 --> 00:05:31.320 symptoms in palliative care setting,
NOTE Confidence: 0.849401545

00:05:31.320 --> 00:05:33.140 but those patients did not
NOTE Confidence: 0.849401545

00:05:33.140 --> 00:05:34.960 necessarily meet criteria for MD.
NOTE Confidence: 0.849401545

00:05:34.960 --> 00:05:35.794 These studies are,
NOTE Confidence: 0.849401545

00:05:35.794 --> 00:05:37.740 this one does not actually include the
NOTE Confidence: 0.849401545

00:05:37.793 --> 00:05:39.599 study that we're going to talk about,

NOTE Confidence: 0.849401545

00:05:39.600 --> 00:05:42.155 but these are the studies that have

NOTE Confidence: 0.849401545

00:05:42.160 --> 00:05:46.280 looked at psilocybin and MDD or and

NOTE Confidence: 0.849401545

00:05:46.280 --> 00:05:50.560 as well as PRD though this table

NOTE Confidence: 0.849401545

00:05:50.560 --> 00:05:53.160 doesn't differentiate between the two.

NOTE Confidence: 0.849401545

00:05:53.160 --> 00:05:56.040 And here in this I think we're going

NOTE Confidence: 0.849401545

00:05:56.040 --> 00:05:59.680 to spend most of the talk looking at

NOTE Confidence: 0.849401545

00:05:59.680 --> 00:06:02.830 the talking about the study at hand

NOTE Confidence: 0.849401545

00:06:02.830 --> 00:06:05.932 which is a single dose of 25 milligram

NOTE Confidence: 0.849401545

00:06:05.932 --> 00:06:09.000 of psilocybin in major depressive disorder.

NOTE Confidence: 0.849401545

00:06:09.000 --> 00:06:11.100 And as Chris mentioned earlier

NOTE Confidence: 0.849401545

00:06:11.100 --> 00:06:13.460 that we were Doctor Senecora and I,

NOTE Confidence: 0.849401545

00:06:13.460 --> 00:06:15.580 we were one of the sites here

NOTE Confidence: 0.849401545

00:06:15.580 --> 00:06:17.800 at Yale and ran the study.

NOTE Confidence: 0.849401545

00:06:17.800 --> 00:06:19.360 And with that any further ado,

NOTE Confidence: 0.849401545

00:06:19.360 --> 00:06:21.142 I'm going to hand this over

NOTE Confidence: 0.849401545

00:06:21.142 --> 00:06:22.033 to Doctor Senecora,
NOTE Confidence: 0.849401545

00:06:22.040 --> 00:06:24.240 He will review the details of the study.
NOTE Confidence: 0.819302645

00:06:24.760 --> 00:06:26.950 Great. Thanks Ben.
NOTE Confidence: 0.819302645

00:06:26.950 --> 00:06:30.940 So just to go through the
NOTE Confidence: 0.819302645

00:06:30.940 --> 00:06:32.640 the purpose of the study,
NOTE Confidence: 0.819302645

00:06:32.640 --> 00:06:35.041 as as Ben said there have been
NOTE Confidence: 0.819302645

00:06:35.041 --> 00:06:36.866 several studies prior to when
NOTE Confidence: 0.819302645

00:06:36.866 --> 00:06:38.318 this study was initiated.
NOTE Confidence: 0.819302645

00:06:38.320 --> 00:06:41.350 But many of the problems with
NOTE Confidence: 0.819302645

00:06:41.350 --> 00:06:42.750 those studies surrounded the
NOTE Confidence: 0.819302645

00:06:42.750 --> 00:06:44.800 small sample sizes that we use,
NOTE Confidence: 0.819302645

00:06:44.800 --> 00:06:46.584 the relatively short term
NOTE Confidence: 0.819302645

00:06:46.584 --> 00:06:49.760 follow up that was used,
NOTE Confidence: 0.819302645

00:06:49.760 --> 00:06:53.500 the the comparative that was used
NOTE Confidence: 0.819302645

00:06:53.500 --> 00:06:55.990 and and the frequent unblinding
NOTE Confidence: 0.819302645

00:06:55.990 --> 00:06:59.680 of the assessors or the raters.

NOTE Confidence: 0.819302645
00:06:59.680 --> 00:07:03.061 So this study aimed to try to
NOTE Confidence: 0.819302645
00:07:03.061 --> 00:07:05.543 address many of those concerns
NOTE Confidence: 0.819302645
00:07:05.543 --> 00:07:07.458 with the original studies when
NOTE Confidence: 0.819302645
00:07:07.458 --> 00:07:09.680 when it was originally initiated.
NOTE Confidence: 0.819302645
00:07:09.680 --> 00:07:12.952 So we'll take quite a bit of time
NOTE Confidence: 0.819302645
00:07:12.952 --> 00:07:15.086 just going through I'm there.
NOTE Confidence: 0.819302645
00:07:15.086 --> 00:07:16.501 I'm getting a message saying
NOTE Confidence: 0.819302645
00:07:16.501 --> 00:07:17.680 it's hard to hear.
NOTE Confidence: 0.819302645
00:07:17.680 --> 00:07:19.217 So I don't know if anything
NOTE Confidence: 0.819302645
00:07:19.217 --> 00:07:21.479 we can do about the volume,
NOTE Confidence: 0.819302645
00:07:21.480 --> 00:07:24.024 but so I wanted to just take
NOTE Confidence: 0.819302645
00:07:24.024 --> 00:07:25.000 some time going through this,
NOTE Confidence: 0.799821575238095
00:07:25.320 --> 00:07:26.888 sorry. We can turn off the room
NOTE Confidence: 0.799821575238095
00:07:26.888 --> 00:07:28.616 mics and you can turn on your
NOTE Confidence: 0.799821575238095
00:07:28.616 --> 00:07:30.320 computer mic and then it'll make it.
NOTE Confidence: 0.74916786

00:07:57.480 --> 00:07:59.639 OK. I don't know
NOTE Confidence: 0.6061673075

00:08:01.800 --> 00:08:02.280 if that's that that
NOTE Confidence: 0.580640235

00:08:12.450 --> 00:08:14.810 that you were getting. I don't know.
NOTE Confidence: 0.94073075

00:08:15.970 --> 00:08:17.894 We didn't get any complaints from others, so.
NOTE Confidence: 0.94073075

00:08:17.894 --> 00:08:20.486 OK. So I I'm assuming people can hear.
NOTE Confidence: 0.94073075

00:08:20.490 --> 00:08:22.290 So I'm going to move forward.
NOTE Confidence: 0.94073075

00:08:22.290 --> 00:08:24.050 It sounds good. OK.
NOTE Confidence: 0.94073075

00:08:24.050 --> 00:08:25.283 Thanks, Jess. Thank you. Good.
NOTE Confidence: 0.94073075

00:08:25.283 --> 00:08:26.770 So we're going to spend a little
NOTE Confidence: 0.94073075

00:08:26.770 --> 00:08:27.985 bit of time going through.
NOTE Confidence: 0.94073075

00:08:27.985 --> 00:08:30.860 This was actually the study
NOTE Confidence: 0.94073075

00:08:30.860 --> 00:08:34.960 design and this was a randomized
NOTE Confidence: 0.94073075

00:08:34.960 --> 00:08:38.293 2 group which was psilocybin 25
NOTE Confidence: 0.94073075

00:08:38.293 --> 00:08:40.558 milligrams or niacin 100 milligrams.
NOTE Confidence: 0.94073075

00:08:40.560 --> 00:08:42.880 Phase two clinical trial designed
NOTE Confidence: 0.94073075

00:08:42.880 --> 00:08:45.760 to evaluate both the efficacy of

NOTE Confidence: 0.94073075

00:08:45.760 --> 00:08:48.332 psilocybin as compared to niacin

NOTE Confidence: 0.94073075

00:08:48.332 --> 00:08:50.424 what it's administered with

NOTE Confidence: 0.94073075

00:08:50.424 --> 00:08:52.605 psychological support in patients

NOTE Confidence: 0.94073075

00:08:52.605 --> 00:08:54.717 with major depressive disorder.

NOTE Confidence: 0.94073075

00:08:54.720 --> 00:08:58.920 It was run at 11 US sites between

NOTE Confidence: 0.94073075

00:08:58.920 --> 00:09:02.106 December 2019 and June 2022 and we'll

NOTE Confidence: 0.94073075

00:09:02.106 --> 00:09:05.118 get a chance to talk a little bit later.

NOTE Confidence: 0.94073075

00:09:05.120 --> 00:09:07.495 That's kind of a interesting

NOTE Confidence: 0.94073075

00:09:07.495 --> 00:09:09.395 period of of history.

NOTE Confidence: 0.94073075

00:09:09.400 --> 00:09:11.668 So we'll get a chance to talk about how

NOTE Confidence: 0.94073075

00:09:11.668 --> 00:09:15.440 that could have had impact on the study.

NOTE Confidence: 0.94073075

00:09:15.440 --> 00:09:18.800 The inclusion criteria for the study,

NOTE Confidence: 0.94073075

00:09:18.800 --> 00:09:21.166 the main the key inclusion criteria was

NOTE Confidence: 0.94073075

00:09:21.166 --> 00:09:23.839 patients had to be medically healthy or

NOTE Confidence: 0.94073075

00:09:23.839 --> 00:09:26.233 participants had to be medically healthy.

NOTE Confidence: 0.94073075

00:09:26.240 --> 00:09:28.112 It was for adults between the
NOTE Confidence: 0.94073075

00:09:28.112 --> 00:09:30.000 ages of 21 and 65 with,
NOTE Confidence: 0.94073075

00:09:30.000 --> 00:09:31.215 as I said,
NOTE Confidence: 0.94073075

00:09:31.215 --> 00:09:34.322 a diagnosis of MDD with the current
NOTE Confidence: 0.94073075

00:09:34.322 --> 00:09:36.298 major depressive episode lasting
NOTE Confidence: 0.94073075

00:09:36.298 --> 00:09:38.958 of two months or or 60 days.
NOTE Confidence: 0.94073075

00:09:38.960 --> 00:09:43.000 They had to have a rating of over
NOTE Confidence: 0.94073075

00:09:43.000 --> 00:09:47.212 28 but equal to or above 28 on the
NOTE Confidence: 0.94073075

00:09:47.212 --> 00:09:49.700 Madras by the Central Rater and
NOTE Confidence: 0.94073075

00:09:49.700 --> 00:09:52.280 they couldn't have more than a 30%
NOTE Confidence: 0.94073075

00:09:52.280 --> 00:09:54.680 drop from the screening to baseline.
NOTE Confidence: 0.94073075

00:09:54.680 --> 00:09:56.080 So that's the inclusion criteria.
NOTE Confidence: 0.94073075

00:09:56.080 --> 00:09:57.400 Key exclusion criteria which are
NOTE Confidence: 0.94073075

00:09:57.400 --> 00:09:59.280 going to be important to remember is
NOTE Confidence: 0.899353404

00:10:01.960 --> 00:10:03.340 the patients had to be
NOTE Confidence: 0.899353404

00:10:03.340 --> 00:10:04.720 able to be medication free.

NOTE Confidence: 0.899353404

00:10:04.720 --> 00:10:06.640 So either not on a medication

NOTE Confidence: 0.899353404

00:10:06.640 --> 00:10:08.720 at the time of screening,

NOTE Confidence: 0.899353404

00:10:08.720 --> 00:10:11.921 or being able to tolerate a taper

NOTE Confidence: 0.899353404

00:10:11.921 --> 00:10:14.567 and a washout during the screening

NOTE Confidence: 0.899353404

00:10:14.567 --> 00:10:16.697 period and remain off antidepressant

NOTE Confidence: 0.899353404

00:10:16.697 --> 00:10:19.679 medications during that period of time.

NOTE Confidence: 0.899353404

00:10:19.680 --> 00:10:22.456 They also needed to be able to go

NOTE Confidence: 0.899353404

00:10:22.456 --> 00:10:24.958 without seeing an outpatient therapist.

NOTE Confidence: 0.899353404

00:10:24.960 --> 00:10:26.857 So if they have not had an

NOTE Confidence: 0.899353404

00:10:26.857 --> 00:10:27.946 outpatient therapist, that was OK,

NOTE Confidence: 0.899353404

00:10:27.946 --> 00:10:29.038 but if they were seeing one,

NOTE Confidence: 0.899353404

00:10:29.040 --> 00:10:31.238 they would have to be able to

NOTE Confidence: 0.899353404

00:10:31.240 --> 00:10:32.535 at least take a hiatus or a

NOTE Confidence: 0.899353404

00:10:32.535 --> 00:10:33.884 pause in their treatment with

NOTE Confidence: 0.899353404

00:10:33.884 --> 00:10:34.997 their outpatient therapist.

NOTE Confidence: 0.952799308461538

00:10:37.120 --> 00:10:39.451 The other part that was important to
NOTE Confidence: 0.952799308461538

00:10:39.451 --> 00:10:41.760 think about is the family history.
NOTE Confidence: 0.952799308461538

00:10:41.760 --> 00:10:43.884 So there was no personal history
NOTE Confidence: 0.952799308461538

00:10:43.884 --> 00:10:45.760 of either psychosis or mania,
NOTE Confidence: 0.952799308461538

00:10:45.760 --> 00:10:48.436 but also no first degree relative
NOTE Confidence: 0.952799308461538

00:10:48.440 --> 00:10:51.384 with psychosis or mania.
NOTE Confidence: 0.952799308461538

00:10:51.384 --> 00:10:55.750 And then the other major important two
NOTE Confidence: 0.952799308461538

00:10:55.750 --> 00:10:58.900 exclusion criterias was having psychedelic
NOTE Confidence: 0.952799308461538

00:10:58.900 --> 00:11:01.640 drug use in the past five years,
NOTE Confidence: 0.952799308461538

00:11:01.640 --> 00:11:04.125 or more than 10 lifetime
NOTE Confidence: 0.952799308461538

00:11:04.125 --> 00:11:06.113 uses of a psychedelic.
NOTE Confidence: 0.952799308461538

00:11:06.120 --> 00:11:07.320 And then finally the last one,
NOTE Confidence: 0.952799308461538

00:11:07.320 --> 00:11:09.590 was anybody with active suicidal
NOTE Confidence: 0.952799308461538

00:11:09.590 --> 00:11:12.695 ideation with intent or a plan within
NOTE Confidence: 0.952799308461538

00:11:12.695 --> 00:11:15.313 the past six months or or self,
NOTE Confidence: 0.952799308461538

00:11:15.320 --> 00:11:17.228 I'm sorry, or suicidal behavior in

NOTE Confidence: 0.952799308461538
00:11:17.228 --> 00:11:19.400 the past 12 months were excluded.
NOTE Confidence: 0.952799308461538
00:11:19.400 --> 00:11:22.352 So just to give you a general sense that
NOTE Confidence: 0.952799308461538
00:11:22.352 --> 00:11:25.317 this was a generally healthy population,
NOTE Confidence: 0.952799308461538
00:11:25.320 --> 00:11:26.680 really screened pretty carefully
NOTE Confidence: 0.952799308461538
00:11:26.680 --> 00:11:28.040 not to have psychosis,
NOTE Confidence: 0.952799308461538
00:11:28.040 --> 00:11:30.060 not even a family history
NOTE Confidence: 0.952799308461538
00:11:30.060 --> 00:11:31.676 of psychosis or mania.
NOTE Confidence: 0.952799308461538
00:11:31.680 --> 00:11:33.360 So well this is the population
NOTE Confidence: 0.952799308461538
00:11:33.360 --> 00:11:35.000 that it was geared towards.
NOTE Confidence: 0.952799308461538
00:11:35.000 --> 00:11:36.000 As you can see here,
NOTE Confidence: 0.952799308461538
00:11:36.000 --> 00:11:39.598 the original plan was to run 80
NOTE Confidence: 0.952799308461538
00:11:39.600 --> 00:11:42.680 participants that was actually updated
NOTE Confidence: 0.530287529
00:11:44.720 --> 00:11:47.924 see if I can. So it was updated to
NOTE Confidence: 0.530287529
00:11:47.924 --> 00:11:49.438 100 in the beginning of the study.
NOTE Confidence: 0.530287529
00:11:49.440 --> 00:11:52.824 The the there was a reanalysis and the and
NOTE Confidence: 0.530287529

00:11:52.824 --> 00:11:56.637 the idea to run 100 patients was was changed.
NOTE Confidence: 0.530287529

00:11:56.640 --> 00:11:58.830 So patients were screened and
NOTE Confidence: 0.530287529

00:11:58.830 --> 00:12:01.426 then after a screening period of
NOTE Confidence: 0.530287529

00:12:01.426 --> 00:12:03.400 lasting either 7 to 35 days,
NOTE Confidence: 0.530287529

00:12:03.400 --> 00:12:04.954 largely depending on if they needed
NOTE Confidence: 0.530287529

00:12:04.954 --> 00:12:06.760 to do a medication washout or not,
NOTE Confidence: 0.530287529

00:12:06.760 --> 00:12:10.724 they would then go into a baseline period.
NOTE Confidence: 0.530287529

00:12:10.724 --> 00:12:12.544 And during that baseline period
NOTE Confidence: 0.530287529

00:12:12.544 --> 00:12:14.000 they would be reassessed.
NOTE Confidence: 0.530287529

00:12:14.000 --> 00:12:14.880 And as I said,
NOTE Confidence: 0.530287529

00:12:14.880 --> 00:12:17.079 if they had more than a 30 point drop,
NOTE Confidence: 0.530287529

00:12:17.080 --> 00:12:18.880 a 30% drop in their ratings,
NOTE Confidence: 0.530287529

00:12:18.880 --> 00:12:20.160 that would be an exclusion,
NOTE Confidence: 0.530287529

00:12:20.160 --> 00:12:22.599 but otherwise they can move on and that would
NOTE Confidence: 0.530287529

00:12:22.599 --> 00:12:25.076 be their baseline ratings moving forward.
NOTE Confidence: 0.530287529

00:12:25.080 --> 00:12:27.240 And then before actual treatment

NOTE Confidence: 0.530287529

00:12:27.240 --> 00:12:29.400 or randomization was a preparatory

NOTE Confidence: 0.530287529

00:12:29.466 --> 00:12:31.466 session and these preparatory sessions

NOTE Confidence: 0.530287529

00:12:31.466 --> 00:12:34.238 were designed to last 6 to 8 hours,

NOTE Confidence: 0.530287529

00:12:34.240 --> 00:12:37.438 usually two sessions prior to the.

NOTE Confidence: 0.530287529

00:12:37.440 --> 00:12:40.040 Originally in this design it was meant to

NOTE Confidence: 0.530287529

00:12:40.040 --> 00:12:42.997 be the person with the two facilitators.

NOTE Confidence: 0.530287529

00:12:43.000 --> 00:12:43.472 The two,

NOTE Confidence: 0.530287529

00:12:43.472 --> 00:12:45.360 the two people in the room in these

NOTE Confidence: 0.530287529

00:12:45.423 --> 00:12:47.199 studies were called facilitators.

NOTE Confidence: 0.530287529

00:12:47.200 --> 00:12:49.927 But again if we go back and look at

NOTE Confidence: 0.530287529

00:12:49.927 --> 00:12:52.877 the timing starting in December 2019,

NOTE Confidence: 0.530287529

00:12:52.880 --> 00:12:53.840 the pandemic hit.

NOTE Confidence: 0.530287529

00:12:53.840 --> 00:12:55.760 So the prep sessions were altered

NOTE Confidence: 0.530287529

00:12:55.760 --> 00:12:57.719 where many of these prep sessions,

NOTE Confidence: 0.530287529

00:12:57.720 --> 00:13:02.400 at least one of them was done virtually.

NOTE Confidence: 0.530287529

00:13:02.400 --> 00:13:04.160 So that little bit different
NOTE Confidence: 0.530287529

00:13:04.160 --> 00:13:05.920 than the original study design.
NOTE Confidence: 0.530287529

00:13:05.920 --> 00:13:07.915 And then then there was day one
NOTE Confidence: 0.530287529

00:13:07.915 --> 00:13:10.102 and that was actually the dosing
NOTE Confidence: 0.530287529

00:13:10.102 --> 00:13:12.207 day and that's where patients
NOTE Confidence: 0.530287529

00:13:12.207 --> 00:13:14.093 or participants were randomized
NOTE Confidence: 0.530287529

00:13:14.093 --> 00:13:16.526 either receive the psilocybin 25
NOTE Confidence: 0.530287529

00:13:16.526 --> 00:13:18.956 milligrams or niacin 100 milligrams.
NOTE Confidence: 0.530287529

00:13:18.960 --> 00:13:21.284 And I'll just say briefly that the
NOTE Confidence: 0.530287529

00:13:21.284 --> 00:13:23.597 idea of using niacin was to have
NOTE Confidence: 0.530287529

00:13:23.600 --> 00:13:26.639 an attempt of at an active placebo,
NOTE Confidence: 0.530287529

00:13:26.639 --> 00:13:29.992 meaning that there may be some sensation
NOTE Confidence: 0.530287529

00:13:29.992 --> 00:13:32.998 from taking 100 milligrams of niacin,
NOTE Confidence: 0.530287529

00:13:33.000 --> 00:13:35.238 there's some flushing.
NOTE Confidence: 0.530287529

00:13:35.240 --> 00:13:37.100 It also offered us the opportunity
NOTE Confidence: 0.530287529

00:13:37.100 --> 00:13:39.024 to provide some equipoise by saying

NOTE Confidence: 0.530287529

00:13:39.024 --> 00:13:40.914 'cause there were there are some

NOTE Confidence: 0.530287529

00:13:40.914 --> 00:13:42.306 studies suggesting niacin may

NOTE Confidence: 0.530287529

00:13:42.306 --> 00:13:43.718 have an antidepressant effect.

NOTE Confidence: 0.530287529

00:13:43.720 --> 00:13:45.472 So you could legitimately tell patients

NOTE Confidence: 0.530287529

00:13:45.472 --> 00:13:47.755 that there is some evidence that they

NOTE Confidence: 0.530287529

00:13:47.755 --> 00:13:49.520 both may have antidepressant effects,

NOTE Confidence: 0.530287529

00:13:49.520 --> 00:13:53.872 but it was still was listed as a placebo,

NOTE Confidence: 0.530287529

00:13:53.872 --> 00:13:57.960 what would be called an active placebo.

NOTE Confidence: 0.530287529

00:13:57.960 --> 00:14:00.991 So dosing was provided on that day

NOTE Confidence: 0.530287529

00:14:00.991 --> 00:14:03.559 and dosing typically lasted 6 hours.

NOTE Confidence: 0.530287529

00:14:03.560 --> 00:14:05.906 And then patients were seen again

NOTE Confidence: 0.530287529

00:14:05.906 --> 00:14:08.260 the day after, which was day two.

NOTE Confidence: 0.530287529

00:14:08.260 --> 00:14:09.440 And on day two,

NOTE Confidence: 0.530287529

00:14:09.440 --> 00:14:12.038 they had their first integration session

NOTE Confidence: 0.530287529

00:14:12.040 --> 00:14:13.462 along with the ratings that were

NOTE Confidence: 0.530287529

00:14:13.462 --> 00:14:15.359 performed And in the integration session,
NOTE Confidence: 0.530287529

00:14:15.360 --> 00:14:16.640 typically lasting about an hour,
NOTE Confidence: 0.530287529

00:14:16.640 --> 00:14:21.320 then I think an hour to two hours depending,
NOTE Confidence: 0.530287529

00:14:21.320 --> 00:14:22.784 they were seen again.
NOTE Confidence: 0.530287529

00:14:22.784 --> 00:14:26.208 And then day five was a telephone interview.
NOTE Confidence: 0.530287529

00:14:26.208 --> 00:14:28.656 The day eight was one of
NOTE Confidence: 0.530287529

00:14:28.656 --> 00:14:29.880 the secondary measures,
NOTE Confidence: 0.530287529

00:14:29.880 --> 00:14:31.160 an outcome at day eight.
NOTE Confidence: 0.530287529

00:14:31.160 --> 00:14:32.786 And again they had their rating
NOTE Confidence: 0.530287529

00:14:32.786 --> 00:14:34.944 scales and you can see all the
NOTE Confidence: 0.530287529

00:14:34.944 --> 00:14:36.639 individual rating skills listed there.
NOTE Confidence: 0.530287529

00:14:36.640 --> 00:14:37.440 Day nine,
NOTE Confidence: 0.530287529

00:14:37.440 --> 00:14:39.440 they came back for another
NOTE Confidence: 0.530287529

00:14:39.440 --> 00:14:40.240 integration session.
NOTE Confidence: 0.530287529

00:14:40.240 --> 00:14:42.928 So this is again a meeting with
NOTE Confidence: 0.530287529

00:14:42.928 --> 00:14:44.504 the two facilitators lasting

NOTE Confidence: 0.530287529

00:14:44.504 --> 00:14:46.954 about an hour to to two hours.

NOTE Confidence: 0.530287529

00:14:46.960 --> 00:14:49.676 And then day 15 were more assessments.

NOTE Confidence: 0.530287529

00:14:49.680 --> 00:14:52.039 And then on day 16 was the

NOTE Confidence: 0.530287529

00:14:52.039 --> 00:14:53.050 3rd integration session

NOTE Confidence: 0.956447876

00:14:53.120 --> 00:14:55.480 meeting with the facilitators again

NOTE Confidence: 0.956447876

00:14:55.480 --> 00:14:56.926 and then there was another exploratory

NOTE Confidence: 0.956447876

00:14:56.926 --> 00:14:58.478 endpoint at the end of the month.

NOTE Confidence: 0.956447876

00:14:58.480 --> 00:15:00.120 And then the primary endpoint,

NOTE Confidence: 0.956447876

00:15:00.120 --> 00:15:02.584 the, the main goal of the study

NOTE Confidence: 0.956447876

00:15:02.584 --> 00:15:04.650 was the look at the response.

NOTE Confidence: 0.956447876

00:15:04.650 --> 00:15:08.101 There were the change in the Madras score on

NOTE Confidence: 0.956447876

00:15:08.101 --> 00:15:11.160 day 43 that was really look at six weeks out.

NOTE Confidence: 0.956447876

00:15:11.160 --> 00:15:13.876 That's the primary endpoint for the study.

NOTE Confidence: 0.956447876

00:15:13.880 --> 00:15:16.757 So let's just look a little bit.

NOTE Confidence: 0.956447876

00:15:16.760 --> 00:15:20.158 The idea of using this, as I said,

NOTE Confidence: 0.956447876

00:15:20.160 --> 00:15:21.580 along with psychological support goes
NOTE Confidence: 0.956447876

00:15:21.580 --> 00:15:24.000 back to the set and setting principles.
NOTE Confidence: 0.956447876

00:15:24.000 --> 00:15:25.176 And then I don't know if you want
NOTE Confidence: 0.956447876

00:15:25.176 --> 00:15:26.476 to talk a little bit about this or.
NOTE Confidence: 0.956447876

00:15:26.480 --> 00:15:29.440 Sure, yeah, yeah. The idea
NOTE Confidence: 0.853937831428571

00:15:29.440 --> 00:15:31.477 behind set and setting is something that
NOTE Confidence: 0.853937831428571

00:15:31.477 --> 00:15:33.680 was taken very seriously in this study.
NOTE Confidence: 0.853937831428571

00:15:33.680 --> 00:15:39.720 And set is defined as mindset sort of the
NOTE Confidence: 0.853937831428571

00:15:39.720 --> 00:15:44.006 your sort of baseline thinking process,
NOTE Confidence: 0.853937831428571

00:15:44.006 --> 00:15:46.805 your mood at the time of dosing and the
NOTE Confidence: 0.853937831428571

00:15:46.805 --> 00:15:48.811 setting typically referred to the environment
NOTE Confidence: 0.853937831428571

00:15:48.811 --> 00:15:52.477 where the dosing occurred in this.
NOTE Confidence: 0.853937831428571

00:15:52.480 --> 00:15:54.635 What's a typical though this
NOTE Confidence: 0.853937831428571

00:15:54.635 --> 00:15:56.790 becoming familiar to this group
NOTE Confidence: 0.853937831428571

00:15:56.865 --> 00:15:59.138 is that these dosing sessions are
NOTE Confidence: 0.853937831428571

00:15:59.138 --> 00:16:01.610 administered in a slightly unusual

NOTE Confidence: 0.853937831428571
00:16:01.610 --> 00:16:04.435 environmental setting where you know,
NOTE Confidence: 0.853937831428571
00:16:04.440 --> 00:16:06.160 as you can see in the picture here,
NOTE Confidence: 0.853937831428571
00:16:06.160 --> 00:16:07.600 there is actually a couch.
NOTE Confidence: 0.815448939230769
00:16:09.880 --> 00:16:12.197 The room is set up to actually
NOTE Confidence: 0.815448939230769
00:16:12.197 --> 00:16:16.680 looks to look like a ordinary room.
NOTE Confidence: 0.815448939230769
00:16:16.680 --> 00:16:18.390 It's typically a little bit
NOTE Confidence: 0.815448939230769
00:16:18.390 --> 00:16:19.633 more aesthetically appealing
NOTE Confidence: 0.815448939230769
00:16:19.633 --> 00:16:22.398 than a typical Med room.
NOTE Confidence: 0.815448939230769
00:16:22.400 --> 00:16:25.208 The other idea behind the setting
NOTE Confidence: 0.815448939230769
00:16:25.208 --> 00:16:27.751 that it would facilitate A
NOTE Confidence: 0.815448939230769
00:16:27.751 --> 00:16:30.103 therapeutic alliance and between
NOTE Confidence: 0.815448939230769
00:16:30.103 --> 00:16:32.132 the facilitators and participants.
NOTE Confidence: 0.815448939230769
00:16:32.132 --> 00:16:35.114 So you know this is what was
NOTE Confidence: 0.815448939230769
00:16:35.114 --> 00:16:38.254 taken into account during both the
NOTE Confidence: 0.815448939230769
00:16:38.254 --> 00:16:40.879 preparation as well as integration.
NOTE Confidence: 0.815448939230769

00:16:40.880 --> 00:16:44.273 This idea of using Satis setting
NOTE Confidence: 0.815448939230769

00:16:44.273 --> 00:16:46.870 as a foundation for where a lot
NOTE Confidence: 0.815448939230769

00:16:46.944 --> 00:16:49.440 of the conversations took place.
NOTE Confidence: 0.815448939230769

00:16:49.440 --> 00:16:51.000 So treatment sessions,
NOTE Confidence: 0.754274631666667

00:16:51.760 --> 00:16:55.360 so, so as Ben described this,
NOTE Confidence: 0.754274631666667

00:16:55.360 --> 00:16:59.568 the idea of using the the integration,
NOTE Confidence: 0.754274631666667

00:16:59.568 --> 00:17:02.816 I'm sorry, the preparatory sessions to help
NOTE Confidence: 0.754274631666667

00:17:02.816 --> 00:17:05.857 work on the set and the actual physical
NOTE Confidence: 0.754274631666667

00:17:05.857 --> 00:17:08.720 setting was was very tightly controlled.
NOTE Confidence: 0.754274631666667

00:17:08.720 --> 00:17:10.685 During dosing sessions participants were
NOTE Confidence: 0.754274631666667

00:17:10.685 --> 00:17:13.689 encouraged to wear eye shades and to listen
NOTE Confidence: 0.754274631666667

00:17:13.689 --> 00:17:15.639 to a curated playlist with headphones.
NOTE Confidence: 0.754274631666667

00:17:15.640 --> 00:17:17.796 So this is again very tightly controlled.
NOTE Confidence: 0.754274631666667

00:17:17.800 --> 00:17:20.611 The setting the leaf facilitators
NOTE Confidence: 0.754274631666667

00:17:20.611 --> 00:17:21.959 were all doctoral level,
NOTE Confidence: 0.754274631666667

00:17:21.960 --> 00:17:26.760 either psychologists or physicians with

NOTE Confidence: 0.754274631666667
00:17:26.760 --> 00:17:28.612 treatment with depression treatment
NOTE Confidence: 0.754274631666667
00:17:28.612 --> 00:17:30.927 experience and the Co facilitators
NOTE Confidence: 0.754274631666667
00:17:30.927 --> 00:17:32.980 held a minimum of a bachelor's degree
NOTE Confidence: 0.754274631666667
00:17:32.980 --> 00:17:34.960 in a mental health related field.
NOTE Confidence: 0.754274631666667
00:17:34.960 --> 00:17:36.175 All facilitators completed
NOTE Confidence: 0.754274631666667
00:17:36.175 --> 00:17:37.795 a study specific training.
NOTE Confidence: 0.754274631666667
00:17:37.800 --> 00:17:40.222 So this is clearly a case where
NOTE Confidence: 0.754274631666667
00:17:40.222 --> 00:17:42.464 there was great effort taken to
NOTE Confidence: 0.754274631666667
00:17:42.464 --> 00:17:44.394 make sure that the facilitating
NOTE Confidence: 0.797341222
00:17:47.320 --> 00:17:49.560 study members were were providing
NOTE Confidence: 0.797341222
00:17:49.560 --> 00:17:52.413 as much of the uniformed experience
NOTE Confidence: 0.797341222
00:17:52.413 --> 00:17:55.478 as possible in the study.
NOTE Confidence: 0.797341222
00:17:55.480 --> 00:17:57.088 Here if we go through looking
NOTE Confidence: 0.797341222
00:17:57.088 --> 00:17:58.160 at the consort diagram.
NOTE Confidence: 0.797341222
00:17:58.160 --> 00:18:00.768 So this is one of the first interesting
NOTE Confidence: 0.797341222

00:18:00.768 --> 00:18:02.958 things to see is that there were
NOTE Confidence: 0.97244084

00:18:05.200 --> 00:18:09.680 15129 people assessed by phone
NOTE Confidence: 0.97244084

00:18:09.680 --> 00:18:12.714 screen with 1425 excluded.
NOTE Confidence: 0.97244084

00:18:12.714 --> 00:18:14.513 So this is one of these studies
NOTE Confidence: 0.97244084

00:18:14.513 --> 00:18:16.002 where there was a lot of interest
NOTE Confidence: 0.97244084

00:18:16.002 --> 00:18:17.754 and a lot of people that wanted to
NOTE Confidence: 0.97244084

00:18:17.754 --> 00:18:19.516 take part in a study and we can
NOTE Confidence: 0.97244084

00:18:19.516 --> 00:18:20.902 talk about the pros and cons of
NOTE Confidence: 0.97244084

00:18:20.902 --> 00:18:22.302 that that that that could either
NOTE Confidence: 0.97244084

00:18:22.302 --> 00:18:23.979 work in your favour or that could
NOTE Confidence: 0.97244084

00:18:23.979 --> 00:18:25.395 work against you in some ways.
NOTE Confidence: 0.97244084

00:18:25.400 --> 00:18:26.412 But for this study,
NOTE Confidence: 0.97244084

00:18:26.412 --> 00:18:28.305 you can see that's a pretty large
NOTE Confidence: 0.97244084

00:18:28.305 --> 00:18:29.875 number that was whittled down
NOTE Confidence: 0.860973454285714

00:18:32.640 --> 00:18:34.880 to to a number.
NOTE Confidence: 0.860973454285714

00:18:34.880 --> 00:18:37.610 As you said, 11182 were excluded

NOTE Confidence: 0.860973454285714
00:18:37.610 --> 00:18:39.360 just on the telephone screen.
NOTE Confidence: 0.860973454285714
00:18:39.360 --> 00:18:41.600 So just on those initial telephone screens,
NOTE Confidence: 0.860973454285714
00:18:41.600 --> 00:18:44.140 it was either the patient didn't meet
NOTE Confidence: 0.860973454285714
00:18:44.140 --> 00:18:46.310 the depression or some of the patients
NOTE Confidence: 0.860973454285714
00:18:46.310 --> 00:18:48.835 would call and didn't have depression at all.
NOTE Confidence: 0.860973454285714
00:18:48.840 --> 00:18:52.000 They just wanted to try study,
NOTE Confidence: 0.860973454285714
00:18:52.000 --> 00:18:53.752 but others had other clear reasons
NOTE Confidence: 0.860973454285714
00:18:53.752 --> 00:18:55.552 where they may have had psychosis
NOTE Confidence: 0.860973454285714
00:18:55.552 --> 00:18:57.274 or a family member had psychosis
NOTE Confidence: 0.860973454285714
00:18:57.274 --> 00:18:59.039 or for a variety of reasons.
NOTE Confidence: 0.860973454285714
00:18:59.040 --> 00:19:01.155 But then it got down to the point where
NOTE Confidence: 0.860973454285714
00:19:01.160 --> 00:19:02.996 240 were actually screened in person.
NOTE Confidence: 0.860973454285714
00:19:03.000 --> 00:19:04.875 And then there were additional
NOTE Confidence: 0.860973454285714
00:19:04.875 --> 00:19:06.784 exclusions down to 104
NOTE Confidence: 0.860973454285714
00:19:06.784 --> 00:19:09.920 participants that were randomized,
NOTE Confidence: 0.860973454285714

00:19:09.920 --> 00:19:13.118 51 randomized to receive the psilocybin
NOTE Confidence: 0.860973454285714

00:19:13.118 --> 00:19:16.680 and 53 randomized to receive niacin.
NOTE Confidence: 0.860973454285714

00:19:16.680 --> 00:19:17.828 In fact, if we,
NOTE Confidence: 0.860973454285714

00:19:17.828 --> 00:19:19.263 if you read the paper
NOTE Confidence: 0.860973454285714

00:19:19.263 --> 00:19:20.920 real quick real closely,
NOTE Confidence: 0.860973454285714

00:19:20.920 --> 00:19:23.237 it actually ended up that 50 people
NOTE Confidence: 0.860973454285714

00:19:23.240 --> 00:19:26.252 actually received the psilocybin and 54
NOTE Confidence: 0.860973454285714

00:19:26.252 --> 00:19:29.840 received niacin because one patient was
NOTE Confidence: 0.860973454285714

00:19:29.840 --> 00:19:33.518 by mistake given the wrong treatment.
NOTE Confidence: 0.860973454285714

00:19:33.520 --> 00:19:37.714 So, so really ended up 50 to to 54
NOTE Confidence: 0.860973454285714

00:19:37.720 --> 00:19:39.736 which is actually not that uncommon
NOTE Confidence: 0.860973454285714

00:19:39.736 --> 00:19:41.794 in in these studies that there's
NOTE Confidence: 0.860973454285714

00:19:41.794 --> 00:19:45.360 one mistake in there somewhere.
NOTE Confidence: 0.860973454285714

00:19:45.360 --> 00:19:47.754 And then you can see the number
NOTE Confidence: 0.860973454285714

00:19:47.754 --> 00:19:49.832 that completed at the day 8
NOTE Confidence: 0.860973454285714

00:19:49.832 --> 00:19:52.760 assessment you had 51 of the of the

NOTE Confidence: 0.860973454285714
00:19:52.760 --> 00:19:55.080 51 patients completing the day at
NOTE Confidence: 0.860973454285714
00:19:55.080 --> 00:19:57.525 and actually 50 of the psilocybin.
NOTE Confidence: 0.860973454285714
00:19:57.525 --> 00:20:00.255 Of the 51 psilocybin patients completing
NOTE Confidence: 0.860973454285714
00:20:00.255 --> 00:20:03.180 the six week evaluation where for the
NOTE Confidence: 0.860973454285714
00:20:03.180 --> 00:20:06.292 niacin you had 48 of the 53 make it
NOTE Confidence: 0.860973454285714
00:20:06.292 --> 00:20:09.360 to day eight and only 42 of the 53
NOTE Confidence: 0.860973454285714
00:20:09.360 --> 00:20:12.118 make it to the six week assessment.
NOTE Confidence: 0.860973454285714
00:20:12.120 --> 00:20:13.896 So you had nine people that
NOTE Confidence: 0.860973454285714
00:20:13.896 --> 00:20:15.360 did not make it on?
NOTE Confidence: 0.860973454285714
00:20:15.360 --> 00:20:16.240 And then on the right,
NOTE Confidence: 0.860973454285714
00:20:16.240 --> 00:20:19.180 you can actually see by the
NOTE Confidence: 0.860973454285714
00:20:19.180 --> 00:20:21.270 individual sites how many people
NOTE Confidence: 0.860973454285714
00:20:21.270 --> 00:20:23.120 were recruited at what percent.
NOTE Confidence: 0.860973454285714
00:20:23.120 --> 00:20:25.064 This is the number of participants
NOTE Confidence: 0.860973454285714
00:20:25.064 --> 00:20:27.303 that each site had and the percent
NOTE Confidence: 0.860973454285714

00:20:27.303 --> 00:20:29.137 of the total that they shared.
NOTE Confidence: 0.860973454285714

00:20:29.137 --> 00:20:32.153 You can see although there was 11 sites,
NOTE Confidence: 0.860973454285714

00:20:32.160 --> 00:20:34.384 really the top five or six accounted for
NOTE Confidence: 0.860973454285714

00:20:34.384 --> 00:20:36.556 the large majority of the patients run
NOTE Confidence: 0.767647362

00:20:39.480 --> 00:20:42.360 Kathleen. At the patient characteristics
NOTE Confidence: 0.767647362

00:20:42.360 --> 00:20:44.467 we can see that there was a
NOTE Confidence: 0.767647362

00:20:44.467 --> 00:20:46.283 pretty much an equal distribution
NOTE Confidence: 0.767647362

00:20:46.283 --> 00:20:48.935 of men and women in this study.
NOTE Confidence: 0.767647362

00:20:48.935 --> 00:20:52.360 The average age was about 41.
NOTE Confidence: 0.767647362

00:20:52.360 --> 00:20:54.640 Ethnicity was something
NOTE Confidence: 0.767647362

00:20:54.640 --> 00:20:57.680 we'll talk about later.
NOTE Confidence: 0.767647362

00:20:57.680 --> 00:21:00.400 There was about 94%
NOTE Confidence: 0.80088173125

00:21:04.000 --> 00:21:08.800 white patients participants in the study
NOTE Confidence: 0.80088173125

00:21:08.800 --> 00:21:14.910 and only less than 3% or about 3% black or
NOTE Confidence: 0.80088173125

00:21:14.910 --> 00:21:18.240 African American participating in this study.
NOTE Confidence: 0.80088173125

00:21:18.240 --> 00:21:23.022 The other things of real interest is only

NOTE Confidence: 0.80088173125

00:21:23.022 --> 00:21:27.234 about 12% had treatment resistant depression.

NOTE Confidence: 0.80088173125

00:21:27.240 --> 00:21:29.656 So this was primarily and the large majority

NOTE Confidence: 0.80088173125

00:21:29.656 --> 00:21:32.543 of these patients were not meeting criteria

NOTE Confidence: 0.80088173125

00:21:32.543 --> 00:21:34.319 for treatment resistant depression,

NOTE Confidence: 0.80088173125

00:21:34.320 --> 00:21:36.427 although there was no cap on the

NOTE Confidence: 0.80088173125

00:21:36.427 --> 00:21:37.920 number of treatment failures.

NOTE Confidence: 0.80088173125

00:21:37.920 --> 00:21:40.656 So, so there were people that could have

NOTE Confidence: 0.80088173125

00:21:40.656 --> 00:21:42.520 had any number of treatment failures.

NOTE Confidence: 0.80088173125

00:21:42.520 --> 00:21:44.280 It was really about 12,

NOTE Confidence: 0.80088173125

00:21:44.280 --> 00:21:46.499 1/2% that would have met the criteria

NOTE Confidence: 0.80088173125

00:21:46.499 --> 00:21:48.080 for treatment resistant depression.

NOTE Confidence: 0.80088173125

00:21:48.080 --> 00:21:50.026 And you can see the number that

NOTE Confidence: 0.80088173125

00:21:50.026 --> 00:21:51.920 actually went through a medication taper

NOTE Confidence: 0.80088173125

00:21:51.920 --> 00:21:53.703 both for psilocybin and for niacin.

NOTE Confidence: 0.80088173125

00:21:53.703 --> 00:21:56.240 But in general it was about one out of

NOTE Confidence: 0.80088173125

00:21:56.240 --> 00:21:58.196 five people required a medication taper,
NOTE Confidence: 0.80088173125

00:21:58.200 --> 00:22:03.000 meaning that about 80% of the patients or
NOTE Confidence: 0.80088173125

00:22:03.000 --> 00:22:05.160 75% of the patients didn't require a taper.
NOTE Confidence: 0.80088173125

00:22:05.160 --> 00:22:07.578 They were not on an antidepressant
NOTE Confidence: 0.80088173125

00:22:07.578 --> 00:22:10.839 at the time of entering the study.
NOTE Confidence: 0.80088173125

00:22:10.840 --> 00:22:11.524 Baseline mattress,
NOTE Confidence: 0.80088173125

00:22:11.524 --> 00:22:14.809 as I said the cut off was was actually
NOTE Confidence: 0.80088173125

00:22:14.809 --> 00:22:17.080 pretty high for the study of 28.
NOTE Confidence: 0.80088173125

00:22:17.080 --> 00:22:19.866 So the the baseline mattress of 35
NOTE Confidence: 0.80088173125

00:22:19.866 --> 00:22:25.300 is is a pretty high average level
NOTE Confidence: 0.80088173125

00:22:25.300 --> 00:22:27.200 of severity for a study.
NOTE Confidence: 0.80088173125

00:22:27.200 --> 00:22:28.936 The SDS scores of,
NOTE Confidence: 0.80088173125

00:22:28.936 --> 00:22:32.061 you know somewhere around seven was actually
NOTE Confidence: 0.80088173125

00:22:32.061 --> 00:22:34.968 a little bit lower then you would expect.
NOTE Confidence: 0.80088173125

00:22:34.968 --> 00:22:36.720 So a lower level that's the
NOTE Confidence: 0.80088173125

00:22:36.779 --> 00:22:38.240 Sheehan disability scale,

NOTE Confidence: 0.80088173125

00:22:38.240 --> 00:22:40.832 a little bit lower level of disability than

NOTE Confidence: 0.80088173125

00:22:40.832 --> 00:22:43.639 you see in most other depression studies.

NOTE Confidence: 0.80088173125

00:22:43.640 --> 00:22:46.336 And then you can see the average length of

NOTE Confidence: 0.80088173125

00:22:46.336 --> 00:22:49.706 depression and number of previous episodes.

NOTE Confidence: 0.80088173125

00:22:49.706 --> 00:22:50.559 Jerry,

NOTE Confidence: 0.84663695

00:22:50.560 --> 00:22:52.296 what do you make of that observation

NOTE Confidence: 0.84663695

00:22:52.296 --> 00:22:54.137 that the the Madras is on the high

NOTE Confidence: 0.84663695

00:22:54.137 --> 00:22:56.040 side but the SDS is on the low side?

NOTE Confidence: 0.84663695

00:22:56.040 --> 00:22:57.951 Is that because so many other possible

NOTE Confidence: 0.84663695

00:22:57.951 --> 00:22:59.598 sources of disability are screened out?

NOTE Confidence: 0.6881867633333333

00:23:00.000 --> 00:23:02.408 I I think that's a a large possibility.

NOTE Confidence: 0.6881867633333333

00:23:02.408 --> 00:23:04.840 The way the way the Sheen is collected

NOTE Confidence: 0.6881867633333333

00:23:04.900 --> 00:23:06.285 may give you some artificial

NOTE Confidence: 0.6881867633333333

00:23:06.285 --> 00:23:08.320 ratings with that too a little bit.

NOTE Confidence: 0.6881867633333333

00:23:08.320 --> 00:23:09.792 But I also think and we'll talk about

NOTE Confidence: 0.6881867633333333

00:23:09.792 --> 00:23:11.519 when we talk about the generalizability,
NOTE Confidence: 0.6881867633333333

00:23:11.520 --> 00:23:13.116 this is kind of a unique population,
NOTE Confidence: 0.6881867633333333

00:23:13.120 --> 00:23:14.203 very highly educated,
NOTE Confidence: 0.6881867633333333

00:23:14.203 --> 00:23:18.176 more so than you would see in a lot of
NOTE Confidence: 0.6881867633333333

00:23:18.176 --> 00:23:20.000 other typical antidepressant trials.
NOTE Confidence: 0.6881867633333333

00:23:20.000 --> 00:23:21.300 And that I think probably
NOTE Confidence: 0.6881867633333333

00:23:21.300 --> 00:23:22.800 attributes to some of that. There
NOTE Confidence: 0.6982936811111111

00:23:22.800 --> 00:23:25.110 is buffer. There are various
NOTE Confidence: 0.6982936811111111

00:23:25.110 --> 00:23:26.496 characteristics that buffer
NOTE Confidence: 0.6982936811111111

00:23:26.496 --> 00:23:27.680 disability in this population.
NOTE Confidence: 0.709693466363636

00:23:28.360 --> 00:23:30.866 Yeah, they have a lot of other
NOTE Confidence: 0.709693466363636

00:23:30.866 --> 00:23:32.982 buffers to to prevent it.
NOTE Confidence: 0.709693466363636

00:23:32.982 --> 00:23:35.274 All right. So here's the data.
NOTE Confidence: 0.709693466363636

00:23:35.280 --> 00:23:37.086 I mean, there's the data is the
NOTE Confidence: 0.709693466363636

00:23:37.086 --> 00:23:38.799 real easy thing to present here.
NOTE Confidence: 0.709693466363636

00:23:38.800 --> 00:23:41.560 This is pretty, pretty clear.

NOTE Confidence: 0.709693466363636
00:23:41.560 --> 00:23:45.124 So that, yeah, that there was no doubt in
NOTE Confidence: 0.709693466363636
00:23:45.124 --> 00:23:48.673 the significance there with AP of 001.001.
NOTE Confidence: 0.709693466363636
00:23:48.673 --> 00:23:52.066 The mean difference was the 12.3 points
NOTE Confidence: 0.709693466363636
00:23:52.066 --> 00:23:54.957 on a mattress which is quite large,
NOTE Confidence: 0.709693466363636
00:23:54.960 --> 00:23:58.360 you know very large.
NOTE Confidence: 0.709693466363636
00:23:58.360 --> 00:24:00.369 So this is really standing out in
NOTE Confidence: 0.709693466363636
00:24:00.369 --> 00:24:02.358 in many ways as a very strong.
NOTE Confidence: 0.709693466363636
00:24:02.360 --> 00:24:03.920 Remember the the primary end
NOTE Confidence: 0.709693466363636
00:24:03.920 --> 00:24:05.480 point here is day 43,
NOTE Confidence: 0.709693466363636
00:24:05.480 --> 00:24:07.436 so it's this last time point.
NOTE Confidence: 0.709693466363636
00:24:07.440 --> 00:24:09.360 If you look look at it other ways,
NOTE Confidence: 0.709693466363636
00:24:09.360 --> 00:24:11.448 this is actually my new favorite
NOTE Confidence: 0.709693466363636
00:24:11.448 --> 00:24:12.840 way to present data.
NOTE Confidence: 0.709693466363636
00:24:12.840 --> 00:24:14.472 It's called the waterfall
NOTE Confidence: 0.709693466363636
00:24:14.472 --> 00:24:16.026 plot where you actually,
NOTE Confidence: 0.709693466363636

00:24:16.026 --> 00:24:18.250 you can see the the mattress
NOTE Confidence: 0.709693466363636

00:24:18.250 --> 00:24:19.975 score that the patients start
NOTE Confidence: 0.709693466363636

00:24:19.975 --> 00:24:23.745 with or the participants and
NOTE Confidence: 0.709693466363636

00:24:23.745 --> 00:24:25.880 then how far they drop or rise.
NOTE Confidence: 0.709693466363636

00:24:25.880 --> 00:24:28.120 For the few people that actually got worse,
NOTE Confidence: 0.709693466363636

00:24:28.120 --> 00:24:30.880 you can see here that there's an increase.
NOTE Confidence: 0.709693466363636

00:24:30.880 --> 00:24:32.800 So you can see the group,
NOTE Confidence: 0.709693466363636

00:24:32.800 --> 00:24:35.764 the orange represents the psilocybin and
NOTE Confidence: 0.709693466363636

00:24:35.764 --> 00:24:38.640 the blue representing the the niacin.
NOTE Confidence: 0.709693466363636

00:24:38.640 --> 00:24:40.474 You can see that the large majority
NOTE Confidence: 0.709693466363636

00:24:40.474 --> 00:24:42.072 of people getting psilocybin
NOTE Confidence: 0.709693466363636

00:24:42.072 --> 00:24:43.680 had significant drops,
NOTE Confidence: 0.709693466363636

00:24:43.680 --> 00:24:45.384 very few that did a little
NOTE Confidence: 0.709693466363636

00:24:45.384 --> 00:24:47.119 bit worse and in fact no,
NOTE Confidence: 0.709693466363636

00:24:47.120 --> 00:24:48.920 no more doing worse than
NOTE Confidence: 0.709693466363636

00:24:48.920 --> 00:24:50.457 the people getting niacin.

NOTE Confidence: 0.709693466363636

00:24:50.457 --> 00:24:53.656 So it's really nice way of looking

NOTE Confidence: 0.709693466363636

00:24:53.656 --> 00:24:56.118 at individual patient data here.

NOTE Confidence: 0.709693466363636

00:24:56.120 --> 00:24:58.164 And then you can see the change

NOTE Confidence: 0.709693466363636

00:24:58.164 --> 00:25:00.319 in Madras from baseline to day 43,

NOTE Confidence: 0.709693466363636

00:25:00.320 --> 00:25:02.635 which was the primary outcome

NOTE Confidence: 0.709693466363636

00:25:02.635 --> 00:25:04.487 measure very clearly hitting

NOTE Confidence: 0.709693466363636

00:25:04.487 --> 00:25:05.880 statistical significance.

NOTE Confidence: 0.709693466363636

00:25:05.880 --> 00:25:08.323 So this data for the primary outcome

NOTE Confidence: 0.709693466363636

00:25:08.323 --> 00:25:11.119 I don't think could be more promising.

NOTE Confidence: 0.709693466363636

00:25:11.120 --> 00:25:13.352 I don't think you could have

NOTE Confidence: 0.709693466363636

00:25:13.352 --> 00:25:14.840 asked for much more.

NOTE Confidence: 0.709693466363636

00:25:14.840 --> 00:25:16.544 Also for secondary measures,

NOTE Confidence: 0.709693466363636

00:25:16.544 --> 00:25:18.674 the data were very strong.

NOTE Confidence: 0.709693466363636

00:25:18.680 --> 00:25:20.498 So here's looking at the Madras

NOTE Confidence: 0.709693466363636

00:25:20.498 --> 00:25:22.000 scores throughout the different days.

NOTE Confidence: 0.709693466363636

00:25:22.000 --> 00:25:23.540 You can see pretty much
NOTE Confidence: 0.709693466363636

00:25:23.540 --> 00:25:25.080 significance all the way through.
NOTE Confidence: 0.709693466363636

00:25:25.080 --> 00:25:26.052 Even the the Sheehan,
NOTE Confidence: 0.709693466363636

00:25:26.052 --> 00:25:27.861 even though I said the the the
NOTE Confidence: 0.709693466363636

00:25:27.861 --> 00:25:29.037 baseline levels of disability
NOTE Confidence: 0.709693466363636

00:25:29.037 --> 00:25:31.098 weren't maybe as high as you may
NOTE Confidence: 0.709693466363636

00:25:31.098 --> 00:25:32.438 expect in some other studies,
NOTE Confidence: 0.709693466363636

00:25:32.440 --> 00:25:35.824 they still showed high levels of change
NOTE Confidence: 0.709693466363636

00:25:35.824 --> 00:25:38.960 significance compared to niacin.
NOTE Confidence: 0.709693466363636

00:25:38.960 --> 00:25:42.656 And if you looked at the sustained
NOTE Confidence: 0.709693466363636

00:25:42.656 --> 00:25:45.240 depressive response at the six week point,
NOTE Confidence: 0.709693466363636

00:25:45.240 --> 00:25:47.480 highly statistically significant remission
NOTE Confidence: 0.709693466363636

00:25:47.480 --> 00:25:50.280 didn't quite hit statistical significance,
NOTE Confidence: 0.709693466363636

00:25:50.280 --> 00:25:51.528 but pretty darn close.
NOTE Confidence: 0.709693466363636

00:25:51.528 --> 00:25:53.827 And you can actually see here sometimes
NOTE Confidence: 0.709693466363636

00:25:53.827 --> 00:25:56.396 visually it's just a little bit better.

NOTE Confidence: 0.709693466363636

00:25:56.400 --> 00:25:58.234 This is looking at the response rates.

NOTE Confidence: 0.709693466363636

00:25:58.240 --> 00:26:00.814 So we had 20 of 48 in psilocybin 5

NOTE Confidence: 0.709693466363636

00:26:00.814 --> 00:26:03.439 and 44 in Niacin meeting response,

NOTE Confidence: 0.709693466363636

00:26:03.440 --> 00:26:05.660 which was a 50% improvement on

NOTE Confidence: 0.709693466363636

00:26:05.660 --> 00:26:08.240 the Madras and an odds ratio

NOTE Confidence: 0.709693466363636

00:26:08.240 --> 00:26:10.502 of response of about 5.6.

NOTE Confidence: 0.709693466363636

00:26:10.502 --> 00:26:13.034 If you look at sustained remission,

NOTE Confidence: 0.709693466363636

00:26:13.040 --> 00:26:15.504 there was 12 of 48 meaning that

NOTE Confidence: 0.709693466363636

00:26:15.504 --> 00:26:18.990 they had a Madras less than 10 for

NOTE Confidence: 0.709693466363636

00:26:18.990 --> 00:26:22.346 psilocybin and only about I think it

NOTE Confidence: 0.709693466363636

00:26:22.346 --> 00:26:25.675 was about 11 percent or so for for

NOTE Confidence: 0.709693466363636

00:26:25.675 --> 00:26:29.155 Niacin with an odd ratio of over three.

NOTE Confidence: 0.709693466363636

00:26:29.160 --> 00:26:31.561 So again very strong data and remember

NOTE Confidence: 0.709693466363636

00:26:31.561 --> 00:26:33.834 this is different than some of the

NOTE Confidence: 0.709693466363636

00:26:33.834 --> 00:26:36.074 other studies that you may have seen

NOTE Confidence: 0.709693466363636

00:26:36.074 --> 00:26:38.119 where there's been repeated dosing.
NOTE Confidence: 0.709693466363636

00:26:38.120 --> 00:26:40.070 This is just that one single
NOTE Confidence: 0.709693466363636

00:26:40.070 --> 00:26:41.370 dose and then these
NOTE Confidence: 0.907544118461538

00:26:41.446 --> 00:26:45.538 measures are made day 43 out other
NOTE Confidence: 0.907544118461538

00:26:45.538 --> 00:26:47.210 secondary measures pretty much
NOTE Confidence: 0.907544118461538

00:26:47.210 --> 00:26:50.156 across the board you you can look at
NOTE Confidence: 0.907544118461538

00:26:50.156 --> 00:26:51.920 and see how they were carried out.
NOTE Confidence: 0.907544118461538

00:26:51.920 --> 00:26:54.557 I did forget to say early on that these
NOTE Confidence: 0.907544118461538

00:26:54.557 --> 00:26:56.918 ratings are all done by central Raiders.
NOTE Confidence: 0.907544118461538

00:26:56.920 --> 00:26:59.680 So in order to address the
NOTE Confidence: 0.907544118461538

00:26:59.680 --> 00:27:03.240 the unblinding of the Raider,
NOTE Confidence: 0.907544118461538

00:27:03.240 --> 00:27:05.298 all these ratings were done by
NOTE Confidence: 0.907544118461538

00:27:05.298 --> 00:27:06.920 somebody off site calling in.
NOTE Confidence: 0.907544118461538

00:27:06.920 --> 00:27:10.591 And it was very rigorous that the
NOTE Confidence: 0.907544118461538

00:27:10.591 --> 00:27:12.697 the person doing the raid and
NOTE Confidence: 0.907544118461538

00:27:12.697 --> 00:27:15.198 did not even know at what point

NOTE Confidence: 0.907544118461538
00:27:15.200 --> 00:27:17.078 the person was in the study.
NOTE Confidence: 0.907544118461538
00:27:17.080 --> 00:27:19.408 So they didn't know if this was a
NOTE Confidence: 0.907544118461538
00:27:19.408 --> 00:27:21.325 baseline rating this is the end of
NOTE Confidence: 0.907544118461538
00:27:21.325 --> 00:27:23.080 study rating if this was a week,
NOTE Confidence: 0.907544118461538
00:27:23.080 --> 00:27:25.320 Week 2 rating, a week 1 rating.
NOTE Confidence: 0.907544118461538
00:27:25.320 --> 00:27:26.760 So they really didn't know
NOTE Confidence: 0.907544118461538
00:27:26.760 --> 00:27:28.200 everybody was treated the same.
NOTE Confidence: 0.907544118461538
00:27:28.200 --> 00:27:30.096 So these these data are pretty
NOTE Confidence: 0.907544118461538
00:27:30.096 --> 00:27:31.360 strong from that perspective.
NOTE Confidence: 0.907544118461538
00:27:31.360 --> 00:27:31.440 So
NOTE Confidence: 0.737113966666667
00:27:31.520 --> 00:27:33.560 it wasn't the same rater consistent,
NOTE Confidence: 0.737113966666667
00:27:33.560 --> 00:27:35.650 it was not the same rater
NOTE Confidence: 0.737113966666667
00:27:35.650 --> 00:27:36.800 consistently which is you know
NOTE Confidence: 0.903351388571428
00:27:36.800 --> 00:27:38.655 when you design studies that's
NOTE Confidence: 0.903351388571428
00:27:38.655 --> 00:27:41.960 something you usually do want to have.
NOTE Confidence: 0.903351388571428

00:27:41.960 --> 00:27:45.200 But in this case to try to prevent
NOTE Confidence: 0.903351388571428

00:27:45.200 --> 00:27:48.200 the functional on blinding of
NOTE Confidence: 0.903351388571428

00:27:48.200 --> 00:27:50.432 the of the rater that was a
NOTE Confidence: 0.903351388571428

00:27:50.432 --> 00:27:52.280 precaution that was put in place was
NOTE Confidence: 0.850431003076923

00:27:52.280 --> 00:27:53.750 there any formal assessment of blind
NOTE Confidence: 0.850431003076923

00:27:53.750 --> 00:27:55.319 for either the subject or the rater.
NOTE Confidence: 0.86438697

00:27:55.440 --> 00:27:56.830 So that we'll we'll talk
NOTE Confidence: 0.86438697

00:27:56.830 --> 00:27:58.641 about that's one of the major
NOTE Confidence: 0.86438697

00:27:58.641 --> 00:28:00.077 limitations of the study.
NOTE Confidence: 0.86438697

00:28:00.080 --> 00:28:01.480 So there there was not
NOTE Confidence: 0.768518528571428

00:28:03.640 --> 00:28:05.953 the other I I I think the the flip
NOTE Confidence: 0.768518528571428

00:28:05.953 --> 00:28:08.000 side the real important data from
NOTE Confidence: 0.768518528571428

00:28:08.000 --> 00:28:10.120 the study of the safety data.
NOTE Confidence: 0.768518528571428

00:28:10.120 --> 00:28:14.274 So here you can see that 88% or
NOTE Confidence: 0.768518528571428

00:28:14.274 --> 00:28:16.896 44 or 50 other participants in
NOTE Confidence: 0.768518528571428

00:28:16.896 --> 00:28:19.848 psilocybin and 33 or out of 54 or

NOTE Confidence: 0.768518528571428
00:28:19.848 --> 00:28:22.088 61% receiving niacin reported at
NOTE Confidence: 0.768518528571428
00:28:22.088 --> 00:28:24.878 least one AE through throughout the
NOTE Confidence: 0.768518528571428
00:28:24.878 --> 00:28:28.840 end of the the 43 day follow up.
NOTE Confidence: 0.768518528571428
00:28:28.840 --> 00:28:32.072 But I think a little bit more important
NOTE Confidence: 0.768518528571428
00:28:32.072 --> 00:28:34.598 is what would be considered drug related.
NOTE Confidence: 0.768518528571428
00:28:34.600 --> 00:28:36.749 So the investigator at least said it's
NOTE Confidence: 0.768518528571428
00:28:36.749 --> 00:28:38.591 possible that that these side effects
NOTE Confidence: 0.768518528571428
00:28:38.591 --> 00:28:40.889 were related to the drug and you can
NOTE Confidence: 0.768518528571428
00:28:40.889 --> 00:28:43.902 say here that it was 41 of the 50 or
NOTE Confidence: 0.768518528571428
00:28:43.902 --> 00:28:47.918 82% of the patients in the psilocybin group.
NOTE Confidence: 0.768518528571428
00:28:47.920 --> 00:28:51.196 And you can see it was 24 of the
NOTE Confidence: 0.768518528571428
00:28:51.196 --> 00:28:54.800 54 or 44% in the niacin group
NOTE Confidence: 0.768518528571428
00:28:54.800 --> 00:28:57.299 experienced at least an AE that was
NOTE Confidence: 0.768518528571428
00:28:57.299 --> 00:28:59.399 thought to be drug related.
NOTE Confidence: 0.768518528571428
00:28:59.400 --> 00:29:01.633 And then even bumping it up a
NOTE Confidence: 0.768518528571428

00:29:01.633 --> 00:29:03.806 little bit more in importance is
NOTE Confidence: 0.768518528571428

00:29:03.806 --> 00:29:06.080 more of the severe related AES.
NOTE Confidence: 0.768518528571428

00:29:06.080 --> 00:29:09.120 And this is through the first week really.
NOTE Confidence: 0.768518528571428

00:29:09.120 --> 00:29:12.312 I reported four out of the 50
NOTE Confidence: 0.768518528571428

00:29:12.312 --> 00:29:14.120 participants receiving psilocybin or 8%
NOTE Confidence: 0.768518528571428

00:29:14.120 --> 00:29:16.796 of those that consisted of migraines,
NOTE Confidence: 0.768518528571428

00:29:16.800 --> 00:29:21.280 headaches, illusions, panic attacks,
NOTE Confidence: 0.768518528571428

00:29:21.280 --> 00:29:22.400 paranoia.
NOTE Confidence: 0.768518528571428

00:29:22.400 --> 00:29:25.557 There were none in the Niacin group,
NOTE Confidence: 0.768518528571428

00:29:25.560 --> 00:29:27.800 but I I'm bumping it up one more
NOTE Confidence: 0.768518528571428

00:29:27.800 --> 00:29:29.820 in level of importance and this
NOTE Confidence: 0.768518528571428

00:29:29.820 --> 00:29:31.070 is obviously the most important
NOTE Confidence: 0.768518528571428

00:29:31.070 --> 00:29:33.193 all of these there were no serious
NOTE Confidence: 0.768518528571428

00:29:33.193 --> 00:29:34.513 treatment emergent adverse events.
NOTE Confidence: 0.768518528571428

00:29:34.520 --> 00:29:36.221 So for those of you that aren't
NOTE Confidence: 0.768518528571428

00:29:36.221 --> 00:29:37.320 familiar with clinical trials,

NOTE Confidence: 0.768518528571428

00:29:37.320 --> 00:29:40.145 a serious adverse event basically

NOTE Confidence: 0.768518528571428

00:29:40.145 --> 00:29:41.275 means hospitalization,

NOTE Confidence: 0.768518528571428

00:29:41.280 --> 00:29:47.088 risk of death or some other long

NOTE Confidence: 0.768518528571428

00:29:47.088 --> 00:29:49.640 term impairment that results from it.

NOTE Confidence: 0.768518528571428

00:29:49.640 --> 00:29:52.358 So nothing along those lines occurred,

NOTE Confidence: 0.578699218888889

00:29:52.920 --> 00:29:54.908 Jerry. Yeah, that difference

NOTE Confidence: 0.578699218888889

00:29:54.908 --> 00:29:57.393 between in in severe aids.

NOTE Confidence: 0.578699218888889

00:29:57.400 --> 00:29:58.672 Well, in both categories,

NOTE Confidence: 0.578699218888889

00:29:58.672 --> 00:29:59.944 I guess you're lumping

NOTE Confidence: 0.578699218888889

00:29:59.944 --> 00:30:01.360 together days one through 9,

NOTE Confidence: 0.578699218888889

00:30:01.360 --> 00:30:03.544 but we expect a lot of

NOTE Confidence: 0.578699218888889

00:30:03.544 --> 00:30:05.000 psychological effects during dosing.

NOTE Confidence: 0.578699218888889

00:30:05.000 --> 00:30:06.708 So I'd be interested in the differences

NOTE Confidence: 0.578699218888889

00:30:06.708 --> 00:30:08.548 on days two through 9 because like

NOTE Confidence: 0.578699218888889

00:30:08.548 --> 00:30:10.200 how much of that difference was,

NOTE Confidence: 0.578699218888889

00:30:10.200 --> 00:30:12.216 you know, panic attack on day one during
NOTE Confidence: 0.578699218888889

00:30:12.216 --> 00:30:13.958 dosing means something very different.
NOTE Confidence: 0.578699218888889

00:30:13.960 --> 00:30:15.640 So many attacks four days later,
NOTE Confidence: 0.8739576575

00:30:15.920 --> 00:30:17.920 so many were here we can go through.
NOTE Confidence: 0.8739576575

00:30:17.920 --> 00:30:19.360 We'll talk a little bit about some of
NOTE Confidence: 0.8739576575

00:30:19.360 --> 00:30:20.746 the more persistent ones or outside
NOTE Confidence: 0.8739576575

00:30:20.746 --> 00:30:22.222 ones because those are obviously the
NOTE Confidence: 0.8739576575

00:30:22.269 --> 00:30:23.679 ones that everybody's worried about.
NOTE Confidence: 0.8739576575

00:30:23.680 --> 00:30:26.640 Yeah, good question.
NOTE Confidence: 0.8739576575

00:30:26.640 --> 00:30:28.159 Yeah, they'll come up a little bit.
NOTE Confidence: 0.772907803076923

00:30:29.120 --> 00:30:30.513 I'm also, I'm a little surprised it's
NOTE Confidence: 0.772907803076923

00:30:30.513 --> 00:30:32.022 reported as day one through 9, yeah.
NOTE Confidence: 0.772907803076923

00:30:32.022 --> 00:30:34.314 Yeah, As opposed to day one, yeah.
NOTE Confidence: 0.772907803076923

00:30:34.314 --> 00:30:35.920 And then day two through nine. Well,
NOTE Confidence: 0.90480678625

00:30:35.920 --> 00:30:38.320 I think that people were worried about that,
NOTE Confidence: 0.90480678625

00:30:38.320 --> 00:30:40.400 that total time frame,

NOTE Confidence: 0.90480678625
00:30:40.400 --> 00:30:45.800 people meeting the regulatory agencies, OK,
NOTE Confidence: 0.90480678625
00:30:45.800 --> 00:30:48.194 when it came to solicited adverse events.
NOTE Confidence: 0.90480678625
00:30:48.200 --> 00:30:50.384 So ones that were adverse events of
NOTE Confidence: 0.90480678625
00:30:50.384 --> 00:30:52.327 interest that was specifically asked
NOTE Confidence: 0.90480678625
00:30:52.327 --> 00:30:54.880 about the most common were headache,
NOTE Confidence: 0.90480678625
00:30:54.880 --> 00:30:59.920 nausea and visual perceptual effects,
NOTE Confidence: 0.90480678625
00:30:59.920 --> 00:31:02.928 which was not so surprising I think and
NOTE Confidence: 0.90480678625
00:31:02.928 --> 00:31:06.316 you can see the relative ratios of them.
NOTE Confidence: 0.90480678625
00:31:06.320 --> 00:31:08.948 The one that may be a little bit more
NOTE Confidence: 0.90480678625
00:31:08.948 --> 00:31:10.823 concerning with the visual and perceptual
NOTE Confidence: 0.90480678625
00:31:10.823 --> 00:31:13.177 effects is 3 of the 50 patients
NOTE Confidence: 0.90480678625
00:31:13.177 --> 00:31:15.530 receiving psilocybin or 6% actually
NOTE Confidence: 0.90480678625
00:31:15.530 --> 00:31:19.580 reported some visual or perceptual
NOTE Confidence: 0.90480678625
00:31:19.580 --> 00:31:22.272 effects occurring after that first day.
NOTE Confidence: 0.90480678625
00:31:22.272 --> 00:31:25.092 So this is getting a little bit more of what
NOTE Confidence: 0.90480678625

00:31:25.092 --> 00:31:27.340 you're talking about S 6% smaller number,
NOTE Confidence: 0.90480678625

00:31:27.340 --> 00:31:30.296 but still a number that I I think
NOTE Confidence: 0.90480678625

00:31:30.296 --> 00:31:32.714 it's important to pay attention to.
NOTE Confidence: 0.90480678625

00:31:32.720 --> 00:31:33.830 And and they all did resolve
NOTE Confidence: 0.90480678625

00:31:33.830 --> 00:31:34.840 by the end of this.
NOTE Confidence: 0.90480678625

00:31:34.840 --> 00:31:34.960 Well
NOTE Confidence: 0.829558628666667

00:31:34.960 --> 00:31:36.850 what was the severity of those Because
NOTE Confidence: 0.829558628666667

00:31:36.850 --> 00:31:38.839 if that's a little flickering on day
NOTE Confidence: 0.829558628666667

00:31:38.839 --> 00:31:40.608 2 versus well formed hallucinations
NOTE Confidence: 0.829558628666667

00:31:40.608 --> 00:31:42.840 of pink elephants on day 20,
NOTE Confidence: 0.829558628666667

00:31:42.840 --> 00:31:43.880 those are very different.
NOTE Confidence: 0.829558628666667

00:31:43.880 --> 00:31:45.160 You know, so remember there were
NOTE Confidence: 0.838822982352941

00:31:45.160 --> 00:31:46.861 no SAES, so nothing reaching the level
NOTE Confidence: 0.838822982352941

00:31:46.861 --> 00:31:49.160 of an SAE and I don't think any of them,
NOTE Confidence: 0.838822982352941

00:31:49.160 --> 00:31:50.875 I I I can't say for sure,
NOTE Confidence: 0.838822982352941

00:31:50.880 --> 00:31:52.410 but I don't think any of them even met

NOTE Confidence: 0.838822982352941
00:31:52.410 --> 00:31:53.839 the level of severe adverse event.
NOTE Confidence: 0.787474428571428
00:31:54.640 --> 00:31:56.856 I I I would have to look at
NOTE Confidence: 0.787474428571428
00:31:56.856 --> 00:31:58.839 the details but I don't recall.
NOTE Confidence: 0.787474428571428
00:31:58.840 --> 00:32:01.560 I would have to agree.
NOTE Confidence: 0.787474428571428
00:32:01.560 --> 00:32:05.172 And these out of those three,
NOTE Confidence: 0.787474428571428
00:32:05.172 --> 00:32:07.491 I think it it's possible we worked
NOTE Confidence: 0.787474428571428
00:32:07.491 --> 00:32:09.504 with one of them and you're right,
NOTE Confidence: 0.787474428571428
00:32:09.504 --> 00:32:12.180 it was more flickering and it was,
NOTE Confidence: 0.787474428571428
00:32:12.180 --> 00:32:14.400 but it was not sustained or
NOTE Confidence: 0.787474428571428
00:32:14.400 --> 00:32:16.546 there was no overt hallucinate,
NOTE Confidence: 0.787474428571428
00:32:16.546 --> 00:32:17.638 frank hallucinations,
NOTE Confidence: 0.64446776
00:32:19.200 --> 00:32:20.480 no more kind of illusions.
NOTE Confidence: 0.925418549090909
00:32:21.600 --> 00:32:26.037 And I think other good news is in terms
NOTE Confidence: 0.925418549090909
00:32:26.037 --> 00:32:29.424 of any suicidal ideation or behavior,
NOTE Confidence: 0.925418549090909
00:32:29.424 --> 00:32:31.040 they're really the main
NOTE Confidence: 0.925418549090909

00:32:31.040 --> 00:32:32.240 thing is right on the bottom,
NOTE Confidence: 0.925418549090909

00:32:32.240 --> 00:32:35.150 there was no real clinically significant
NOTE Confidence: 0.925418549090909

00:32:35.150 --> 00:32:37.919 increases in suicidal ideation or behavior
NOTE Confidence: 0.925418549090909

00:32:37.920 --> 00:32:39.798 associated with the study, if any.
NOTE Confidence: 0.925418549090909

00:32:39.800 --> 00:32:42.194 If you look at any patients that
NOTE Confidence: 0.925418549090909

00:32:42.194 --> 00:32:44.124 even had some level of increase
NOTE Confidence: 0.925418549090909

00:32:44.124 --> 00:32:46.400 throughout the main part of the study,
NOTE Confidence: 0.925418549090909

00:32:46.400 --> 00:32:48.300 there were numerically more
NOTE Confidence: 0.925418549090909

00:32:48.300 --> 00:32:50.200 in the niacin group,
NOTE Confidence: 0.925418549090909

00:32:50.200 --> 00:32:52.318 about 10% compared to about 2%.
NOTE Confidence: 0.925418549090909

00:32:52.320 --> 00:32:54.870 There are just five people versus
NOTE Confidence: 0.925418549090909

00:32:54.870 --> 00:32:58.188 one person that showed some increase
NOTE Confidence: 0.925418549090909

00:32:58.188 --> 00:33:01.273 in suicidality or suicidal ideation.
NOTE Confidence: 0.925418549090909

00:33:01.280 --> 00:33:04.271 This I I think is becoming more
NOTE Confidence: 0.925418549090909

00:33:04.271 --> 00:33:05.897 important as we realize how to
NOTE Confidence: 0.925418549090909

00:33:05.897 --> 00:33:07.480 move forward with these studies.

NOTE Confidence: 0.925418549090909
00:33:07.480 --> 00:33:08.915 At least for me running these studies,
NOTE Confidence: 0.925418549090909
00:33:08.920 --> 00:33:10.426 I worry less about the people
NOTE Confidence: 0.925418549090909
00:33:10.426 --> 00:33:11.961 getting the active drug than the
NOTE Confidence: 0.925418549090909
00:33:11.961 --> 00:33:13.160 people being randomized,
NOTE Confidence: 0.925418549090909
00:33:13.160 --> 00:33:14.924 especially if there's functional
NOTE Confidence: 0.925418549090909
00:33:14.924 --> 00:33:16.868 on blinding for the patient that
NOTE Confidence: 0.925418549090909
00:33:16.868 --> 00:33:18.753 you really have to worry about
NOTE Confidence: 0.925418549090909
00:33:18.753 --> 00:33:20.463 how they're going to react to
NOTE Confidence: 0.925418549090909
00:33:20.463 --> 00:33:22.239 feeling they didn't get treatment.
NOTE Confidence: 0.925418549090909
00:33:22.240 --> 00:33:23.912 So now if we just talk about the
NOTE Confidence: 0.925418549090909
00:33:23.912 --> 00:33:25.398 limitations of the study quickly and then
NOTE Confidence: 0.925418549090909
00:33:25.398 --> 00:33:27.015 we'll be able to go into discussions
NOTE Confidence: 0.925418549090909
00:33:27.015 --> 00:33:28.919 because I think that's the main thing.
NOTE Confidence: 0.925418549090909
00:33:28.920 --> 00:33:30.976 So one of the biggest limitations and we
NOTE Confidence: 0.925418549090909
00:33:30.976 --> 00:33:32.760 mentioned it one or two times already,
NOTE Confidence: 0.925418549090909

00:33:32.760 --> 00:33:35.388 was this study was conducted between
NOTE Confidence: 0.925418549090909

00:33:35.388 --> 00:33:38.165 December of 2019 and June of 2022.
NOTE Confidence: 0.925418549090909

00:33:38.165 --> 00:33:40.595 Here is just the death rate
NOTE Confidence: 0.925418549090909

00:33:40.600 --> 00:33:42.358 due to COVID during that time.
NOTE Confidence: 0.925418549090909

00:33:42.360 --> 00:33:44.826 I mean this was smack, we started it.
NOTE Confidence: 0.925418549090909

00:33:44.826 --> 00:33:47.220 I think we were ready to run
NOTE Confidence: 0.925418549090909

00:33:47.301 --> 00:33:49.953 our first patient in like March
NOTE Confidence: 0.925418549090909

00:33:49.953 --> 00:33:52.920 just as everything was hidden.
NOTE Confidence: 0.925418549090909

00:33:52.920 --> 00:33:56.280 So this study was thrown on its head.
NOTE Confidence: 0.925418549090909

00:33:56.280 --> 00:33:57.360 That being said,
NOTE Confidence: 0.925418549090909

00:33:57.360 --> 00:33:59.160 it's a limitation because the
NOTE Confidence: 0.925418549090909

00:33:59.160 --> 00:34:01.038 study was run during COVID.
NOTE Confidence: 0.925418549090909

00:34:01.040 --> 00:34:03.420 Almost most studies run during
NOTE Confidence: 0.925418549090909

00:34:03.420 --> 00:34:06.600 this time has had major problems
NOTE Confidence: 0.925418549090909

00:34:06.600 --> 00:34:08.205 because the placebo response rates
NOTE Confidence: 0.925418549090909

00:34:08.205 --> 00:34:10.140 have been astronomical and most of

NOTE Confidence: 0.925418549090909

00:34:10.140 --> 00:34:11.520 these studies run during COVID.

NOTE Confidence: 0.925418549090909

00:34:11.520 --> 00:34:13.400 And we're not quite sure why that is.

NOTE Confidence: 0.925418549090909

00:34:13.400 --> 00:34:15.500 But many of the studies run in

NOTE Confidence: 0.925418549090909

00:34:15.500 --> 00:34:17.108 depression during the summer has

NOTE Confidence: 0.925418549090909

00:34:17.108 --> 00:34:18.810 had very abnormal elevated and

NOTE Confidence: 0.925418549090909

00:34:18.810 --> 00:34:20.635 it may be the people.

NOTE Confidence: 0.925418549090909

00:34:20.640 --> 00:34:22.754 Some of the hypothesis that's been put

NOTE Confidence: 0.925418549090909

00:34:22.754 --> 00:34:25.400 out is that people are really more

NOTE Confidence: 0.925418549090909

00:34:25.400 --> 00:34:27.440 starved for socialization and just

NOTE Confidence: 0.925418549090909

00:34:27.513 --> 00:34:29.816 the fact of getting out was actually

NOTE Confidence: 0.925418549090909

00:34:29.816 --> 00:34:32.360 working a lot you know in their favor

NOTE Confidence: 0.925418549090909

00:34:32.360 --> 00:34:34.640 but against the study drugs favor.

NOTE Confidence: 0.925418549090909

00:34:34.640 --> 00:34:36.440 And in this case if anything

NOTE Confidence: 0.925418549090909

00:34:36.440 --> 00:34:37.702 this would be something.

NOTE Confidence: 0.925418549090909

00:34:37.702 --> 00:34:39.688 The fact that we did see

NOTE Confidence: 0.925418549090909

00:34:39.688 --> 00:34:41.488 such a big effect is,
NOTE Confidence: 0.925418549090909

00:34:41.488 --> 00:34:43.456 is probably not a major limitation
NOTE Confidence: 0.925418549090909

00:34:43.456 --> 00:34:45.892 that we would use that we would
NOTE Confidence: 0.925418549090909

00:34:45.892 --> 00:34:47.280 worry about going forward.
NOTE Confidence: 0.925418549090909

00:34:47.280 --> 00:34:49.576 But there are some other limitations and
NOTE Confidence: 0.925418549090909

00:34:49.576 --> 00:34:53.080 and I think maybe the the biggest one
NOTE Confidence: 0.925418549090909

00:34:53.080 --> 00:34:55.600 moving forward is the generalizability.
NOTE Confidence: 0.925418549090909

00:34:55.600 --> 00:34:57.000 So as I mentioned this,
NOTE Confidence: 0.925418549090909

00:34:57.000 --> 00:34:58.701 you know ways that this differs than
NOTE Confidence: 0.925418549090909

00:34:58.701 --> 00:35:00.340 many of the other depression trials
NOTE Confidence: 0.925418549090909

00:35:00.340 --> 00:35:02.328 is that the male to female ratio
NOTE Confidence: 0.925418549090909

00:35:02.383 --> 00:35:04.159 is different than we typically see.
NOTE Confidence: 0.925418549090909

00:35:04.160 --> 00:35:06.400 We typically see a one to two
NOTE Confidence: 0.925418549090909

00:35:06.400 --> 00:35:07.880 male to female ratio.
NOTE Confidence: 0.925418549090909

00:35:07.880 --> 00:35:09.800 This was pretty much a one to one.
NOTE Confidence: 0.925418549090909

00:35:09.800 --> 00:35:12.088 The age was a little bit younger in

NOTE Confidence: 0.925418549090909

00:35:12.088 --> 00:35:14.192 this group and typically in trials

NOTE Confidence: 0.925418549090909

00:35:14.192 --> 00:35:16.394 of antidepressants the age is about

NOTE Confidence: 0.859233582962963

00:35:16.458 --> 00:35:18.194 45 and here it was about 41.

NOTE Confidence: 0.859233582962963

00:35:18.200 --> 00:35:20.120 I don't know if that's going

NOTE Confidence: 0.859233582962963

00:35:20.120 --> 00:35:22.240 to make any major difference,

NOTE Confidence: 0.859233582962963

00:35:22.240 --> 00:35:26.386 but there really was a failure to

NOTE Confidence: 0.859233582962963

00:35:26.386 --> 00:35:28.864 have adequate representation of the US

NOTE Confidence: 0.859233582962963

00:35:28.864 --> 00:35:31.197 population in general in this study,

NOTE Confidence: 0.859233582962963

00:35:31.200 --> 00:35:33.375 mainly in terms of race

NOTE Confidence: 0.859233582962963

00:35:33.375 --> 00:35:35.115 and even in ethnicity.

NOTE Confidence: 0.859233582962963

00:35:35.120 --> 00:35:36.578 89% were white,

NOTE Confidence: 0.859233582962963

00:35:36.578 --> 00:35:39.080 with only 3% black in the study.

NOTE Confidence: 0.922497596363636

00:35:41.280 --> 00:35:43.776 And even if you broke it down during

NOTE Confidence: 0.922497596363636

00:35:43.776 --> 00:35:45.932 other things like other demographic

NOTE Confidence: 0.922497596363636

00:35:45.932 --> 00:35:48.980 and social demographic factors in

NOTE Confidence: 0.922497596363636

00:35:48.980 --> 00:35:52.080 terms of income, education level,
NOTE Confidence: 0.922497596363636

00:35:52.080 --> 00:35:53.990 this probably wasn't as representative
NOTE Confidence: 0.922497596363636

00:35:53.990 --> 00:35:56.680 of the population as we'd like to see.
NOTE Confidence: 0.922497596363636

00:35:56.680 --> 00:35:58.588 And and I think ultimately regulators
NOTE Confidence: 0.922497596363636

00:35:58.588 --> 00:36:00.916 would like to say that this,
NOTE Confidence: 0.922497596363636

00:36:00.920 --> 00:36:03.062 this is a treatment if approved
NOTE Confidence: 0.922497596363636

00:36:03.062 --> 00:36:05.402 would work in the large majority
NOTE Confidence: 0.922497596363636

00:36:05.402 --> 00:36:08.190 or would the data they would have
NOTE Confidence: 0.922497596363636

00:36:08.190 --> 00:36:10.557 would be representative of the large
NOTE Confidence: 0.922497596363636

00:36:10.557 --> 00:36:12.916 majority of of people in the US.
NOTE Confidence: 0.922497596363636

00:36:12.920 --> 00:36:14.672 The other point was that the
NOTE Confidence: 0.922497596363636

00:36:14.672 --> 00:36:16.588 majority of these people were not
NOTE Confidence: 0.922497596363636

00:36:16.588 --> 00:36:18.318 taking antidepressants at the time.
NOTE Confidence: 0.922497596363636

00:36:18.320 --> 00:36:21.016 So you're really looking at a group of
NOTE Confidence: 0.922497596363636

00:36:21.016 --> 00:36:22.770 either antidepressant naive patients
NOTE Confidence: 0.922497596363636

00:36:22.770 --> 00:36:25.824 or at least patients that weren't at

NOTE Confidence: 0.922497596363636

00:36:25.824 --> 00:36:28.288 the level of severity that they were

NOTE Confidence: 0.922497596363636

00:36:28.288 --> 00:36:30.240 seeking treatment in some of the way.

NOTE Confidence: 0.922497596363636

00:36:30.240 --> 00:36:30.360 And

NOTE Confidence: 0.811201895

00:36:32.600 --> 00:36:34.595 then there were also patients are able

NOTE Confidence: 0.811201895

00:36:34.595 --> 00:36:36.760 to remain free of both medication and

NOTE Confidence: 0.811201895

00:36:36.760 --> 00:36:38.632 free of contact with the therapist.

NOTE Confidence: 0.811201895

00:36:38.640 --> 00:36:40.624 So these are all issues that in the

NOTE Confidence: 0.811201895

00:36:40.624 --> 00:36:42.290 future we're going to have to either

NOTE Confidence: 0.811201895

00:36:42.290 --> 00:36:44.320 determine if we're going to limit it

NOTE Confidence: 0.811201895

00:36:44.320 --> 00:36:46.840 to that or we're going to broaden the

NOTE Confidence: 0.811201895

00:36:46.840 --> 00:36:49.125 categories and and figure out other ways

NOTE Confidence: 0.811201895

00:36:49.125 --> 00:36:51.393 of managing the risk associated with that.

NOTE Confidence: 0.811201895

00:36:51.400 --> 00:36:53.080 As I said, the baseline SES was

NOTE Confidence: 0.811201895

00:36:53.080 --> 00:36:54.550 slightly lower and that that could

NOTE Confidence: 0.811201895

00:36:54.550 --> 00:36:55.954 be for a variety of reasons.

NOTE Confidence: 0.811201895

00:36:55.960 --> 00:36:57.717 I wouldn't worry too much about that.
NOTE Confidence: 0.811201895

00:36:57.720 --> 00:37:00.184 And the other main limitation is this
NOTE Confidence: 0.811201895

00:37:00.184 --> 00:37:02.342 was probably over represented for the
NOTE Confidence: 0.811201895

00:37:02.342 --> 00:37:05.331 age group and the number of people that
NOTE Confidence: 0.811201895

00:37:05.331 --> 00:37:07.836 have had previous psychedelic experiences.
NOTE Confidence: 0.811201895

00:37:07.840 --> 00:37:09.840 And as we know this can have a major impact.
NOTE Confidence: 0.811201895

00:37:09.840 --> 00:37:13.040 I've been a major impact on the validity
NOTE Confidence: 0.811201895

00:37:13.040 --> 00:37:16.039 of some of these measures that those
NOTE Confidence: 0.811201895

00:37:16.039 --> 00:37:17.992 first things I said were more related
NOTE Confidence: 0.811201895

00:37:17.992 --> 00:37:20.039 to the ecological validity of the
NOTE Confidence: 0.811201895

00:37:20.039 --> 00:37:23.277 study which we can fix by study design.
NOTE Confidence: 0.811201895

00:37:23.280 --> 00:37:25.443 And some of these future ones I'll
NOTE Confidence: 0.811201895

00:37:25.443 --> 00:37:27.822 talk about is related to really the
NOTE Confidence: 0.811201895

00:37:27.822 --> 00:37:29.677 the interpretive validity of study
NOTE Confidence: 0.811201895

00:37:29.677 --> 00:37:32.345 is you know is this due to the drug
NOTE Confidence: 0.811201895

00:37:32.345 --> 00:37:34.319 or some of the other effects.

NOTE Confidence: 0.811201895

00:37:34.320 --> 00:37:36.336 And one of the major ones that

NOTE Confidence: 0.811201895

00:37:36.336 --> 00:37:37.200 continues to the

NOTE Confidence: 0.843312586666667

00:37:39.400 --> 00:37:41.848 be dealt with or or the to be

NOTE Confidence: 0.843312586666667

00:37:41.848 --> 00:37:44.892 difficult to deal with is the struggle

NOTE Confidence: 0.843312586666667

00:37:44.892 --> 00:37:47.712 around blinding or or unmasking

NOTE Confidence: 0.843312586666667

00:37:47.712 --> 00:37:51.480 of allocation, group allocation.

NOTE Confidence: 0.843312586666667

00:37:51.480 --> 00:37:53.447 You know it's really hard when you

NOTE Confidence: 0.843312586666667

00:37:53.447 --> 00:37:56.832 have a drug that has very prominent

NOTE Confidence: 0.843312586666667

00:37:56.832 --> 00:37:59.096 and acute psychoactive effects

NOTE Confidence: 0.843312586666667

00:37:59.096 --> 00:38:01.036 in comparing it to something.

NOTE Confidence: 0.843312586666667

00:38:01.040 --> 00:38:03.077 So Niacin was used as an attempt.

NOTE Confidence: 0.843312586666667

00:38:03.080 --> 00:38:04.040 But as Chris mentioned,

NOTE Confidence: 0.843312586666667

00:38:04.040 --> 00:38:05.960 you know if you really ask somebody,

NOTE Confidence: 0.843312586666667

00:38:05.960 --> 00:38:07.920 I don't know I I can say now

NOTE Confidence: 0.843312586666667

00:38:07.920 --> 00:38:09.517 the study's over at our site.

NOTE Confidence: 0.843312586666667

00:38:09.520 --> 00:38:12.024 I think we may have had one person
NOTE Confidence: 0.843312586666667

00:38:12.024 --> 00:38:14.540 in each group that I think it's fair
NOTE Confidence: 0.843312586666667

00:38:14.540 --> 00:38:17.108 to say that we had one person that I
NOTE Confidence: 0.843312586666667

00:38:17.108 --> 00:38:18.568 think the staff would have guessed,
NOTE Confidence: 0.843312586666667

00:38:18.568 --> 00:38:19.840 got it and didn't get it.
NOTE Confidence: 0.843312586666667

00:38:19.840 --> 00:38:21.136 And I think there was one person who
NOTE Confidence: 0.843312586666667

00:38:21.136 --> 00:38:22.599 got it and staff would have guessed,
NOTE Confidence: 0.843312586666667

00:38:22.600 --> 00:38:23.080 didn't get
NOTE Confidence: 0.831229068

00:38:23.080 --> 00:38:25.920 it. It was only one exactly.
NOTE Confidence: 0.47046827

00:38:25.920 --> 00:38:27.146 So probably many one on
NOTE Confidence: 0.47046827

00:38:27.146 --> 00:38:27.876 each time. What, I don't
NOTE Confidence: 0.549498885

00:38:28.000 --> 00:38:30.120 know how many time I need to deal with 15.
NOTE Confidence: 0.645553602857143

00:38:30.120 --> 00:38:32.717 Yeah, I think 1350, I can't remember.
NOTE Confidence: 0.645553602857143

00:38:32.720 --> 00:38:35.078 Yeah. So I mean there are,
NOTE Confidence: 0.645553602857143

00:38:35.080 --> 00:38:36.040 there are cases of it,
NOTE Confidence: 0.645553602857143

00:38:36.040 --> 00:38:38.960 but I think the large majority of people,

NOTE Confidence: 0.645553602857143
00:38:38.960 --> 00:38:43.240 both participants and site members,
NOTE Confidence: 0.645553602857143
00:38:43.240 --> 00:38:44.610 staff members would would guess
NOTE Confidence: 0.645553602857143
00:38:44.610 --> 00:38:46.280 for the majority of the point
NOTE Confidence: 0.645553602857143
00:38:46.280 --> 00:38:48.170 And just an idea of how important
NOTE Confidence: 0.645553602857143
00:38:48.170 --> 00:38:50.040 some of this unmasking can be.
NOTE Confidence: 0.645553602857143
00:38:50.040 --> 00:38:51.612 There was a recent paper that
NOTE Confidence: 0.645553602857143
00:38:51.612 --> 00:38:53.689 some of you may have seen that
NOTE Confidence: 0.645553602857143
00:38:53.689 --> 00:38:55.615 came out of Stanford Boris Boris
NOTE Confidence: 0.645553602857143
00:38:55.615 --> 00:38:57.696 Heifetz paper where he was getting
NOTE Confidence: 0.645553602857143
00:38:57.696 --> 00:38:59.924 ketamine under adequate blinding
NOTE Confidence: 0.645553602857143
00:38:59.924 --> 00:39:02.436 because they're under anaesthesia.
NOTE Confidence: 0.645553602857143
00:39:02.440 --> 00:39:04.384 So he actually these are people
NOTE Confidence: 0.645553602857143
00:39:04.384 --> 00:39:05.680 going for surgical procedures
NOTE Confidence: 0.645553602857143
00:39:05.738 --> 00:39:07.703 where they were getting general
NOTE Confidence: 0.645553602857143
00:39:07.703 --> 00:39:09.275 anaesthesia during their anaesthesia
NOTE Confidence: 0.645553602857143

00:39:09.275 --> 00:39:11.068 given .5 milligrams per kilogram
NOTE Confidence: 0.645553602857143

00:39:11.068 --> 00:39:12.743 ketamine over 40 minutes during
NOTE Confidence: 0.645553602857143

00:39:12.743 --> 00:39:14.688 that time with the idea of seeing
NOTE Confidence: 0.645553602857143

00:39:14.688 --> 00:39:16.926 one of the questions is how much
NOTE Confidence: 0.645553602857143

00:39:16.926 --> 00:39:18.398 does that psychological experience,
NOTE Confidence: 0.645553602857143

00:39:18.400 --> 00:39:20.822 how necessary is that for ketamine but
NOTE Confidence: 0.645553602857143

00:39:20.822 --> 00:39:23.758 also if it's blinded will it still work.
NOTE Confidence: 0.645553602857143

00:39:23.760 --> 00:39:26.385 And I think that the results were
NOTE Confidence: 0.645553602857143

00:39:26.385 --> 00:39:28.526 actually shocking to to some people
NOTE Confidence: 0.645553602857143

00:39:28.526 --> 00:39:31.360 was and this is like a rush of
NOTE Confidence: 0.645553602857143

00:39:31.360 --> 00:39:33.358 test on itself what people see
NOTE Confidence: 0.645553602857143

00:39:33.358 --> 00:39:35.194 here and the date is small,
NOTE Confidence: 0.645553602857143

00:39:35.200 --> 00:39:36.380 so you may not see anything here.
NOTE Confidence: 0.645553602857143

00:39:36.380 --> 00:39:37.930 But I can tell you,
NOTE Confidence: 0.645553602857143

00:39:37.930 --> 00:39:41.760 I can tell you what is here is you know
NOTE Confidence: 0.645553602857143

00:39:41.760 --> 00:39:45.648 there was over 50% response at day

NOTE Confidence: 0.645553602857143
00:39:45.648 --> 00:39:47.880 one to the group that got ketamine,
NOTE Confidence: 0.645553602857143
00:39:47.880 --> 00:39:49.886 but there was nearly 50% to
NOTE Confidence: 0.645553602857143
00:39:49.886 --> 00:39:51.316 the group that got placebo.
NOTE Confidence: 0.645553602857143
00:39:51.320 --> 00:39:54.155 So there was just a huge placebo
NOTE Confidence: 0.645553602857143
00:39:54.160 --> 00:39:56.176 response rate in this in in in
NOTE Confidence: 0.645553602857143
00:39:56.176 --> 00:39:58.320 terms of their depression ratings.
NOTE Confidence: 0.645553602857143
00:39:58.320 --> 00:40:00.133 We we published sort of the commentary
NOTE Confidence: 0.645553602857143
00:40:00.133 --> 00:40:02.359 on this and there are a lot of
NOTE Confidence: 0.645553602857143
00:40:02.359 --> 00:40:03.475 problems with the interpretation
NOTE Confidence: 0.645553602857143
00:40:03.475 --> 00:40:05.481 of this study and we can go through
NOTE Confidence: 0.645553602857143
00:40:05.481 --> 00:40:06.696 that from the mechanistic side,
NOTE Confidence: 0.645553602857143
00:40:06.696 --> 00:40:08.680 but that's not the main point of it.
NOTE Confidence: 0.645553602857143
00:40:08.680 --> 00:40:11.245 I think the main point of it is to
NOTE Confidence: 0.645553602857143
00:40:11.245 --> 00:40:12.893 realize how important expectations
NOTE Confidence: 0.645553602857143
00:40:12.893 --> 00:40:15.371 and especially if you have unmasking
NOTE Confidence: 0.645553602857143

00:40:15.371 --> 00:40:16.880 going on can be.
NOTE Confidence: 0.645553602857143

00:40:16.880 --> 00:40:19.080 So if you look at the HEIFET study,
NOTE Confidence: 0.645553602857143

00:40:19.080 --> 00:40:21.680 if you look at the 19 patients that
NOTE Confidence: 0.645553602857143

00:40:21.680 --> 00:40:24.367 got ketamine and 19 patients that got
NOTE Confidence: 0.645553602857143

00:40:24.367 --> 00:40:27.080 the placebo during their surgical procedure.
NOTE Confidence: 0.645553602857143

00:40:27.080 --> 00:40:28.640 Really no difference in their
NOTE Confidence: 0.645553602857143

00:40:28.640 --> 00:40:30.600 their Madras score at two weeks.
NOTE Confidence: 0.645553602857143

00:40:30.600 --> 00:40:31.452 But if you add,
NOTE Confidence: 0.645553602857143

00:40:31.452 --> 00:40:33.324 but when they went back and asked the
NOTE Confidence: 0.645553602857143

00:40:33.324 --> 00:40:34.956 people what they thought they got,
NOTE Confidence: 0.645553602857143

00:40:34.960 --> 00:40:36.360 then you can see dramatic.
NOTE Confidence: 0.645553602857143

00:40:36.360 --> 00:40:37.886 So the people that thought they got
NOTE Confidence: 0.645553602857143

00:40:37.886 --> 00:40:39.818 it on average had a remission and the
NOTE Confidence: 0.645553602857143

00:40:39.818 --> 00:40:41.292 people that thought they didn't get
NOTE Confidence: 0.645553602857143

00:40:41.292 --> 00:40:42.876 it or didn't know didn't do so well.
NOTE Confidence: 0.645553602857143

00:40:42.880 --> 00:40:45.176 So this idea of you know what you

NOTE Confidence: 0.645553602857143

00:40:45.176 --> 00:40:47.176 think you got an expectancy really

NOTE Confidence: 0.645553602857143

00:40:47.176 --> 00:40:50.480 plays a big role although that

NOTE Confidence: 0.81315615

00:40:50.480 --> 00:40:51.400 it could work either way.

NOTE Confidence: 0.81315615

00:40:51.400 --> 00:40:52.708 It could be that they thought

NOTE Confidence: 0.81315615

00:40:52.708 --> 00:40:53.884 they didn't get ketamine because

NOTE Confidence: 0.81315615

00:40:53.884 --> 00:40:54.920 they didn't stay better.

NOTE Confidence: 0.85909678

00:40:54.920 --> 00:40:57.320 So one, one of the big problems with

NOTE Confidence: 0.85909678

00:40:57.320 --> 00:40:59.741 this study and several of the other

NOTE Confidence: 0.85909678

00:40:59.741 --> 00:41:01.818 studies including the next one I'll

NOTE Confidence: 0.85909678

00:41:01.818 --> 00:41:03.860 show you is the asking post hoc

NOTE Confidence: 0.85909678

00:41:03.860 --> 00:41:06.240 asking you really have to ask these.

NOTE Confidence: 0.85909678

00:41:06.240 --> 00:41:08.538 And that's some of the questions

NOTE Confidence: 0.85909678

00:41:08.538 --> 00:41:10.493 moving forward that we're trying

NOTE Confidence: 0.85909678

00:41:10.493 --> 00:41:12.985 to really do is to assess both

NOTE Confidence: 0.85909678

00:41:12.985 --> 00:41:14.759 expectations and and preference.

NOTE Confidence: 0.85909678

00:41:14.760 --> 00:41:17.216 And also what you think you got much
NOTE Confidence: 0.85909678

00:41:17.216 --> 00:41:19.511 closer to the time because this is
NOTE Confidence: 0.85909678

00:41:19.511 --> 00:41:22.319 this is very flawed doing it this way.
NOTE Confidence: 0.85909678

00:41:22.320 --> 00:41:23.320 Was there any
NOTE Confidence: 0.809475735

00:41:24.000 --> 00:41:25.715 Cyril asked this question in the chat
NOTE Confidence: 0.809475735

00:41:25.715 --> 00:41:27.848 earlier, was there any assessment
NOTE Confidence: 0.809475735

00:41:27.848 --> 00:41:30.680 of expectation in the Usona study?
NOTE Confidence: 0.730384885

00:41:30.960 --> 00:41:33.000 No, no, no, no.
NOTE Confidence: 0.567767144

00:41:35.080 --> 00:41:37.211 So and this is another, this is actually
NOTE Confidence: 0.567767144

00:41:37.211 --> 00:41:38.993 a study that we recently completed.
NOTE Confidence: 0.567767144

00:41:39.000 --> 00:41:42.486 This was the study funded by Pecorie
NOTE Confidence: 0.567767144

00:41:42.486 --> 00:41:45.560 which was ECT versus IV Ketamine,
NOTE Confidence: 0.567767144

00:41:45.560 --> 00:41:47.320 a 400 person study,
NOTE Confidence: 0.567767144

00:41:47.320 --> 00:41:49.080 actually a little bit more than 400 people,
NOTE Confidence: 0.567767144

00:41:49.080 --> 00:41:52.440 200 approximately 200 randomized C CHARM.
NOTE Confidence: 0.567767144

00:41:52.440 --> 00:41:55.548 You can see where there's really not

NOTE Confidence: 0.567767144

00:41:55.548 --> 00:41:57.912 the dramatic difference between the two.

NOTE Confidence: 0.567767144

00:41:57.912 --> 00:41:59.654 Actually. Numerically ketamine looked

NOTE Confidence: 0.567767144

00:41:59.654 --> 00:42:03.639 like it did a little bit better than ECT,

NOTE Confidence: 0.567767144

00:42:03.640 --> 00:42:05.754 but this was a non inferiority study.

NOTE Confidence: 0.567767144

00:42:05.760 --> 00:42:07.937 So all we can say with confidence

NOTE Confidence: 0.567767144

00:42:07.937 --> 00:42:09.872 is that ketamine was non inferior

NOTE Confidence: 0.567767144

00:42:09.872 --> 00:42:11.512 to ECT in this study.

NOTE Confidence: 0.567767144

00:42:11.520 --> 00:42:14.428 But the main point here was when we

NOTE Confidence: 0.567767144

00:42:14.428 --> 00:42:16.612 went back to look at how preference.

NOTE Confidence: 0.567767144

00:42:16.612 --> 00:42:18.754 So the idea of what they wanted

NOTE Confidence: 0.567767144

00:42:18.754 --> 00:42:20.440 and impacted how they did.

NOTE Confidence: 0.567767144

00:42:20.440 --> 00:42:22.448 And again, this is limited by the fact

NOTE Confidence: 0.567767144

00:42:22.448 --> 00:42:24.478 that we asked the preference after,

NOTE Confidence: 0.567767144

00:42:24.480 --> 00:42:28.560 not before. But if you look here,

NOTE Confidence: 0.567767144

00:42:28.560 --> 00:42:32.976 you know the the percent, I'm sorry,

NOTE Confidence: 0.567767144

00:42:32.976 --> 00:42:36.420 the percent of responders who preferred
NOTE Confidence: 0.567767144

00:42:36.514 --> 00:42:40.154 ECT but got ketamine was only about 20%.
NOTE Confidence: 0.567767144

00:42:40.160 --> 00:42:41.972 But the ones that preferred ketamine
NOTE Confidence: 0.567767144

00:42:41.972 --> 00:42:43.875 and got ketamine was nearly 60%.
NOTE Confidence: 0.567767144

00:42:43.875 --> 00:42:44.820 So the how,
NOTE Confidence: 0.567767144

00:42:44.820 --> 00:42:47.150 how the impact of preference may play
NOTE Confidence: 0.567767144

00:42:47.150 --> 00:42:49.358 on how well somebody actually does.
NOTE Confidence: 0.567767144

00:42:49.360 --> 00:42:51.160 So these are all things when
NOTE Confidence: 0.567767144

00:42:51.160 --> 00:42:52.631 we're designing these studies we
NOTE Confidence: 0.567767144

00:42:52.631 --> 00:42:53.915 just can't be blind to it.
NOTE Confidence: 0.567767144

00:42:53.920 --> 00:42:56.652 And again these are very flawed analysis
NOTE Confidence: 0.567767144

00:42:56.652 --> 00:42:58.560 because they were done post doc as it is.
NOTE Confidence: 0.567767144

00:42:58.560 --> 00:43:01.272 So the first thing is to make sure
NOTE Confidence: 0.567767144

00:43:01.272 --> 00:43:03.360 you're doing it earlier on and
NOTE Confidence: 0.567767144

00:43:03.360 --> 00:43:05.760 it's not just how it complicates,
NOTE Confidence: 0.567767144

00:43:05.760 --> 00:43:06.935 well maybe they wanted it

NOTE Confidence: 0.567767144

00:43:06.935 --> 00:43:07.875 and they got better.

NOTE Confidence: 0.567767144

00:43:07.880 --> 00:43:09.524 It's really important if you have

NOTE Confidence: 0.567767144

00:43:09.524 --> 00:43:11.177 differential drop out and in this

NOTE Confidence: 0.567767144

00:43:11.177 --> 00:43:12.934 case I did mention there was some

NOTE Confidence: 0.567767144

00:43:12.934 --> 00:43:14.196 differential drop out the people

NOTE Confidence: 0.567767144

00:43:14.196 --> 00:43:16.092 that really yeah they these people

NOTE Confidence: 0.567767144

00:43:16.092 --> 00:43:18.315 came in my guess is the majority

NOTE Confidence: 0.567767144

00:43:18.315 --> 00:43:20.812 really one and we didn't ask but I

NOTE Confidence: 0.567767144

00:43:20.812 --> 00:43:23.008 think the probably very likely they

NOTE Confidence: 0.567767144

00:43:23.008 --> 00:43:25.594 were very much looking forward to

NOTE Confidence: 0.567767144

00:43:25.594 --> 00:43:27.344 getting psilocybin when they didn't.

NOTE Confidence: 0.567767144

00:43:27.344 --> 00:43:29.679 There was some evidence of a little

NOTE Confidence: 0.567767144

00:43:29.679 --> 00:43:31.279 bit of a differential dropout

NOTE Confidence: 0.567767144

00:43:31.279 --> 00:43:33.478 because when you do look at day 48,

NOTE Confidence: 0.567767144

00:43:33.480 --> 00:43:35.258 there was only one of the psilocybin

NOTE Confidence: 0.567767144

00:43:35.258 --> 00:43:36.743 patients at day 43 that didn't
NOTE Confidence: 0.567767144

00:43:36.743 --> 00:43:37.635 make it that far,
NOTE Confidence: 0.567767144

00:43:37.640 --> 00:43:39.464 but nine of the niacin and that remember
NOTE Confidence: 0.567767144

00:43:39.464 --> 00:43:41.277 that was the primary outcome measure.
NOTE Confidence: 0.567767144

00:43:41.280 --> 00:43:43.666 So now you have to impute their data.
NOTE Confidence: 0.567767144

00:43:43.666 --> 00:43:45.904 So it really makes these trials
NOTE Confidence: 0.567767144

00:43:45.904 --> 00:43:48.398 complicated that when when you don't have
NOTE Confidence: 0.567767144

00:43:48.398 --> 00:43:50.640 data on everybody starting at the end.
NOTE Confidence: 0.567767144

00:43:50.640 --> 00:43:51.166 So yeah,
NOTE Confidence: 0.567767144

00:43:51.166 --> 00:43:52.481 to manage those preferences is
NOTE Confidence: 0.567767144

00:43:52.481 --> 00:43:54.199 going to be really important.
NOTE Confidence: 0.84524183

00:43:54.320 --> 00:43:56.540 There was no opportunity for
NOTE Confidence: 0.84524183

00:43:56.540 --> 00:43:58.760 open label follow up and
NOTE Confidence: 0.762519378

00:43:56.960 --> 00:43:59.680 there was no open label. Yeah, I
NOTE Confidence: 0.801699842

00:43:58.760 --> 00:43:59.680 know the answer to that. Yeah,
NOTE Confidence: 0.67909231

00:43:59.680 --> 00:44:00.760 there was, there was no.

NOTE Confidence: 0.67909231

00:44:00.760 --> 00:44:02.629 So that's why these things are so

NOTE Confidence: 0.67909231

00:44:02.629 --> 00:44:04.227 important in the clinical trial design

NOTE Confidence: 0.67909231

00:44:04.227 --> 00:44:07.000 is to do what you can to make sure you're

NOTE Confidence: 0.67909231

00:44:07.000 --> 00:44:08.992 not getting this differential drop out.

NOTE Confidence: 0.67909231

00:44:09.000 --> 00:44:12.492 And then lastly there there was

NOTE Confidence: 0.67909231

00:44:12.492 --> 00:44:14.696 some questions or concerns about

NOTE Confidence: 0.67909231

00:44:14.696 --> 00:44:16.716 the onset of antidepressant action.

NOTE Confidence: 0.67909231

00:44:16.720 --> 00:44:18.904 So I don't know if you, I can flip back,

NOTE Confidence: 0.67909231

00:44:18.904 --> 00:44:22.072 but take my word for it on day one,

NOTE Confidence: 0.67909231

00:44:22.072 --> 00:44:24.077 day two, one day after,

NOTE Confidence: 0.67909231

00:44:24.077 --> 00:44:25.872 there really wasn't a big

NOTE Confidence: 0.67909231

00:44:25.872 --> 00:44:27.274 antidepressant response to either

NOTE Confidence: 0.67909231

00:44:27.274 --> 00:44:28.834 the niacin or the psilocybin.

NOTE Confidence: 0.67909231

00:44:28.840 --> 00:44:31.048 And people were questioning how come

NOTE Confidence: 0.67909231

00:44:31.048 --> 00:44:33.335 most of these other studies showed

NOTE Confidence: 0.67909231

00:44:33.335 --> 00:44:35.555 a more rapid onset of benefit.
NOTE Confidence: 0.67909231

00:44:35.560 --> 00:44:36.484 And it's difficult.
NOTE Confidence: 0.67909231

00:44:36.484 --> 00:44:39.640 We don't know 100% why that happened here.
NOTE Confidence: 0.67909231

00:44:39.640 --> 00:44:41.734 But I think it's quite likely that
NOTE Confidence: 0.67909231

00:44:41.734 --> 00:44:43.839 one of the main contributors to that
NOTE Confidence: 0.67909231

00:44:43.839 --> 00:44:46.680 finding was the way that study was designed.
NOTE Confidence: 0.67909231

00:44:46.680 --> 00:44:48.448 I I mentioned in in order to try
NOTE Confidence: 0.67909231

00:44:48.448 --> 00:44:50.017 to preserve the blinding of the
NOTE Confidence: 0.67909231

00:44:50.017 --> 00:44:51.357 Raiders as much as possible.
NOTE Confidence: 0.67909231

00:44:51.360 --> 00:44:53.754 These were remote Raiders done off site.
NOTE Confidence: 0.67909231

00:44:53.760 --> 00:44:55.794 They didn't know at what point
NOTE Confidence: 0.67909231

00:44:55.794 --> 00:44:57.520 in their assessments they were.
NOTE Confidence: 0.67909231

00:44:57.520 --> 00:44:59.417 They were at one point in their
NOTE Confidence: 0.67909231

00:44:59.417 --> 00:45:01.079 treatment they were assessing them.
NOTE Confidence: 0.67909231

00:45:01.080 --> 00:45:05.800 So they asked very carefully and very
NOTE Confidence: 0.67909231

00:45:05.800 --> 00:45:08.386 rigorous about saying tell me how

NOTE Confidence: 0.67909231

00:45:08.386 --> 00:45:11.760 you've been doing for the last seven days.

NOTE Confidence: 0.67909231

00:45:11.760 --> 00:45:13.224 But so if they only received

NOTE Confidence: 0.67909231

00:45:13.224 --> 00:45:14.640 the drug one day before,

NOTE Confidence: 0.67909231

00:45:14.640 --> 00:45:16.096 you know six of those seven days

NOTE Confidence: 0.67909231

00:45:16.096 --> 00:45:17.319 were before they got the drug.

NOTE Confidence: 0.67909231

00:45:17.320 --> 00:45:19.920 So that may account for part of it.

NOTE Confidence: 0.67909231

00:45:19.920 --> 00:45:21.495 So it's so when but when was

NOTE Confidence: 0.67909231

00:45:21.495 --> 00:45:23.720 the integration session,

NOTE Confidence: 0.67909231

00:45:23.720 --> 00:45:25.160 the integration session was right,

NOTE Confidence: 0.67909231

00:45:25.160 --> 00:45:27.320 was after this right after.

NOTE Confidence: 0.67909231

00:45:27.320 --> 00:45:27.840 Yeah.

NOTE Confidence: 0.67909231

00:45:27.840 --> 00:45:29.118 Question the 1st

NOTE Confidence: 0.816027681333333

00:45:29.680 --> 00:45:31.732 in your, I mean in my experience when you

NOTE Confidence: 0.816027681333333

00:45:31.732 --> 00:45:33.790 ask patients about seven days they use

NOTE Confidence: 0.816027681333333

00:45:33.790 --> 00:45:35.722 pecan heuristic but you know they talk

NOTE Confidence: 0.816027681333333

00:45:35.722 --> 00:45:37.678 about how they are now and yeah so I I

NOTE Confidence: 0.4780106025

00:45:38.400 --> 00:45:40.240 don't get incredibly powerful

NOTE Confidence: 0.602498116666667

00:45:40.240 --> 00:45:42.316 effect made accurately across the week.

NOTE Confidence: 0.72090336625

00:45:42.400 --> 00:45:44.885 Yeah no that I mean that's why

NOTE Confidence: 0.72090336625

00:45:44.885 --> 00:45:46.950 these type of measures like the

NOTE Confidence: 0.72090336625

00:45:46.950 --> 00:45:50.120 mattress says serious limitations.

NOTE Confidence: 0.72090336625

00:45:50.120 --> 00:45:53.160 So I think this contributes part to it.

NOTE Confidence: 0.72090336625

00:45:53.160 --> 00:45:55.136 I I'm not sure what some some of

NOTE Confidence: 0.72090336625

00:45:55.136 --> 00:45:57.159 the other reason could just be that

NOTE Confidence: 0.72090336625

00:45:57.160 --> 00:45:58.870 by really blinding the Raiders

NOTE Confidence: 0.72090336625

00:45:58.870 --> 00:46:02.200 you do have some effect that way.

NOTE Confidence: 0.72090336625

00:46:02.200 --> 00:46:03.244 We don't know that,

NOTE Confidence: 0.72090336625

00:46:03.244 --> 00:46:05.143 but it's one of the things moving

NOTE Confidence: 0.72090336625

00:46:05.143 --> 00:46:07.075 forward that we have to look at.

NOTE Confidence: 0.72090336625

00:46:07.080 --> 00:46:09.312 And then the last thing that I I just

NOTE Confidence: 0.72090336625

00:46:09.312 --> 00:46:11.592 put in here is to put it in perspective

NOTE Confidence: 0.72090336625

00:46:11.592 --> 00:46:16.280 a way or or to give some context to it,

NOTE Confidence: 0.72090336625

00:46:16.280 --> 00:46:18.638 'cause these results are pretty amazing.

NOTE Confidence: 0.72090336625

00:46:18.640 --> 00:46:21.040 I mean there's there's you would be hard

NOTE Confidence: 0.72090336625

00:46:21.040 --> 00:46:23.360 pressed to say these aren't impressive.

NOTE Confidence: 0.72090336625

00:46:23.360 --> 00:46:25.004 These are they're they're

NOTE Confidence: 0.72090336625

00:46:25.004 --> 00:46:26.237 pretty big findings.

NOTE Confidence: 0.72090336625

00:46:26.240 --> 00:46:27.671 But I I did want to put them in

NOTE Confidence: 0.72090336625

00:46:27.671 --> 00:46:28.677 perspective to some of the others.

NOTE Confidence: 0.72090336625

00:46:28.680 --> 00:46:31.767 So this is to one of the ASPIRE studies

NOTE Confidence: 0.72090336625

00:46:31.767 --> 00:46:34.179 done with S ketamine looking at treatment,

NOTE Confidence: 0.72090336625

00:46:34.179 --> 00:46:35.817 the green line here I have

NOTE Confidence: 0.72090336625

00:46:35.817 --> 00:46:37.277 it faded out so you can,

NOTE Confidence: 0.72090336625

00:46:37.280 --> 00:46:39.520 I can superimpose them over.

NOTE Confidence: 0.72090336625

00:46:39.520 --> 00:46:42.250 But you can see that the response

NOTE Confidence: 0.72090336625

00:46:42.250 --> 00:46:44.388 to ketamine over S ketamine

NOTE Confidence: 0.72090336625

00:46:44.388 --> 00:46:46.690 sporvato over that time and going
NOTE Confidence: 0.72090336625

00:46:46.690 --> 00:46:48.280 out over the same time period,
NOTE Confidence: 0.72090336625

00:46:48.280 --> 00:46:49.440 that's on the graph here
NOTE Confidence: 0.857619112307692

00:46:52.280 --> 00:46:54.933 with with the extra going on for
NOTE Confidence: 0.857619112307692

00:46:54.933 --> 00:46:57.118 for the psilocybin in the study.
NOTE Confidence: 0.857619112307692

00:46:57.120 --> 00:46:58.236 But you can see, you know,
NOTE Confidence: 0.857619112307692

00:46:58.240 --> 00:47:00.438 really on par with what we're seeing.
NOTE Confidence: 0.857619112307692

00:47:00.440 --> 00:47:02.470 And remember this is just a single
NOTE Confidence: 0.857619112307692

00:47:02.470 --> 00:47:05.040 dose of psilocybin witnesses
NOTE Confidence: 0.857619112307692

00:47:05.040 --> 00:47:08.840 8 doses of esketamine here.
NOTE Confidence: 0.857619112307692

00:47:08.840 --> 00:47:11.549 But you know, one of the caveats is look
NOTE Confidence: 0.857619112307692

00:47:11.549 --> 00:47:14.720 at where the standard of care treatment is.
NOTE Confidence: 0.857619112307692

00:47:14.720 --> 00:47:16.452 Yeah, which was another,
NOTE Confidence: 0.857619112307692

00:47:16.452 --> 00:47:19.048 which was anti standard of
NOTE Confidence: 0.857619112307692

00:47:19.048 --> 00:47:20.766 antidepressant treatments, big gap.
NOTE Confidence: 0.857619112307692

00:47:20.766 --> 00:47:21.332 The biggest,

NOTE Confidence: 0.857619112307692
00:47:21.332 --> 00:47:23.030 the thing that stands alone here
NOTE Confidence: 0.857619112307692
00:47:23.083 --> 00:47:24.757 is the psilocybin is the niacin,
NOTE Confidence: 0.857619112307692
00:47:24.760 --> 00:47:26.212 not the psilocybin.
NOTE Confidence: 0.857619112307692
00:47:26.212 --> 00:47:29.667 So it's it's a lesson that for
NOTE Confidence: 0.857619112307692
00:47:29.667 --> 00:47:31.855 many trials in antidepressants
NOTE Confidence: 0.857619112307692
00:47:31.855 --> 00:47:35.110 what determines if a drug works
NOTE Confidence: 0.857619112307692
00:47:35.110 --> 00:47:37.235 or not is the placebo.
NOTE Confidence: 0.857619112307692
00:47:37.240 --> 00:47:39.580 If you if if you get a large placebo
NOTE Confidence: 0.857619112307692
00:47:39.580 --> 00:47:41.240 response it's very difficult what
NOTE Confidence: 0.855400474666667
00:47:41.240 --> 00:47:42.570 is the there are two lines there
NOTE Confidence: 0.855400474666667
00:47:42.570 --> 00:47:44.276 is one of those placebo and one of
NOTE Confidence: 0.880318715833333
00:47:44.280 --> 00:47:46.275 those. So one of these is the
NOTE Confidence: 0.880318715833333
00:47:46.275 --> 00:47:47.733 actual esketamine is the green
NOTE Confidence: 0.880318715833333
00:47:47.733 --> 00:47:49.644 line and the blue line is actually
NOTE Confidence: 0.880318715833333
00:47:49.644 --> 00:47:51.517 the standard of care treatment.
NOTE Confidence: 0.469375076666667

00:47:52.720 --> 00:47:54.196 We what it mean well in
NOTE Confidence: 0.786984796

00:47:54.200 --> 00:47:55.740 in those esketamine studies they
NOTE Confidence: 0.786984796

00:47:55.740 --> 00:47:57.280 were allowed to have regulated.
NOTE Confidence: 0.786984796

00:47:57.280 --> 00:47:59.080 So a new antidepressant was
NOTE Confidence: 0.786984796

00:47:59.080 --> 00:48:01.000 started, OK, so that that's
NOTE Confidence: 0.70193819

00:48:01.000 --> 00:48:03.920 and they had intensive interventions
NOTE Confidence: 0.79561184

00:48:04.000 --> 00:48:05.998 and they had very intensive interventions.
NOTE Confidence: 0.910782351428571

00:48:08.400 --> 00:48:12.216 And this is just comparing it to one of the
NOTE Confidence: 0.910782351428571

00:48:12.216 --> 00:48:14.424 TRANSFORM studies with S ketamine seeing
NOTE Confidence: 0.910782351428571

00:48:14.424 --> 00:48:16.944 here you don't see that rapid effects These
NOTE Confidence: 0.910782351428571

00:48:16.944 --> 00:48:18.954 those other patients were hospitalized,
NOTE Confidence: 0.910782351428571

00:48:18.960 --> 00:48:21.840 these patients weren't hospitalized.
NOTE Confidence: 0.910782351428571

00:48:21.840 --> 00:48:23.946 But again, you can see looking
NOTE Confidence: 0.910782351428571

00:48:23.946 --> 00:48:26.200 at that endpoint of four weeks,
NOTE Confidence: 0.910782351428571

00:48:26.200 --> 00:48:29.830 you you can see where where it is overall
NOTE Confidence: 0.910782351428571

00:48:29.830 --> 00:48:31.720 compared to the effect with S ketamine.

NOTE Confidence: 0.910782351428571
00:48:31.720 --> 00:48:32.860 And again these are
NOTE Confidence: 0.910782351428571
00:48:32.860 --> 00:48:34.000 comparing apples and oranges.
NOTE Confidence: 0.910782351428571
00:48:34.000 --> 00:48:35.866 You have to be really careful
NOTE Confidence: 0.910782351428571
00:48:35.866 --> 00:48:36.799 comparing across studies.
NOTE Confidence: 0.910782351428571
00:48:36.800 --> 00:48:39.635 This is just to give some level of context,
NOTE Confidence: 0.910782351428571
00:48:39.640 --> 00:48:41.784 but again really to show where the big
NOTE Confidence: 0.910782351428571
00:48:41.784 --> 00:48:43.319 difference lies is with the Niacin.
NOTE Confidence: 0.910782351428571
00:48:43.320 --> 00:48:45.736 So we really have to be careful you
NOTE Confidence: 0.910782351428571
00:48:45.736 --> 00:48:48.413 know and moving forward not to get so
NOTE Confidence: 0.910782351428571
00:48:48.413 --> 00:48:50.434 confident that these effects sizes are
NOTE Confidence: 0.910782351428571
00:48:50.434 --> 00:48:53.088 going to hold up as you move forward.
NOTE Confidence: 0.910782351428571
00:48:53.088 --> 00:48:55.669 There's there's a real trend in all
NOTE Confidence: 0.910782351428571
00:48:55.669 --> 00:48:58.027 clinical trials work that going from
NOTE Confidence: 0.910782351428571
00:48:58.027 --> 00:49:00.800 phase two studies to phase three studies
NOTE Confidence: 0.910782351428571
00:49:00.800 --> 00:49:03.428 that delta between the active and and
NOTE Confidence: 0.910782351428571

00:49:03.428 --> 00:49:05.516 sham or placebo treatment narrows down.

NOTE Confidence: 0.910782351428571

00:49:05.520 --> 00:49:07.606 So you really want to make sure

NOTE Confidence: 0.910782351428571

00:49:07.606 --> 00:49:09.600 that's managed as much as possible.

NOTE Confidence: 0.910782351428571

00:49:09.600 --> 00:49:10.440 And then finally, Curtis,

NOTE Confidence: 0.828912814285714

00:49:10.480 --> 00:49:12.106 how would you interpret that the

NOTE Confidence: 0.828912814285714

00:49:12.106 --> 00:49:13.879 fact that there's such a such a low,

NOTE Confidence: 0.828912814285714

00:49:13.880 --> 00:49:15.959 is that because of a nocebo effect,

NOTE Confidence: 0.828912814285714

00:49:15.960 --> 00:49:17.920 because of functional unblinding? Well

NOTE Confidence: 0.83094684

00:49:18.000 --> 00:49:20.184 I I mean I don't it's hard for

NOTE Confidence: 0.83094684

00:49:20.184 --> 00:49:22.480 me to know for sure. I think

NOTE Confidence: 0.775509430909091

00:49:24.640 --> 00:49:27.210 I I think functional unblinding

NOTE Confidence: 0.775509430909091

00:49:27.210 --> 00:49:30.720 probably plays a a large role in it.

NOTE Confidence: 0.908663905882353

00:49:32.840 --> 00:49:35.252 So we don't really know but I think we

NOTE Confidence: 0.908663905882353

00:49:35.252 --> 00:49:37.634 have to be careful as we move forward.

NOTE Confidence: 0.908663905882353

00:49:37.640 --> 00:49:40.200 I think expectations, functional

NOTE Confidence: 0.908663905882353

00:49:40.200 --> 00:49:42.400 unblinding play a large role and

NOTE Confidence: 0.701243884285714
00:49:42.400 --> 00:49:43.492 lack of open labels.
NOTE Confidence: 0.701243884285714
00:49:43.492 --> 00:49:45.130 So patients had really fun to
NOTE Confidence: 0.701243884285714
00:49:45.193 --> 00:49:46.997 because of functional unblinding,
NOTE Confidence: 0.648720745625
00:49:47.000 --> 00:49:48.135 that's going to increase the
NOTE Confidence: 0.648720745625
00:49:48.135 --> 00:49:49.043 nocebo effect because there's
NOTE Confidence: 0.648720745625
00:49:49.043 --> 00:49:50.160 going to be a sense of despair
NOTE Confidence: 0.511063413333333
00:49:50.200 --> 00:49:53.838 and betrayal. Yeah. So the questions yeah,
NOTE Confidence: 0.727109286666667
00:49:54.480 --> 00:49:55.116 I was going to ask you,
NOTE Confidence: 0.727109286666667
00:49:55.120 --> 00:49:57.060 how do you feel as well about the psycho,
NOTE Confidence: 0.727109286666667
00:49:57.060 --> 00:49:58.960 like the control psychotherapy?
NOTE Confidence: 0.727109286666667
00:49:58.960 --> 00:50:01.940 Because there's recently being an editorial.
NOTE Confidence: 0.727109286666667
00:50:01.940 --> 00:50:03.520 Right. Unless psychiatry on this.
NOTE Confidence: 0.727109286666667
00:50:03.520 --> 00:50:05.200 And I think it's really valuable because
NOTE Confidence: 0.9191937875
00:50:05.600 --> 00:50:07.230 what the control condition is
NOTE Confidence: 0.9191937875
00:50:07.230 --> 00:50:09.466 given the patients is this form
NOTE Confidence: 0.9191937875

00:50:09.466 --> 00:50:11.096 of psychotherapy that has never
NOTE Confidence: 0.9191937875

00:50:11.096 --> 00:50:13.960 been tested for anything. So
NOTE Confidence: 0.8443720125

00:50:13.960 --> 00:50:15.880 it's it is truly placebo
NOTE Confidence: 0.904478775

00:50:15.880 --> 00:50:17.504 in no way because it's not like
NOTE Confidence: 0.904478775

00:50:17.504 --> 00:50:18.679 we're giving them let's say
NOTE Confidence: 0.658188848333333

00:50:18.880 --> 00:50:22.480 CBT active behavioral therapy you know
NOTE Confidence: 0.658188848333333

00:50:22.480 --> 00:50:24.398 behavioral activation story and then
NOTE Confidence: 0.586092604

00:50:24.400 --> 00:50:25.420 giving them you know you
NOTE Confidence: 0.586092604

00:50:25.420 --> 00:50:26.440 know silos happened or not.
NOTE Confidence: 0.586092604

00:50:26.440 --> 00:50:28.000 I wonder how you think you can
NOTE Confidence: 0.8445947

00:50:28.040 --> 00:50:29.816 we can address that in the future if
NOTE Confidence: 0.8445947

00:50:29.816 --> 00:50:31.648 there is room for that even because
NOTE Confidence: 0.8445947

00:50:31.648 --> 00:50:33.400 I assume that user institute is
NOTE Confidence: 0.8445947

00:50:33.400 --> 00:50:35.220 pushing forward that we test silo
NOTE Confidence: 0.8445947

00:50:35.220 --> 00:50:37.240 cyber under that specific model.
NOTE Confidence: 0.674281812666667

00:50:37.720 --> 00:50:39.720 So I've been maybe you want to answer

NOTE Confidence: 0.674281812666667

00:50:39.720 --> 00:50:42.006 too I mean I'll stop by saying this

NOTE Confidence: 0.674281812666667

00:50:42.006 --> 00:50:43.962 is a very complicated issue because

NOTE Confidence: 0.674281812666667

00:50:43.962 --> 00:50:45.989 it actually comes to what's in your

NOTE Confidence: 0.674281812666667

00:50:45.989 --> 00:50:48.040 label when when you do get approved.

NOTE Confidence: 0.674281812666667

00:50:48.040 --> 00:50:51.524 So it it gets very complicated and has a

NOTE Confidence: 0.674281812666667

00:50:51.524 --> 00:50:55.044 lot of legal and regulatory meanings to it.

NOTE Confidence: 0.674281812666667

00:50:55.044 --> 00:50:56.984 This study was really with

NOTE Confidence: 0.674281812666667

00:50:56.984 --> 00:50:57.760 psychological assistance.

NOTE Confidence: 0.674281812666667

00:50:57.760 --> 00:51:01.560 It it was not and very clearly not

NOTE Confidence: 0.674281812666667

00:51:01.560 --> 00:51:04.960 done to augment a psychotherapy.

NOTE Confidence: 0.674281812666667

00:51:04.960 --> 00:51:05.440 And Ben I

NOTE Confidence: 0.702470845

00:51:05.480 --> 00:51:07.181 don't know if they used a model

NOTE Confidence: 0.702470845

00:51:07.181 --> 00:51:08.463 that's called non directive

NOTE Confidence: 0.702470845

00:51:08.463 --> 00:51:10.320 supportive therapy which really does

NOTE Confidence: 0.702470845

00:51:10.320 --> 00:51:12.120 not necessarily meet criteria of

NOTE Confidence: 0.702470845

00:51:12.120 --> 00:51:13.600 structured targeted psychotherapy
NOTE Confidence: 0.702470845

00:51:13.600 --> 00:51:16.360 that we would see conventionally.
NOTE Confidence: 0.702470845

00:51:16.360 --> 00:51:18.580 It still is a form of supportive
NOTE Confidence: 0.702470845

00:51:18.580 --> 00:51:20.448 therapy which really includes
NOTE Confidence: 0.702470845

00:51:20.448 --> 00:51:22.680 checking in during preparation,
NOTE Confidence: 0.702470845

00:51:22.680 --> 00:51:27.880 building rapport and getting to become
NOTE Confidence: 0.702470845

00:51:27.880 --> 00:51:30.760 familiar with the facilitators.
NOTE Confidence: 0.702470845

00:51:30.760 --> 00:51:34.036 So it that's what non directive supportive
NOTE Confidence: 0.883295029444444

00:51:36.440 --> 00:51:38.784 therapy is as opposed to in some other
NOTE Confidence: 0.883295029444444

00:51:38.784 --> 00:51:40.630 studies where it's psilocybin assisted
NOTE Confidence: 0.883295029444444

00:51:40.630 --> 00:51:42.635 psychotherapy which is much more
NOTE Confidence: 0.65621504

00:51:45.160 --> 00:51:47.764 with kind of a conventional therapy
NOTE Confidence: 0.65621504

00:51:47.764 --> 00:51:49.678 heavy as opposed to what was it here.
NOTE Confidence: 0.65621504

00:51:49.680 --> 00:51:51.780 So they try to actually use the
NOTE Confidence: 0.65621504

00:51:51.780 --> 00:51:53.808 the reason they use this non
NOTE Confidence: 0.65621504

00:51:53.808 --> 00:51:56.204 directive approach is to minimize

NOTE Confidence: 0.65621504

00:51:56.204 --> 00:51:59.600 the psychotherapy as a variable to

NOTE Confidence: 0.65621504

00:51:59.600 --> 00:52:01.960 whatever as much as it's feasible.

NOTE Confidence: 0.65621504

00:52:01.960 --> 00:52:06.517 But it remains to be seen what

NOTE Confidence: 0.65621504

00:52:06.520 --> 00:52:08.776 eventually how this will be packaged

NOTE Confidence: 0.65621504

00:52:08.776 --> 00:52:11.423 in the label and how this what

NOTE Confidence: 0.65621504

00:52:11.423 --> 00:52:13.553 sort of training will be required

NOTE Confidence: 0.65621504

00:52:13.553 --> 00:52:15.984 to say that you are certified to

NOTE Confidence: 0.65621504

00:52:15.984 --> 00:52:18.144 actually deliver this treatment with

NOTE Confidence: 0.65621504

00:52:18.144 --> 00:52:20.320 a non directive supported therapy.

NOTE Confidence: 0.65621504

00:52:20.320 --> 00:52:24.478 I that that remains to be seen

NOTE Confidence: 0.847678335833333

00:52:24.960 --> 00:52:27.616 there there. I mean the first thing is

NOTE Confidence: 0.847678335833333

00:52:27.616 --> 00:52:30.120 let's just demonstrate that there is

NOTE Confidence: 0.847678335833333

00:52:30.120 --> 00:52:32.160 efficacy in general safety and then

NOTE Confidence: 0.847678335833333

00:52:32.160 --> 00:52:34.880 I think decomposing the treatments.

NOTE Confidence: 0.847678335833333

00:52:34.880 --> 00:52:37.193 But as you see right down at the bottom

NOTE Confidence: 0.847678335833333

00:52:37.200 --> 00:52:38.838 just very quickly things moving forward.

NOTE Confidence: 0.847678335833333

00:52:38.840 --> 00:52:40.905 I think the field needs to address

NOTE Confidence: 0.847678335833333

00:52:40.905 --> 00:52:42.344 this to address the generalizability.

NOTE Confidence: 0.847678335833333

00:52:42.344 --> 00:52:44.358 And I can tell you future studies

NOTE Confidence: 0.847678335833333

00:52:44.358 --> 00:52:46.052 are working hard to make sure it's

NOTE Confidence: 0.847678335833333

00:52:46.052 --> 00:52:48.400 a more representative sample of

NOTE Confidence: 0.847678335833333

00:52:48.400 --> 00:52:50.640 participants address the safety.

NOTE Confidence: 0.847678335833333

00:52:50.640 --> 00:52:51.399 And longer term,

NOTE Confidence: 0.847678335833333

00:52:51.399 --> 00:52:53.170 you know if these are going to

NOTE Confidence: 0.847678335833333

00:52:53.230 --> 00:52:56.680 move forward to approval,

NOTE Confidence: 0.847678335833333

00:52:56.680 --> 00:52:58.318 the clinicians and the regulators are

NOTE Confidence: 0.847678335833333

00:52:58.318 --> 00:53:00.704 going to want to have some more information

NOTE Confidence: 0.847678335833333

00:53:00.704 --> 00:53:02.558 about what if the person relapses,

NOTE Confidence: 0.847678335833333

00:53:02.560 --> 00:53:04.220 can you do this again, how,

NOTE Confidence: 0.847678335833333

00:53:04.220 --> 00:53:06.320 how do you manage this?

NOTE Confidence: 0.847678335833333

00:53:06.320 --> 00:53:07.545 So that's more data that's

NOTE Confidence: 0.847678335833333

00:53:07.545 --> 00:53:09.360 going to need to be collected,

NOTE Confidence: 0.847678335833333

00:53:09.360 --> 00:53:10.685 a better understanding of the

NOTE Confidence: 0.847678335833333

00:53:10.685 --> 00:53:11.480 dose response relationship,

NOTE Confidence: 0.847678335833333

00:53:11.480 --> 00:53:13.600 that's only just one dose.

NOTE Confidence: 0.847678335833333

00:53:13.600 --> 00:53:14.656 We're going to have to have

NOTE Confidence: 0.847678335833333

00:53:14.656 --> 00:53:15.360 a better understanding is,

NOTE Confidence: 0.847678335833333

00:53:15.360 --> 00:53:18.640 is that dose necessary.

NOTE Confidence: 0.847678335833333

00:53:18.640 --> 00:53:21.066 And then as I said,

NOTE Confidence: 0.847678335833333

00:53:21.066 --> 00:53:22.340 we're going to have to figure out

NOTE Confidence: 0.847678335833333

00:53:22.385 --> 00:53:23.743 how to manage some of the trial

NOTE Confidence: 0.847678335833333

00:53:23.743 --> 00:53:25.118 design issues a little bit better,

NOTE Confidence: 0.847678335833333

00:53:25.120 --> 00:53:27.100 asking about expectations,

NOTE Confidence: 0.847678335833333

00:53:27.100 --> 00:53:29.080 asking about preference,

NOTE Confidence: 0.847678335833333

00:53:29.080 --> 00:53:32.540 that guess of a study assignment,

NOTE Confidence: 0.847678335833333

00:53:32.540 --> 00:53:34.446 those will all be studies.

NOTE Confidence: 0.847678335833333

00:53:34.446 --> 00:53:36.276 And then really moving forward,

NOTE Confidence: 0.847678335833333

00:53:36.280 --> 00:53:38.180 getting more to your question

NOTE Confidence: 0.847678335833333

00:53:38.180 --> 00:53:40.520 is you know how are we going to

NOTE Confidence: 0.847678335833333

00:53:40.520 --> 00:53:41.480 address the cost effectiveness?

NOTE Confidence: 0.847678335833333

00:53:41.480 --> 00:53:42.149 You know how,

NOTE Confidence: 0.847678335833333

00:53:42.149 --> 00:53:44.125 how are we going to do this in a

NOTE Confidence: 0.847678335833333

00:53:44.125 --> 00:53:45.998 way that would increase access

NOTE Confidence: 0.847678335833333

00:53:45.998 --> 00:53:49.034 and and reduce the overall cost.

NOTE Confidence: 0.847678335833333

00:53:49.040 --> 00:53:50.510 And then a better understanding of

NOTE Confidence: 0.847678335833333

00:53:50.510 --> 00:53:51.774 the mechanism, mechanistic actions.

NOTE Confidence: 0.847678335833333

00:53:51.774 --> 00:53:54.096 And then a quick review of

NOTE Confidence: 0.847678335833333

00:53:54.096 --> 00:53:55.240 everything just here.

NOTE Confidence: 0.847678335833333

00:53:55.240 --> 00:53:56.120 This is the main point.

NOTE Confidence: 0.847678335833333

00:53:56.120 --> 00:53:59.444 It's a huge, that's a huge effect size.

NOTE Confidence: 0.847678335833333

00:53:59.444 --> 00:54:04.680 I mean it the this is is encouraging.

NOTE Confidence: 0.847678335833333

00:54:04.680 --> 00:54:05.520 Those of you that know me,

NOTE Confidence: 0.847678335833333

00:54:05.520 --> 00:54:07.975 I'm quite sceptical coming into

NOTE Confidence: 0.847678335833333

00:54:07.975 --> 00:54:10.680 this and my scepticism share is

NOTE Confidence: 0.847678335833333

00:54:10.680 --> 00:54:12.960 across the board for everything.

NOTE Confidence: 0.847678335833333

00:54:12.960 --> 00:54:14.647 I think that's the way you should

NOTE Confidence: 0.847678335833333

00:54:14.647 --> 00:54:16.200 be running these clinical trials.

NOTE Confidence: 0.847678335833333

00:54:16.200 --> 00:54:18.416 But I have to say this is quite

NOTE Confidence: 0.847678335833333

00:54:18.416 --> 00:54:20.479 encouraging that you know it's it's there.

NOTE Confidence: 0.847678335833333

00:54:20.480 --> 00:54:22.262 I think there's very little doubt

NOTE Confidence: 0.847678335833333

00:54:22.262 --> 00:54:24.199 that in this study design this,

NOTE Confidence: 0.847678335833333

00:54:24.200 --> 00:54:27.432 it's a very large effect and then

NOTE Confidence: 0.847678335833333

00:54:27.432 --> 00:54:29.364 really the support of so many people

NOTE Confidence: 0.847678335833333

00:54:29.364 --> 00:54:31.064 that just at this yell site just

NOTE Confidence: 0.847678335833333

00:54:31.064 --> 00:54:33.327 to give you a sense of how many

NOTE Confidence: 0.847678335833333

00:54:33.327 --> 00:54:34.952 people contribute to these studies.

NOTE Confidence: 0.847678335833333

00:54:34.960 --> 00:54:37.860 So this is 11 sites doing this

NOTE Confidence: 0.847678335833333

00:54:37.860 --> 00:54:39.240 and this is just an oversight.
NOTE Confidence: 0.8476783358333333

00:54:39.240 --> 00:54:42.012 So and really the patients participating
NOTE Confidence: 0.8476783358333333

00:54:42.012 --> 00:54:44.722 in this especially during the pandemic
NOTE Confidence: 0.8476783358333333

00:54:44.722 --> 00:54:46.600 were were really troopers and and
NOTE Confidence: 0.8476783358333333

00:54:46.600 --> 00:54:48.394 they all needed support systems that
NOTE Confidence: 0.8476783358333333

00:54:48.394 --> 00:54:49.960 would bring them back and forth.
NOTE Confidence: 0.8476783358333333

00:54:49.960 --> 00:54:51.544 So really required a lot of
NOTE Confidence: 0.8476783358333333

00:54:51.544 --> 00:54:53.005 work both from the participants
NOTE Confidence: 0.8476783358333333

00:54:53.005 --> 00:54:55.300 and all the staff and faculty.
NOTE Confidence: 0.8476783358333333

00:54:55.300 --> 00:54:57.400 So I'll stop with that.
NOTE Confidence: 0.8476783358333333

00:54:57.400 --> 00:55:00.079 All right, thanks.
NOTE Confidence: 0.8476783358333333

00:55:00.080 --> 00:55:00.810 Any question?
NOTE Confidence: 0.8476783358333333

00:55:00.810 --> 00:55:01.175 Oh,
NOTE Confidence: 0.8476783358333333

00:55:01.175 --> 00:55:02.635 I should probably go
NOTE Confidence: 0.6107969266666667

00:55:09.810 --> 00:55:12.846 kind of are there any questions,
NOTE Confidence: 0.6107969266666667

00:55:12.850 --> 00:55:13.588 Terry, why don't you stop

NOTE Confidence: 0.610796926666667
00:55:13.588 --> 00:55:14.698 the share so we can see it?
NOTE Confidence: 0.610796926666667
00:55:14.698 --> 00:55:15.930 Yeah, let me stop share
NOTE Confidence: 0.815341936111111
00:55:31.280 --> 00:55:33.278 the questions about the blinding and
NOTE Confidence: 0.815341936111111
00:55:33.278 --> 00:55:35.425 expectancy that I think we've addressed
NOTE Confidence: 0.815341936111111
00:55:35.425 --> 00:55:37.675 a question about modifying the Madras,
NOTE Confidence: 0.815341936111111
00:55:37.680 --> 00:55:40.299 the time frame of the Madras in order to
NOTE Confidence: 0.815341936111111
00:55:40.299 --> 00:55:43.920 try to avoid that day two point but that
NOTE Confidence: 0.897001289285714
00:55:44.880 --> 00:55:48.723 So I I think honestly if for these clinical
NOTE Confidence: 0.897001289285714
00:55:48.723 --> 00:55:51.560 trials that are going through FDA,
NOTE Confidence: 0.897001289285714
00:55:51.560 --> 00:55:54.640 the FDA doesn't care about day two,
NOTE Confidence: 0.897001289285714
00:55:54.640 --> 00:55:56.060 right. It's just not it.
NOTE Confidence: 0.897001289285714
00:55:56.060 --> 00:55:57.960 You know they're much more
NOTE Confidence: 0.897001289285714
00:55:57.960 --> 00:55:59.480 interested in this endpoint.
NOTE Confidence: 0.897001289285714
00:55:59.480 --> 00:56:01.000 So you can modify it if you want,
NOTE Confidence: 0.897001289285714
00:56:01.000 --> 00:56:03.072 but it's always going to be a
NOTE Confidence: 0.897001289285714

00:56:03.072 --> 00:56:06.220 secondary or an exploratory aim here
NOTE Confidence: 0.897001289285714

00:56:06.220 --> 00:56:09.175 because the I don't think the FDA
NOTE Confidence: 0.897001289285714

00:56:09.175 --> 00:56:10.670 is interested in giving approval
NOTE Confidence: 0.897001289285714

00:56:10.727 --> 00:56:12.343 for rapid acting specifically
NOTE Confidence: 0.897001289285714

00:56:12.343 --> 00:56:14.363 that's they're they're much more
NOTE Confidence: 0.897001289285714

00:56:14.363 --> 00:56:15.879 interested in the longer term.
NOTE Confidence: 0.875681539285714

00:56:17.000 --> 00:56:18.456 It was an early question I don't
NOTE Confidence: 0.875681539285714

00:56:18.456 --> 00:56:20.078 think we talked about about the music.
NOTE Confidence: 0.875681539285714

00:56:20.080 --> 00:56:21.872 I think you everyone listens to the
NOTE Confidence: 0.875681539285714

00:56:21.872 --> 00:56:24.160 same playlist in the same order. Yes.
NOTE Confidence: 0.875681539285714

00:56:24.160 --> 00:56:26.360 And sequence which is an interesting,
NOTE Confidence: 0.875681539285714

00:56:26.360 --> 00:56:28.060 I mean that's going to be more that's going
NOTE Confidence: 0.875681539285714

00:56:28.060 --> 00:56:29.722 to be a perfect fit for some people and
NOTE Confidence: 0.875681539285714

00:56:29.722 --> 00:56:31.360 kind of weird for other people, right.
NOTE Confidence: 0.875681539285714

00:56:31.360 --> 00:56:35.250 So that's some participants and that may
NOTE Confidence: 0.875681539285714

00:56:35.250 --> 00:56:36.800 interact with ethnicity or cultural.

NOTE Confidence: 0.875681539285714
00:56:36.800 --> 00:56:38.240 Absolutely. Yep.
NOTE Confidence: 0.847593806
00:56:39.800 --> 00:56:40.480 And I think this is,
NOTE Confidence: 0.847593806
00:56:40.480 --> 00:56:41.960 this is the general idea,
NOTE Confidence: 0.847593806
00:56:41.960 --> 00:56:43.640 the whole idea of generalizability.
NOTE Confidence: 0.76980329
00:56:43.960 --> 00:56:44.840 And and I think
NOTE Confidence: 0.902346942857143
00:56:45.120 --> 00:56:47.115 one of the comments in the bottom,
NOTE Confidence: 0.902346942857143
00:56:47.120 --> 00:56:48.560 Chris, if you don't mind scrolling,
NOTE Confidence: 0.902346942857143
00:56:48.560 --> 00:56:51.670 Steve made a really good
NOTE Confidence: 0.902346942857143
00:56:51.670 --> 00:56:55.954 point which ties into, yeah,
NOTE Confidence: 0.902346942857143
00:56:55.954 --> 00:56:58.078 that's the one, ties into the
NOTE Confidence: 0.664185198571429
00:57:00.920 --> 00:57:02.975 functional unblinding potential
NOTE Confidence: 0.664185198571429
00:57:02.975 --> 00:57:07.355 NASEBO effect is that because of the
NOTE Confidence: 0.664185198571429
00:57:07.355 --> 00:57:10.080 uniqueness of the study designed
NOTE Confidence: 0.664185198571429
00:57:10.080 --> 00:57:12.313 patients who knew that they did not
NOTE Confidence: 0.664185198571429
00:57:12.313 --> 00:57:14.225 receive psilocybin and they had to stay
NOTE Confidence: 0.664185198571429

00:57:14.225 --> 00:57:16.238 in the study for a total of 6 weeks.
NOTE Confidence: 0.664185198571429

00:57:16.240 --> 00:57:18.360 That was very challenging also on the staff,
NOTE Confidence: 0.664185198571429

00:57:18.360 --> 00:57:22.332 both on the facilitators as well
NOTE Confidence: 0.664185198571429

00:57:22.332 --> 00:57:24.896 as research assistants who work
NOTE Confidence: 0.664185198571429

00:57:24.896 --> 00:57:26.885 more closely with participants.
NOTE Confidence: 0.664185198571429

00:57:26.885 --> 00:57:30.275 It was just a shared heartfelt
NOTE Confidence: 0.664185198571429

00:57:30.280 --> 00:57:31.200 experience. And
NOTE Confidence: 0.8450657375

00:57:31.440 --> 00:57:33.008 when it's easy to imagine there being
NOTE Confidence: 0.8450657375

00:57:33.008 --> 00:57:34.240 a feedback process that's right,
NOTE Confidence: 0.8450657375

00:57:34.240 --> 00:57:36.008 the distress and the staff and the distress
NOTE Confidence: 0.8450657375

00:57:36.008 --> 00:57:37.798 and the patient over that time period,
NOTE Confidence: 0.8450657375

00:57:37.800 --> 00:57:39.150 well that's the importance of
NOTE Confidence: 0.8450657375

00:57:39.150 --> 00:57:40.960 the of the raters being blinded.
NOTE Confidence: 0.8450657375

00:57:40.960 --> 00:57:43.396 But that's that's that could have been.
NOTE Confidence: 0.89831017375

00:57:44.040 --> 00:57:45.640 So thank you Steve for bringing that up.
NOTE Confidence: 0.762250812

00:57:48.680 --> 00:57:51.080 Any other questions or hi,

NOTE Confidence: 0.762250812

00:57:51.080 --> 00:57:53.352 I have a question.

NOTE Confidence: 0.762250812

00:57:53.352 --> 00:57:57.568 So in the paper there's mention of three

NOTE Confidence: 0.762250812

00:57:57.568 --> 00:58:00.200 people in both groups that started

NOTE Confidence: 0.762250812

00:58:00.200 --> 00:58:02.520 an antidepressant during the study.

NOTE Confidence: 0.762250812

00:58:02.520 --> 00:58:06.158 To clarify, for any of the participants

NOTE Confidence: 0.762250812

00:58:06.160 --> 00:58:07.720 that were on a medication,

NOTE Confidence: 0.762250812

00:58:07.720 --> 00:58:11.540 did they restart or was it only patients

NOTE Confidence: 0.762250812

00:58:11.540 --> 00:58:14.600 that had not been on a medication

NOTE Confidence: 0.951262945

00:58:14.600 --> 00:58:15.600 prior to the study?

NOTE Confidence: 0.807810771818182

00:58:17.640 --> 00:58:19.890 So I I I can only I think we

NOTE Confidence: 0.807810771818182

00:58:19.890 --> 00:58:22.628 had one at our site and I think

NOTE Confidence: 0.807810771818182

00:58:22.628 --> 00:58:24.520 that patient was a restart.

NOTE Confidence: 0.807810771818182

00:58:24.520 --> 00:58:26.256 I the other ones, I don't know

NOTE Confidence: 0.807810771818182

00:58:26.256 --> 00:58:28.140 for sure, I can't answer.

NOTE Confidence: 0.807810771818182

00:58:28.140 --> 00:58:31.088 But these are people that you know

NOTE Confidence: 0.807810771818182

00:58:31.088 --> 00:58:33.356 the the way the protocol was written
NOTE Confidence: 0.807810771818182

00:58:33.356 --> 00:58:35.780 was patients were supposed to
NOTE Confidence: 0.807810771818182

00:58:35.780 --> 00:58:38.280 remain off all oral antidepressant
NOTE Confidence: 0.807810771818182

00:58:38.280 --> 00:58:41.040 medications or all types of medication
NOTE Confidence: 0.807810771818182

00:58:41.040 --> 00:58:44.958 antidepressant treatments from
NOTE Confidence: 0.807810771818182

00:58:44.960 --> 00:58:47.678 baseline through or at least be
NOTE Confidence: 0.807810771818182

00:58:47.678 --> 00:58:50.857 at least five half lives prior to
NOTE Confidence: 0.807810771818182

00:58:50.857 --> 00:58:55.200 dosing until the end of day 4243.
NOTE Confidence: 0.807810771818182

00:58:55.200 --> 00:58:57.528 So that meant they were supposed
NOTE Confidence: 0.807810771818182

00:58:57.528 --> 00:58:59.982 to not have any contact with
NOTE Confidence: 0.807810771818182

00:58:59.982 --> 00:59:02.237 their therapist or to have
NOTE Confidence: 0.838044001666667

00:59:04.440 --> 00:59:06.678 any antidepressant used during that time.
NOTE Confidence: 0.838044001666667

00:59:06.680 --> 00:59:08.060 And some people just said
NOTE Confidence: 0.838044001666667

00:59:08.060 --> 00:59:09.440 they couldn't stay that long.
NOTE Confidence: 0.838044001666667

00:59:09.440 --> 00:59:11.960 It's going six weeks without any treatment.
NOTE Confidence: 0.838044001666667

00:59:11.960 --> 00:59:14.608 So I I don't know to answer your

NOTE Confidence: 0.838044001666667
00:59:14.608 --> 00:59:16.027 question specifically if these
NOTE Confidence: 0.838044001666667
00:59:16.027 --> 00:59:17.747 are people that were restarting
NOTE Confidence: 0.838044001666667
00:59:17.747 --> 00:59:19.639 or starting new ones offhand.
NOTE Confidence: 0.838044001666667
00:59:19.640 --> 00:59:22.280 And do you, did
NOTE Confidence: 0.970164023333333
00:59:22.280 --> 00:59:23.480 you account for
NOTE Confidence: 0.814345113333333
00:59:23.480 --> 00:59:26.226 how starting or restarting a
NOTE Confidence: 0.814345113333333
00:59:26.226 --> 00:59:29.520 medication affected their outcomes
NOTE Confidence: 0.8388164
00:59:29.520 --> 00:59:33.376 in terms of how the Mitras was affected
NOTE Confidence: 0.8388164
00:59:33.376 --> 00:59:36.440 with those medications? We we do not,
NOTE Confidence: 0.8388164
00:59:36.440 --> 00:59:38.360 I I'm sure we can get that data.
NOTE Confidence: 0.8388164
00:59:38.360 --> 00:59:39.638 We don't have that data offhand,
NOTE Confidence: 0.8388164
00:59:39.640 --> 00:59:42.200 but it's it's probably going to be difficult.
NOTE Confidence: 0.8388164
00:59:42.200 --> 00:59:43.892 I mean it's a small number of people and
NOTE Confidence: 0.8388164
00:59:43.892 --> 00:59:46.664 they, I think all of these people started the
NOTE Confidence: 0.8388164
00:59:46.664 --> 00:59:48.839 medication 'cause they weren't feeling well.
NOTE Confidence: 0.8388164

00:59:48.840 --> 00:59:50.864 So you're going to have a lot of
NOTE Confidence: 0.8388164

00:59:50.864 --> 00:59:52.745 regression to the mean going on things
NOTE Confidence: 0.8388164

00:59:52.745 --> 00:59:54.692 that I don't think such a small number.
NOTE Confidence: 0.8388164

00:59:54.692 --> 00:59:55.584 It's probably not going
NOTE Confidence: 0.8388164

00:59:55.584 --> 00:59:56.840 to be that informative.
NOTE Confidence: 0.8388164

00:59:56.840 --> 00:59:59.960 All right, thank you.
NOTE Confidence: 0.8388164

00:59:59.960 --> 01:00:00.760 Thank you.
NOTE Confidence: 0.8388164

01:00:00.760 --> 01:00:02.200 Hey, Jerry, thanks.
NOTE Confidence: 0.89939721

01:00:02.200 --> 01:00:04.140 Thanks for the talk question
NOTE Confidence: 0.89939721

01:00:04.140 --> 01:00:07.638 about the 22% who had previous
NOTE Confidence: 0.89939721

01:00:07.638 --> 01:00:09.276 experience with psychedelics.
NOTE Confidence: 0.89939721

01:00:09.280 --> 01:00:11.464 Was it sensitivity analysis conducted
NOTE Confidence: 0.89939721

01:00:11.464 --> 01:00:14.840 where if you took out those 22%, was there
NOTE Confidence: 0.969305176666667

01:00:14.840 --> 01:00:18.040 any change in the responses?
NOTE Confidence: 0.969305176666667

01:00:18.040 --> 01:00:19.284 Great question.
NOTE Confidence: 0.969305176666667

01:00:19.284 --> 01:00:22.394 I don't remember seeing that,

NOTE Confidence: 0.483684598

01:00:22.840 --> 01:00:24.251 Ben. I don't think so, no.

NOTE Confidence: 0.483684598

01:00:24.251 --> 01:00:26.106 But that's a really good

NOTE Confidence: 0.483684598

01:00:26.106 --> 01:00:28.355 question to what because that's a

NOTE Confidence: 0.483684598

01:00:28.355 --> 01:00:29.480 disproportionately high number.

NOTE Confidence: 0.483684598

01:00:29.480 --> 01:00:31.280 It's something it's above

NOTE Confidence: 0.483684598

01:00:31.280 --> 01:00:33.360 general population as well. So,

NOTE Confidence: 0.800592818333333

01:00:36.240 --> 01:00:38.118 and one would imagine that that

NOTE Confidence: 0.800592818333333

01:00:38.120 --> 01:00:40.960 those 22% would have the highest

NOTE Confidence: 0.857146686666667

01:00:40.960 --> 01:00:43.608 expectancies and that's why they've

NOTE Confidence: 0.857146686666667

01:00:43.608 --> 01:00:45.750 taken part in the study and it would be

NOTE Confidence: 0.857146686666667

01:00:45.750 --> 01:00:48.413 interesting to look at that. So go ahead.

NOTE Confidence: 0.857146686666667

01:00:48.413 --> 01:00:50.480 No, I'll let you go first. Yeah.

NOTE Confidence: 0.901413523

01:00:50.480 --> 01:00:54.764 No, I think there's actually a bit of a

NOTE Confidence: 0.901413523

01:00:54.764 --> 01:00:56.864 paradox when it comes to think how we

NOTE Confidence: 0.901413523

01:00:56.864 --> 01:00:59.890 think about the participant participants

NOTE Confidence: 0.901413523

01:00:59.890 --> 01:01:02.853 who had prior exposure to psilocybin
NOTE Confidence: 0.901413523

01:01:02.853 --> 01:01:04.518 and psychedelics for that matter.
NOTE Confidence: 0.901413523

01:01:04.520 --> 01:01:08.194 Specifically what I'm getting at is those
NOTE Confidence: 0.901413523

01:01:08.194 --> 01:01:13.224 who actually had prior exposure and ended
NOTE Confidence: 0.901413523

01:01:13.224 --> 01:01:17.360 up getting either psilocybin or niacin,
NOTE Confidence: 0.901413523

01:01:17.360 --> 01:01:19.864 they had less of AI should say it
NOTE Confidence: 0.901413523

01:01:19.864 --> 01:01:22.212 called a disappointment or a let
NOTE Confidence: 0.901413523

01:01:22.212 --> 01:01:24.672 down and they were less likely.
NOTE Confidence: 0.901413523

01:01:24.680 --> 01:01:27.340 Again, this is just based on my
NOTE Confidence: 0.901413523

01:01:27.340 --> 01:01:30.035 general impression in the from the
NOTE Confidence: 0.901413523

01:01:30.035 --> 01:01:33.296 participants that we worked with to have
NOTE Confidence: 0.901413523

01:01:33.296 --> 01:01:35.612 an exaggerated response because they
NOTE Confidence: 0.901413523

01:01:35.612 --> 01:01:39.000 knew sort of they were already familiar
NOTE Confidence: 0.901413523

01:01:39.085 --> 01:01:41.900 with the the effect of psilocybin.
NOTE Confidence: 0.901413523

01:01:41.900 --> 01:01:44.995 So they actually based in this conversation,
NOTE Confidence: 0.901413523

01:01:44.995 --> 01:01:47.130 they were much more realistic about what

NOTE Confidence: 0.901413523

01:01:47.184 --> 01:01:49.032 to expect that they did not come into

NOTE Confidence: 0.901413523

01:01:49.032 --> 01:01:50.854 the study thinking that this was going

NOTE Confidence: 0.901413523

01:01:50.854 --> 01:01:52.582 to be a Tennessee unlike participants

NOTE Confidence: 0.901413523

01:01:52.582 --> 01:01:55.114 with no prior exposure to psilocybin,

NOTE Confidence: 0.901413523

01:01:55.120 --> 01:01:57.538 their sense of expectancy was really

NOTE Confidence: 0.901413523

01:01:57.538 --> 01:02:00.041 through the roof and we struggled

NOTE Confidence: 0.901413523

01:02:00.041 --> 01:02:02.911 to really bring down that sense of

NOTE Confidence: 0.901413523

01:02:02.911 --> 01:02:04.834 expectancy which was really being

NOTE Confidence: 0.901413523

01:02:04.834 --> 01:02:07.673 enforced by you know as it's become

NOTE Confidence: 0.901413523

01:02:07.673 --> 01:02:11.038 dubbed as the pollen effect.

NOTE Confidence: 0.901413523

01:02:11.040 --> 01:02:13.536 Michael Pollan that is.

NOTE Confidence: 0.901413523

01:02:13.536 --> 01:02:18.336 And so it's I think prior exposure

NOTE Confidence: 0.901413523

01:02:18.336 --> 01:02:22.443 to psilocybin actually is going to

NOTE Confidence: 0.901413523

01:02:22.443 --> 01:02:24.448 give participants a more realistic

NOTE Confidence: 0.901413523

01:02:24.448 --> 01:02:26.527 expectation rather than the ones who

NOTE Confidence: 0.901413523

01:02:26.527 --> 01:02:29.108 have not had any prior exposure who are
NOTE Confidence: 0.901413523

01:02:29.108 --> 01:02:31.558 coming in really expecting a silver bullet.
NOTE Confidence: 0.901413523

01:02:31.560 --> 01:02:33.931 Like there really are just it's spelled
NOTE Confidence: 0.901413523

01:02:33.931 --> 01:02:36.157 silver bullet when they're coming in.
NOTE Confidence: 0.901413523

01:02:36.160 --> 01:02:38.998 Yeah perhaps it perhaps it contributes
NOTE Confidence: 0.901413523

01:02:39.000 --> 01:02:42.439 to lower side effects profile
NOTE Confidence: 0.877725054

01:02:43.040 --> 01:02:44.280 because they've had some experience
NOTE Confidence: 0.868966675

01:02:44.280 --> 01:02:46.000 with it before it's
NOTE Confidence: 0.784048702857143

01:02:46.200 --> 01:02:49.035 that's very likely. Yes I I absolutely.
NOTE Confidence: 0.71114726

01:02:49.720 --> 01:02:52.576 Yeah. So I I actually we've been
NOTE Confidence: 0.71114726

01:02:52.576 --> 01:02:53.824 I hadn't really talked about this
NOTE Confidence: 0.71114726

01:02:53.824 --> 01:02:55.076 before but I I I agree with Ben.
NOTE Confidence: 0.71114726

01:02:55.080 --> 01:02:57.752 I think in general people came in with
NOTE Confidence: 0.71114726

01:02:57.752 --> 01:03:00.110 such a high level of expectation that
NOTE Confidence: 0.71114726

01:03:00.110 --> 01:03:02.617 I'm not sure there was a big difference
NOTE Confidence: 0.71114726

01:03:02.617 --> 01:03:05.473 between the previous exposed and the audit.

NOTE Confidence: 0.71114726

01:03:05.480 --> 01:03:07.083 And if anything, they may have had

NOTE Confidence: 0.71114726

01:03:07.083 --> 01:03:08.680 a more realistic expectation. So,

NOTE Confidence: 0.441312625

01:03:12.360 --> 01:03:14.972 hey, Jerry, I have a question for you.

NOTE Confidence: 0.441312625

01:03:14.972 --> 01:03:16.860 You or Ben, You know,

NOTE Confidence: 0.441312625

01:03:16.860 --> 01:03:18.910 talking about the issues of

NOTE Confidence: 0.441312625

01:03:18.910 --> 01:03:20.504 generalizability and the fact that

NOTE Confidence: 0.441312625

01:03:20.504 --> 01:03:24.000 most of your population was Caucasian,

NOTE Confidence: 0.441312625

01:03:24.000 --> 01:03:27.531 what strategies you think may need to

NOTE Confidence: 0.441312625

01:03:27.531 --> 01:03:30.016 be done to actually improve on that?

NOTE Confidence: 0.441312625

01:03:30.016 --> 01:03:32.990 I mean is, are these like echoes of Tuskegee

NOTE Confidence: 0.441312625

01:03:32.990 --> 01:03:37.038 or what are your thoughts about that?

NOTE Confidence: 0.441312625

01:03:37.040 --> 01:03:39.040 Hey Gerard, first of all great great hearing.

NOTE Confidence: 0.666594638

01:03:41.160 --> 01:03:44.480 I I I think it's you know

NOTE Confidence: 0.666594638

01:03:44.480 --> 01:03:46.448 it's a multi fold issue.

NOTE Confidence: 0.666594638

01:03:46.448 --> 01:03:49.616 I mean it's always difficult getting

NOTE Confidence: 0.666594638

01:03:49.616 --> 01:03:51.676 adequate representation from all
NOTE Confidence: 0.666594638

01:03:51.676 --> 01:03:54.434 aspects of the you know the population.
NOTE Confidence: 0.666594638

01:03:54.440 --> 01:03:55.780 I think this study was
NOTE Confidence: 0.666594638

01:03:55.780 --> 01:03:57.120 unique in some ways that
NOTE Confidence: 0.8844757575

01:03:59.280 --> 01:04:02.705 among certain groups this psychedelics
NOTE Confidence: 0.8844757575

01:04:02.705 --> 01:04:06.960 is much more talked about and and much
NOTE Confidence: 0.8844757575

01:04:06.960 --> 01:04:09.170 more seen in a favourable light than
NOTE Confidence: 0.8844757575

01:04:09.170 --> 01:04:11.439 than in other groups in the population.
NOTE Confidence: 0.8844757575

01:04:11.440 --> 01:04:13.515 And I think that contributes
NOTE Confidence: 0.8844757575

01:04:13.515 --> 01:04:14.760 specifically to this.
NOTE Confidence: 0.8844757575

01:04:14.760 --> 01:04:15.640 But I think, you know,
NOTE Confidence: 0.8844757575

01:04:15.640 --> 01:04:17.832 obviously having adequate representation
NOTE Confidence: 0.8844757575

01:04:17.832 --> 01:04:22.239 is a struggle in in all of medicine.
NOTE Confidence: 0.8844757575

01:04:22.240 --> 01:04:24.392 How to how to improve that I I
NOTE Confidence: 0.8844757575

01:04:24.392 --> 01:04:26.318 think is really to make sure
NOTE Confidence: 0.867791065666667

01:04:28.560 --> 01:04:30.842 the sites are chosen well that that

NOTE Confidence: 0.867791065666667
01:04:30.842 --> 01:04:32.188 there's really adequate representation
NOTE Confidence: 0.867791065666667
01:04:32.188 --> 01:04:34.924 at each site and and I don't mean that's
NOTE Confidence: 0.867791065666667
01:04:34.987 --> 01:04:36.979 going in and just going into an area
NOTE Confidence: 0.867791065666667
01:04:36.979 --> 01:04:39.185 that has an over represented group of
NOTE Confidence: 0.867791065666667
01:04:39.185 --> 01:04:41.893 anywhere but trying to go into groups
NOTE Confidence: 0.867791065666667
01:04:41.893 --> 01:04:44.238 that have adequate broad representation.
NOTE Confidence: 0.689564852
01:04:46.880 --> 01:04:49.399 And I and I think you know getting
NOTE Confidence: 0.689564852
01:04:49.399 --> 01:04:51.064 realistic information out and getting
NOTE Confidence: 0.689564852
01:04:51.064 --> 01:04:52.925 real patience I mean that that I think
NOTE Confidence: 0.689564852
01:04:52.925 --> 01:04:54.658 is going to be that's the thing I've
NOTE Confidence: 0.689564852
01:04:54.658 --> 01:04:56.572 said over and over is you know we're
NOTE Confidence: 0.689564852
01:04:56.572 --> 01:04:58.434 we're going to really need to get
NOTE Confidence: 0.689564852
01:04:58.440 --> 01:05:00.248 active treatment seeking patients.
NOTE Confidence: 0.689564852
01:05:00.248 --> 01:05:02.508 So people that are responding
NOTE Confidence: 0.689564852
01:05:02.508 --> 01:05:04.211 specifically for a research study
NOTE Confidence: 0.689564852

01:05:04.211 --> 01:05:06.233 already are quite a unique group.
NOTE Confidence: 0.689564852

01:05:06.240 --> 01:05:07.452 So you're going to really want
NOTE Confidence: 0.689564852

01:05:07.452 --> 01:05:09.014 to get people that are much more
NOTE Confidence: 0.689564852

01:05:09.014 --> 01:05:10.199 coming out of clinical settings.
NOTE Confidence: 0.689564852

01:05:10.200 --> 01:05:11.320 So Ben, I don't know.
NOTE Confidence: 0.645885562857143

01:05:11.480 --> 01:05:15.297 Yeah, no I I think Jerry you that covers it.
NOTE Confidence: 0.645885562857143

01:05:15.297 --> 01:05:17.353 I don't think I have anything to add.
NOTE Confidence: 0.645885562857143

01:05:17.360 --> 01:05:19.920 There is actually in our group and I do not
NOTE Confidence: 0.645885562857143

01:05:19.920 --> 01:05:24.040 want to single out that tan single amount.
NOTE Confidence: 0.645885562857143

01:05:24.040 --> 01:05:25.885 I'm going to single him up.
NOTE Confidence: 0.645885562857143

01:05:25.885 --> 01:05:28.810 But who has done a lot of work in
NOTE Confidence: 0.645885562857143

01:05:28.898 --> 01:05:31.138 this area and to explore different
NOTE Confidence: 0.645885562857143

01:05:31.138 --> 01:05:34.237 strategies and how is it that we can
NOTE Confidence: 0.645885562857143

01:05:34.237 --> 01:05:37.660 actually improve our recruitment,
NOTE Confidence: 0.645885562857143

01:05:37.660 --> 01:05:40.431 diversify our recruitment and what sort
NOTE Confidence: 0.645885562857143

01:05:40.431 --> 01:05:42.813 of outreach programs that have been

NOTE Confidence: 0.645885562857143

01:05:42.813 --> 01:05:45.748 more successful in achieving this goal

NOTE Confidence: 0.645885562857143

01:05:45.748 --> 01:05:48.124 versus some that have been tried and

NOTE Confidence: 0.645885562857143

01:05:48.124 --> 01:05:50.560 have proven not to be as successful.

NOTE Confidence: 0.645885562857143

01:05:50.560 --> 01:05:54.452 And but you can most definitely look

NOTE Confidence: 0.645885562857143

01:05:54.452 --> 01:05:58.957 up some of his work on Pub Med and

NOTE Confidence: 0.645885562857143

01:05:58.960 --> 01:06:01.080 and I think I'll leave it at that,

NOTE Confidence: 0.645885562857143

01:06:01.080 --> 01:06:03.600 but I do not have anything else to add.

NOTE Confidence: 0.56885285

01:06:04.280 --> 01:06:05.318 What Jerry said,

NOTE Confidence: 0.833750174615385

01:06:06.960 --> 01:06:08.360 Jared, do you want to speak to

NOTE Confidence: 0.833750174615385

01:06:08.360 --> 01:06:10.080 that at all briefly or not today?

NOTE Confidence: 0.8471053

01:06:12.160 --> 01:06:16.400 Thank you for singling me out just really

NOTE Confidence: 0.911767558333333

01:06:16.400 --> 01:06:20.115 briefly. I think it is really it's

NOTE Confidence: 0.911767558333333

01:06:20.115 --> 01:06:23.160 already so hard to run a clinical

NOTE Confidence: 0.911767558333333

01:06:23.160 --> 01:06:26.000 trial navigating all of the regulatory

NOTE Confidence: 0.911767558333333

01:06:26.000 --> 01:06:28.200 demands and everything on top of what

NOTE Confidence: 0.911767558333333

01:06:28.200 --> 01:06:30.260 needs to be done in the study itself

NOTE Confidence: 0.911767558333333

01:06:30.260 --> 01:06:32.620 that it requires tremendous effort.

NOTE Confidence: 0.911767558333333

01:06:32.620 --> 01:06:35.740 On top of all of that to conduct

NOTE Confidence: 0.911767558333333

01:06:35.740 --> 01:06:37.280 really intensive outreach

NOTE Confidence: 0.911767558333333

01:06:37.280 --> 01:06:40.542 efforts to communities of color,

NOTE Confidence: 0.911767558333333

01:06:40.542 --> 01:06:42.186 particularly the black and

NOTE Confidence: 0.911767558333333

01:06:42.186 --> 01:06:44.320 brown population in New Haven.

NOTE Confidence: 0.781737085

01:06:46.720 --> 01:06:48.190 You know, just really pointing to

NOTE Confidence: 0.781737085

01:06:48.190 --> 01:06:50.000 the cultural ambassadors program.

NOTE Confidence: 0.781737085

01:06:50.000 --> 01:06:54.000 We've been in touch with them,

NOTE Confidence: 0.781737085

01:06:54.000 --> 01:06:55.916 Jordan, Jordan Slowshower is also

NOTE Confidence: 0.781737085

01:06:55.916 --> 01:06:57.862 on this call and we've had a

NOTE Confidence: 0.781737085

01:06:57.862 --> 01:06:59.959 couple of conversations with them.

NOTE Confidence: 0.781737085

01:06:59.960 --> 01:07:01.120 They've never done anything

NOTE Confidence: 0.781737085

01:07:01.120 --> 01:07:02.560 psychedelic related.

NOTE Confidence: 0.781737085

01:07:02.560 --> 01:07:04.056 So it's all just going to

NOTE Confidence: 0.781737085
01:07:04.056 --> 01:07:05.996 be new frontiers for us.
NOTE Confidence: 0.781737085
01:07:06.000 --> 01:07:08.198 It's worth the effort,
NOTE Confidence: 0.781737085
01:07:08.200 --> 01:07:11.373 but have haven't worked on an
NOTE Confidence: 0.781737085
01:07:11.373 --> 01:07:14.438 MDMA trial looking to exclusively
NOTE Confidence: 0.781737085
01:07:14.440 --> 01:07:16.546 recruit by POC participants.
NOTE Confidence: 0.781737085
01:07:16.546 --> 01:07:18.678 I can say that
NOTE Confidence: 0.8552270525
01:07:19.440 --> 01:07:21.840 it it really is a very
NOTE Confidence: 0.8552270525
01:07:21.840 --> 01:07:23.078 tremendous effort that is
NOTE Confidence: 0.60940523
01:07:23.080 --> 01:07:24.559 required to to
NOTE Confidence: 0.952292257142857
01:07:24.560 --> 01:07:27.710 even bring people into the consent
NOTE Confidence: 0.952292257142857
01:07:27.710 --> 01:07:29.360 call to run through screening
NOTE Confidence: 0.952292257142857
01:07:29.360 --> 01:07:30.690 procedures with them because
NOTE Confidence: 0.952292257142857
01:07:30.690 --> 01:07:32.440 there's just so many considerations
NOTE Confidence: 0.929026266666667
01:07:32.440 --> 01:07:33.358 that people have
NOTE Confidence: 0.959145885
01:07:33.880 --> 01:07:36.640 about completing the trial.
NOTE Confidence: 0.959145885

01:07:36.640 --> 01:07:38.760 So all of these are worthwhile issues
NOTE Confidence: 0.9759234

01:07:38.760 --> 01:07:42.160 to think about that I think you know
NOTE Confidence: 0.841209316666667

01:07:42.200 --> 01:07:44.396 it. If the team is aligned
NOTE Confidence: 0.886591456666667

01:07:44.960 --> 01:07:45.758 in making this
NOTE Confidence: 0.971988522

01:07:45.760 --> 01:07:48.728 work, I think we can move the needle
NOTE Confidence: 0.971988522

01:07:48.728 --> 01:07:50.156 on making sure that we have more
NOTE Confidence: 0.971988522

01:07:50.160 --> 01:07:51.640 diverse samples moving forward.
NOTE Confidence: 0.6524430525

01:07:54.120 --> 01:07:56.739 Jerry, I had a Jerry and Ben I had
NOTE Confidence: 0.6524430525

01:07:56.739 --> 01:07:58.880 a question about generalizability.
NOTE Confidence: 0.87500643

01:07:58.880 --> 01:07:59.712 So all these patients
NOTE Confidence: 0.87500643

01:07:59.712 --> 01:08:01.160 were off antidepressants.
NOTE Confidence: 0.937361836

01:08:01.600 --> 01:08:04.660 How compelling is the data that
NOTE Confidence: 0.937361836

01:08:04.660 --> 01:08:06.430 antidepressants would interfere
NOTE Confidence: 0.937361836

01:08:06.430 --> 01:08:08.320 with the effects of these drugs?
NOTE Confidence: 0.937361836

01:08:08.320 --> 01:08:11.296 And in future trials is the
NOTE Confidence: 0.937361836

01:08:11.296 --> 01:08:13.000 FDA going to, you know,

NOTE Confidence: 0.949613468

01:08:13.400 --> 01:08:17.640 allow patients who are taking antidepressants

NOTE Confidence: 0.949613468

01:08:17.640 --> 01:08:20.226 participate in studies with with the

NOTE Confidence: 0.949613468

01:08:20.226 --> 01:08:21.680 serotonergic psych psychedelics?

NOTE Confidence: 0.647898276666667

01:08:23.320 --> 01:08:23.719 It's a really

NOTE Confidence: 0.919367392

01:08:23.720 --> 01:08:27.363 good question and I think there is some

NOTE Confidence: 0.919367392

01:08:27.363 --> 01:08:30.750 work being done in this area to explore

NOTE Confidence: 0.919367392

01:08:30.750 --> 01:08:35.320 the extent to which the presence of

NOTE Confidence: 0.919367392

01:08:35.320 --> 01:08:38.300 antidepressants at the time of dosing

NOTE Confidence: 0.919367392

01:08:38.300 --> 01:08:42.834 versus a six week follow up versus 5

NOTE Confidence: 0.919367392

01:08:42.834 --> 01:08:46.224 half life paper is impacting outcome.

NOTE Confidence: 0.919367392

01:08:46.224 --> 01:08:49.560 I think in this particular study

NOTE Confidence: 0.919367392

01:08:49.658 --> 01:08:53.852 Particip I many participants did not.

NOTE Confidence: 0.919367392

01:08:53.852 --> 01:08:55.236 We're not an antidepressants.

NOTE Confidence: 0.919367392

01:08:55.240 --> 01:08:57.354 And the ones that were on antidepressants,

NOTE Confidence: 0.919367392

01:08:57.360 --> 01:09:00.741 they were actually tapered and they had

NOTE Confidence: 0.919367392

01:09:00.741 --> 01:09:03.560 to have been off of antidepressants,
NOTE Confidence: 0.919367392

01:09:03.560 --> 01:09:08.600 fully tapered for six weeks prior to
NOTE Confidence: 0.670358304

01:09:10.880 --> 01:09:12.032 baseline rating.
NOTE Confidence: 0.670358304

01:09:12.032 --> 01:09:14.510 I believe it's five half lives.
NOTE Confidence: 0.670358304

01:09:14.510 --> 01:09:16.480 Five half lives, five half lives.
NOTE Confidence: 0.670358304

01:09:16.480 --> 01:09:19.626 So, yeah, so there's basic and
NOTE Confidence: 0.670358304

01:09:19.626 --> 01:09:22.356 recently there's been a paper
NOTE Confidence: 0.670358304

01:09:22.356 --> 01:09:25.186 that has and more specifically,
NOTE Confidence: 0.670358304

01:09:25.186 --> 01:09:28.724 there is a a company that's received,
NOTE Confidence: 0.670358304

01:09:28.724 --> 01:09:30.276 I believe mind Mendapatent,
NOTE Confidence: 0.670358304

01:09:30.280 --> 01:09:32.160 where they actually are used
NOTE Confidence: 0.670358304

01:09:32.160 --> 01:09:36.850 using psilocybin with Lexapro to
NOTE Confidence: 0.670358304

01:09:36.850 --> 01:09:39.100 enhance the effect of psilocybin
NOTE Confidence: 0.670358304

01:09:39.100 --> 01:09:41.600 by minimizing its side effects.
NOTE Confidence: 0.670358304

01:09:41.600 --> 01:09:43.520 And so there is some preliminary
NOTE Confidence: 0.670358304

01:09:43.520 --> 01:09:45.193 data showing that the presence

NOTE Confidence: 0.670358304

01:09:45.193 --> 01:09:48.505 of SSRI in fact might actually

NOTE Confidence: 0.670358304

01:09:48.505 --> 01:09:52.089 enhance the effect of psilocybin by

NOTE Confidence: 0.670358304

01:09:52.089 --> 01:09:54.760 mitigating some of the side effects.

NOTE Confidence: 0.670358304

01:09:54.760 --> 01:09:56.800 But that is just very clear.

NOTE Confidence: 0.670358304

01:09:56.800 --> 01:09:59.400 But it does go back to your question,

NOTE Confidence: 0.670358304

01:09:59.400 --> 01:10:02.324 Doctor Sousa and that is we do

NOTE Confidence: 0.670358304

01:10:02.324 --> 01:10:04.256 not to say that actually patients

NOTE Confidence: 0.670358304

01:10:04.256 --> 01:10:06.200 should be off of SSR is to

NOTE Confidence: 0.670358304

01:10:06.200 --> 01:10:07.320 participate in these studies.

NOTE Confidence: 0.670358304

01:10:07.320 --> 01:10:13.072 I think that is not any more supported.

NOTE Confidence: 0.670358304

01:10:13.072 --> 01:10:15.358 And it's also the COMPASS

NOTE Confidence: 0.670358304

01:10:15.360 --> 01:10:17.000 and the COMPASS open label.

NOTE Confidence: 0.670358304

01:10:17.000 --> 01:10:17.630 That's right.

NOTE Confidence: 0.670358304

01:10:17.630 --> 01:10:19.960 So there's been a couple of studies and

NOTE Confidence: 0.758118894

01:10:20.440 --> 01:10:21.260 there are two different.

NOTE Confidence: 0.758118894

01:10:21.260 --> 01:10:22.982 I mean you said that that having Lexapro

NOTE Confidence: 0.758118894

01:10:22.982 --> 01:10:24.758 on board at the time may not interfere,

NOTE Confidence: 0.758118894

01:10:24.760 --> 01:10:26.794 may actually enhance, but that's a

NOTE Confidence: 0.758118894

01:10:26.794 --> 01:10:28.640 very different question from chronic.

NOTE Confidence: 0.758118894

01:10:28.640 --> 01:10:29.835 There's evidence from animal studies

NOTE Confidence: 0.758118894

01:10:29.835 --> 01:10:31.926 and I think this is where some of the

NOTE Confidence: 0.758118894

01:10:31.926 --> 01:10:33.465 thinking came from that you need to

NOTE Confidence: 0.758118894

01:10:33.465 --> 01:10:34.956 be off is edited from animal studies.

NOTE Confidence: 0.758118894

01:10:34.960 --> 01:10:37.104 The chronic SSRI use leads to a down

NOTE Confidence: 0.758118894

01:10:37.104 --> 01:10:38.719 regulation of the two a receptor.

NOTE Confidence: 0.758118894

01:10:38.720 --> 01:10:41.520 That's right. And so it may,

NOTE Confidence: 0.758118894

01:10:41.520 --> 01:10:43.440 which is a different question from

NOTE Confidence: 0.758118894

01:10:43.440 --> 01:10:44.868 what's the effect of having an SSRI

NOTE Confidence: 0.758118894

01:10:44.868 --> 01:10:46.360 on board at the time of dosing.

NOTE Confidence: 0.758118894

01:10:46.360 --> 01:10:48.208 Yeah. So the five half lives is

NOTE Confidence: 0.758118894

01:10:48.208 --> 01:10:50.034 calculated to make sure it's out of

NOTE Confidence: 0.758118894

01:10:50.034 --> 01:10:51.780 the system so that there's no SSRI

NOTE Confidence: 0.758118894

01:10:51.780 --> 01:10:53.040 on board at the time of dosing.

NOTE Confidence: 0.758118894

01:10:53.040 --> 01:10:54.714 But that may not be enough time to reverse

NOTE Confidence: 0.758118894

01:10:54.714 --> 01:10:56.517 the down regulation of the two A receptors.

NOTE Confidence: 0.758118894

01:10:56.520 --> 01:10:57.360 So two different.

NOTE Confidence: 0.758118894

01:10:57.360 --> 01:10:59.040 And then the third mechanistic concern

NOTE Confidence: 0.758118894

01:10:59.040 --> 01:11:00.455 I've heard is serotonin syndrome

NOTE Confidence: 0.758118894

01:11:00.455 --> 01:11:02.081 as a risk of serotonin syndrome

NOTE Confidence: 0.758118894

01:11:02.129 --> 01:11:03.802 that I think is overblown and I've

NOTE Confidence: 0.758118894

01:11:03.802 --> 01:11:05.660 seen that's right to suggest that,

NOTE Confidence: 0.758118894

01:11:05.660 --> 01:11:07.160 that's a realistic concern.

NOTE Confidence: 0.758118894

01:11:07.160 --> 01:11:08.680 But those are kind of the three different.

NOTE Confidence: 0.611833374

01:11:08.720 --> 01:11:10.600 Yeah, I I mean, I

NOTE Confidence: 0.844377282

01:11:10.600 --> 01:11:13.040 think this is a major,

NOTE Confidence: 0.844377282

01:11:13.040 --> 01:11:14.760 I mean this is going to be a big issue

NOTE Confidence: 0.844377282

01:11:14.806 --> 01:11:16.112 moving forward in implementation.
NOTE Confidence: 0.844377282

01:11:16.112 --> 01:11:19.440 This is going to be a big deal exactly
NOTE Confidence: 0.844377282

01:11:19.440 --> 01:11:21.720 in generalizability too. Yeah.
NOTE Confidence: 0.844377282

01:11:21.720 --> 01:11:24.052 Yeah, yeah, You know, I think this,
NOTE Confidence: 0.844377282

01:11:24.052 --> 01:11:27.790 I mean my take on it is, you know,
NOTE Confidence: 0.844377282

01:11:27.790 --> 01:11:29.995 it's really the lore in the fields.
NOTE Confidence: 0.844377282

01:11:30.000 --> 01:11:31.152 I I think the studies will
NOTE Confidence: 0.844377282

01:11:31.152 --> 01:11:32.479 have to be done to test it.
NOTE Confidence: 0.844377282

01:11:32.480 --> 01:11:35.312 There is the preclinical data that
NOTE Confidence: 0.844377282

01:11:35.312 --> 01:11:37.959 suggests that maybe but and now there's
NOTE Confidence: 0.844377282

01:11:37.960 --> 01:11:39.164 exactly companies are in an open label
NOTE Confidence: 0.844377282

01:11:39.164 --> 01:11:40.477 city where it didn't seem to make and
NOTE Confidence: 0.64826884625

01:11:40.560 --> 01:11:42.320 if it is a 2A down regulation issue,
NOTE Confidence: 0.64826884625

01:11:42.320 --> 01:11:44.110 it may just be an issue of dose, maybe you
NOTE Confidence: 0.64826884625

01:11:44.110 --> 01:11:46.040 need a slightly higher dose and that's
NOTE Confidence: 0.521396071

01:11:47.200 --> 01:11:50.688 so the pre the preclinical data on down

NOTE Confidence: 0.521396071

01:11:50.688 --> 01:11:53.480 regulation of 2A receptors preclinically

NOTE Confidence: 0.521396071

01:11:53.480 --> 01:11:55.480 is actually really quite spotty.

NOTE Confidence: 0.521396071

01:11:55.480 --> 01:11:58.740 There's really and is down regulation

NOTE Confidence: 0.521396071

01:11:58.740 --> 01:12:01.290 really meaningful or are there alter

NOTE Confidence: 0.521396071

01:12:01.369 --> 01:12:04.399 other alterations in 2A receptor function?

NOTE Confidence: 0.521396071

01:12:04.400 --> 01:12:06.650 I looked at, I looked at it once in a

NOTE Confidence: 0.521396071

01:12:06.650 --> 01:12:08.692 very small population of that's and

NOTE Confidence: 0.521396071

01:12:08.692 --> 01:12:10.990 I saw absolutely nothing with them

NOTE Confidence: 0.521396071

01:12:11.060 --> 01:12:14.320 getting reasonable exposures of an SSRI.

NOTE Confidence: 0.521396071

01:12:14.320 --> 01:12:16.360 So it's, it's questionable I think.

NOTE Confidence: 0.77581198

01:12:16.560 --> 01:12:17.436 Yeah. Thank you, Jerry.

NOTE Confidence: 0.77581198

01:12:17.436 --> 01:12:18.750 It's like it's like 2 papers

NOTE Confidence: 0.77581198

01:12:18.799 --> 01:12:20.080 from 20 years ago. Yeah, that's

NOTE Confidence: 0.73265141375

01:12:21.320 --> 01:12:23.480 right. I mean this is something I think

NOTE Confidence: 0.73265141375

01:12:23.480 --> 01:12:25.626 that feels it's like all of all of,

NOTE Confidence: 0.73265141375

01:12:25.626 --> 01:12:26.718 you know, biomedical science,
NOTE Confidence: 0.73265141375

01:12:26.720 --> 01:12:28.560 but especially in this area,
NOTE Confidence: 0.73265141375

01:12:28.560 --> 01:12:30.954 there's a lot of war that's hard to overcome.
NOTE Confidence: 0.73265141375

01:12:30.960 --> 01:12:33.148 I mean this is things, you know,
NOTE Confidence: 0.73265141375

01:12:33.148 --> 01:12:35.512 people have been doing this since
NOTE Confidence: 0.73265141375

01:12:35.512 --> 01:12:38.712 the 1950s and to try to to try to
NOTE Confidence: 0.73265141375

01:12:38.712 --> 01:12:40.936 change that with data requires data.
NOTE Confidence: 0.73265141375

01:12:40.936 --> 01:12:44.200 So, and I I do respect that companies
NOTE Confidence: 0.73265141375

01:12:44.284 --> 01:12:47.374 being nervous about jumping right into
NOTE Confidence: 0.73265141375

01:12:47.374 --> 01:12:50.480 those studies first, first prove,
NOTE Confidence: 0.73265141375

01:12:50.480 --> 01:12:52.280 you know, proof of concept under those
NOTE Confidence: 0.73265141375

01:12:52.280 --> 01:12:53.880 conditions and then going forward.
NOTE Confidence: 0.73265141375

01:12:53.880 --> 01:12:55.750 But I agree with you.
NOTE Confidence: 0.73265141375

01:12:55.750 --> 01:12:57.160 Yeah, I think and
NOTE Confidence: 0.80547798

01:12:58.960 --> 01:13:03.400 not too dissimilar from the REMS that
NOTE Confidence: 0.80547798

01:13:03.400 --> 01:13:06.396 was eventually was attached to the label.

NOTE Confidence: 0.80547798

01:13:06.400 --> 01:13:08.308 As for Avada, where if patients

NOTE Confidence: 0.80547798

01:13:08.308 --> 01:13:11.443 did not at the time of dosing with

NOTE Confidence: 0.80547798

01:13:11.443 --> 01:13:13.673 Astravada were not on antidepressants,

NOTE Confidence: 0.80547798

01:13:13.680 --> 01:13:17.238 they had to start an antidepressant.

NOTE Confidence: 0.80547798

01:13:17.240 --> 01:13:20.920 And I would not be surprised to that if

NOTE Confidence: 0.80547798

01:13:20.920 --> 01:13:24.404 the data begins to show that the Pres

NOTE Confidence: 0.80547798

01:13:24.404 --> 01:13:26.400 adding an antidepressant before dosing,

NOTE Confidence: 0.80547798

01:13:26.400 --> 01:13:29.871 whether it is for treatment issues

NOTE Confidence: 0.80547798

01:13:29.871 --> 01:13:34.008 or to have something on board when

NOTE Confidence: 0.80547798

01:13:34.008 --> 01:13:36.785 psilocybin wears off or to really

NOTE Confidence: 0.80547798

01:13:36.785 --> 01:13:39.155 just minimize some of the adverse

NOTE Confidence: 0.80547798

01:13:39.155 --> 01:13:41.640 events associated with psilocybin.

NOTE Confidence: 0.80547798

01:13:41.640 --> 01:13:44.144 I would not be surprised that we will

NOTE Confidence: 0.80547798

01:13:44.144 --> 01:13:46.479 be seeing more of those studies in

NOTE Confidence: 0.80547798

01:13:46.479 --> 01:13:48.922 the next couple of years to really

NOTE Confidence: 0.80547798

01:13:48.922 --> 01:13:50.772 either optimize the treatment or
NOTE Confidence: 0.80547798

01:13:50.772 --> 01:13:54.354 sort of provide an antidepressant
NOTE Confidence: 0.80547798

01:13:54.354 --> 01:13:57.198 on board as a safety mechanism.
NOTE Confidence: 0.882593879130435

01:14:00.680 --> 01:14:02.010 I do want to put in a plug to the
NOTE Confidence: 0.882593879130435

01:14:02.053 --> 01:14:03.183 issue of the regulatory framework
NOTE Confidence: 0.882593879130435

01:14:03.183 --> 01:14:04.880 and what this is going to look like.
NOTE Confidence: 0.882593879130435

01:14:04.880 --> 01:14:06.672 Will there be a Rams and so forth
NOTE Confidence: 0.882593879130435

01:14:06.672 --> 01:14:08.373 has come up several times at
NOTE Confidence: 0.882593879130435

01:14:08.373 --> 01:14:10.038 this seminar time in February.
NOTE Confidence: 0.882593879130435

01:14:10.040 --> 01:14:12.400 We're going to have a panel of folks
NOTE Confidence: 0.882593879130435

01:14:12.400 --> 01:14:13.969 discussing precisely that issue including
NOTE Confidence: 0.882593879130435

01:14:13.969 --> 01:14:16.111 two folks whose names I don't remember
NOTE Confidence: 0.882593879130435

01:14:16.166 --> 01:14:18.122 off the top of my head, Sean Bulan.
NOTE Confidence: 0.882593879130435

01:14:18.122 --> 01:14:20.027 But leaders from the Department
NOTE Confidence: 0.882593879130435

01:14:20.027 --> 01:14:22.239 of Health and Human Services,
NOTE Confidence: 0.882593879130435

01:14:22.240 --> 01:14:24.103 who are the people who are going to be

NOTE Confidence: 0.882593879130435
01:14:24.103 --> 01:14:25.756 running the process of developing this,
NOTE Confidence: 0.882593879130435
01:14:25.760 --> 01:14:28.256 as well as Jerry and a couple other
NOTE Confidence: 0.882593879130435
01:14:28.256 --> 01:14:30.146 discussions from from a couple
NOTE Confidence: 0.882593879130435
01:14:30.146 --> 01:14:32.640 universities across the country who are
NOTE Confidence: 0.882593879130435
01:14:32.640 --> 01:14:34.640 thinking deeply about these issues.
NOTE Confidence: 0.882593879130435
01:14:34.640 --> 01:14:36.524 There's a call for commentary open
NOTE Confidence: 0.882593879130435
01:14:36.524 --> 01:14:38.908 right now from the Department of Health
NOTE Confidence: 0.882593879130435
01:14:38.908 --> 01:14:41.038 and Human Services asking for people
NOTE Confidence: 0.882593879130435
01:14:41.038 --> 01:14:43.358 in the field to put in information,
NOTE Confidence: 0.882593879130435
01:14:43.360 --> 01:14:45.760 questions and concerns on these issues.
NOTE Confidence: 0.882593879130435
01:14:45.760 --> 01:14:46.980 And we've timed that round
NOTE Confidence: 0.882593879130435
01:14:46.980 --> 01:14:47.956 table discussion in February.
NOTE Confidence: 0.882593879130435
01:14:47.960 --> 01:14:49.093 So it'll be shortly before
NOTE Confidence: 0.882593879130435
01:14:49.093 --> 01:14:50.788 the close of that call.
NOTE Confidence: 0.882593879130435
01:14:50.788 --> 01:14:53.000 So there will be discussion of the,
NOTE Confidence: 0.882593879130435

01:14:53.000 --> 01:14:54.519 you know the the in the early.
NOTE Confidence: 0.882593879130435

01:14:54.520 --> 01:14:56.506 I think it's the first public
NOTE Confidence: 0.882593879130435

01:14:56.506 --> 01:14:57.904 synthesis of the information that
NOTE Confidence: 0.882593879130435

01:14:57.904 --> 01:15:00.159 has come in over the next two months.
NOTE Confidence: 0.882593879130435

01:15:00.160 --> 01:15:02.848 And then after that there will be still
NOTE Confidence: 0.882593879130435

01:15:02.848 --> 01:15:05.648 another week I think to where one can can,
NOTE Confidence: 0.882593879130435

01:15:05.648 --> 01:15:07.184 where where folks can after hearing
NOTE Confidence: 0.882593879130435

01:15:07.184 --> 01:15:08.680 that discussion continue to put
NOTE Confidence: 0.882593879130435

01:15:08.680 --> 01:15:10.160 in information on that call.
NOTE Confidence: 0.882593879130435

01:15:10.160 --> 01:15:11.600 So I think it'll be and we're going
NOTE Confidence: 0.882593879130435

01:15:11.600 --> 01:15:12.599 to advertise this nationally.
NOTE Confidence: 0.882593879130435

01:15:12.600 --> 01:15:14.822 I think it's going to be an important event.
NOTE Confidence: 0.882593879130435

01:15:14.822 --> 01:15:16.920 So I just want to put in the plug for that.
NOTE Confidence: 0.882593879130435

01:15:16.920 --> 01:15:18.355 It'll be at the usual time the,
NOTE Confidence: 0.882593879130435

01:15:18.360 --> 01:15:21.797 you know, the the third Friday in
NOTE Confidence: 0.882593879130435

01:15:21.797 --> 01:15:24.120 February publicity going out soon.

NOTE Confidence: 0.882593879130435
01:15:24.120 --> 01:15:24.600 OK, That's right.
NOTE Confidence: 0.882593879130435
01:15:25.120 --> 01:15:26.640 Well,
NOTE Confidence: 0.728572491666667
01:15:26.640 --> 01:15:29.464 OK, just all right.
NOTE Confidence: 0.728572491666667
01:15:29.464 --> 01:15:32.839 Any other any other questions?
NOTE Confidence: 0.893077625714286
01:15:34.560 --> 01:15:36.648 What's on the horizon for the
NOTE Confidence: 0.893077625714286
01:15:36.648 --> 01:15:42.480 Yasona program? So. So I think
NOTE Confidence: 0.839974988947368
01:15:44.880 --> 01:15:46.856 clearly it is a plan for a phase
NOTE Confidence: 0.839974988947368
01:15:46.856 --> 01:15:48.365 three study that hopefully will
NOTE Confidence: 0.839974988947368
01:15:48.365 --> 01:15:50.273 be underway in the near future.
NOTE Confidence: 0.8338018225
01:15:53.520 --> 01:15:56.480 So the the goal is to move forward. Yeah.
NOTE Confidence: 0.6366484
01:15:59.000 --> 01:16:01.626 OK. Chris, anything else?
NOTE Confidence: 0.6366484
01:16:01.626 --> 01:16:02.520 Anything else? Yeah.
NOTE Confidence: 0.677138975
01:16:02.800 --> 01:16:03.360 Thank you both. Thank
NOTE Confidence: 0.677138975
01:16:03.360 --> 01:16:06.000 you all. Thank you. Bye.