WEBVTT

NOTE duration: "01:13:49"

NOTE language:en-us

NOTE Confidence: 0.86898927

 $00:00:00.000 \longrightarrow 00:00:02.403$ We have a lot to go through today and

NOTE Confidence: 0.86898927

 $00{:}02.403 \dashrightarrow 00{:}05.019$ some exciting science to hear from some

NOTE Confidence: 0.86898927

 $00:00:05.019 \longrightarrow 00:00:07.310$ wonderful trainees in our department.

NOTE Confidence: 0.86898927

00:00:07.310 --> 00:00:09.998 I want to welcome you to grand rounds

NOTE Confidence: 0.86898927

 $00:00:09.998 \dashrightarrow 00:00:13.862$ into the 2021 Seymour Lustman Memorial

NOTE Confidence: 0.86898927

00:00:13.862 --> 00:00:17.054 Awards in psychiatric research.

NOTE Confidence: 0.86898927

 $00{:}00{:}17.060 \longrightarrow 00{:}00{:}19.681$ This I have to say is one of my favorite

NOTE Confidence: 0.86898927

00:00:19.681 --> 00:00:22.051 days in the life of our department

NOTE Confidence: 0.86898927

 $00:00:22.051 \longrightarrow 00:00:24.265$ and in our grand rounds series.

NOTE Confidence: 0.86898927

 $00:00:24.270 \longrightarrow 00:00:26.475$ As we acknowledge the work of trainees

NOTE Confidence: 0.86898927

 $00{:}00{:}26.475 \dashrightarrow 00{:}00{:}28.473$ in our department and the research

NOTE Confidence: 0.86898927

 $00:00:28.473 \longrightarrow 00:00:30.286$ that they've done, and we also

NOTE Confidence: 0.86898927

 $00:00:30.286 \longrightarrow 00:00:32.389$ celebrate the legacy of our department,

NOTE Confidence: 0.86898927

 $00:00:32.389 \longrightarrow 00:00:35.280$ the Lastman Awards has been his allotment.

00:00:35.280 --> 00:00:37.130 Orders been given since 1973,

NOTE Confidence: 0.86898927

 $00{:}00{:}37.130 \dashrightarrow 00{:}00{:}38.880$ and if you look over the list

NOTE Confidence: 0.86898927

 $00:00:38.880 \longrightarrow 00:00:39.630$ of past winners.

NOTE Confidence: 0.86898927

 $00:00:39.630 \longrightarrow 00:00:42.045$ Who are in an engraved plaque outside

NOTE Confidence: 0.86898927

 $00:00:42.045 \longrightarrow 00:00:43.869$ the the auditorium where unfortunately

NOTE Confidence: 0.86898927

 $00:00:43.869 \longrightarrow 00:00:46.410$ not able to meet in person today.

NOTE Confidence: 0.86898927

00:00:46.410 --> 00:00:49.122 It's really an extraordinary list and we have

NOTE Confidence: 0.86898927

 $00:00:49.122 \longrightarrow 00:00:52.067$ some new people to add to that plaque today.

NOTE Confidence: 0.86898927

 $00:00:52.070 \longrightarrow 00:00:54.002$ I want to thank the

NOTE Confidence: 0.86898927

00:00:54.002 --> 00:00:56.530 Lustman family Susan Katz,

NOTE Confidence: 0.86898927

00:00:56.530 --> 00:00:57.354 Jeffrey Lastman,

NOTE Confidence: 0.86898927

 $00:00:57.354 \longrightarrow 00:00:59.414$ Seymour Lessmann's children and the

NOTE Confidence: 0.86898927

00:00:59.414 --> 00:01:01.898 less than Family Foundation who have

NOTE Confidence: 0.86898927

00:01:01.898 --> 00:01:03.474 generously provided support for

NOTE Confidence: 0.86898927

 $00:01:03.474 \longrightarrow 00:01:06.234$ this award for decades now and have

 $00:01:06.234 \longrightarrow 00:01:08.124$ therefore contributed to an enabled.

NOTE Confidence: 0.86898927

 $00{:}01{:}08.130 \dashrightarrow 00{:}01{:}12.892$ The history that I referred to.

NOTE Confidence: 0.86898927

 $00:01:12.892 \dashrightarrow 00:01:16.732$ So see more lost men. Make me you you.

NOTE Confidence: 0.86898927

00:01:16.732 --> 00:01:17.864 You might not feel,

NOTE Confidence: 0.86898927

 $00:01:17.870 \longrightarrow 00:01:18.854$ especially if this is the first

NOTE Confidence: 0.86898927

 $00:01:18.854 \longrightarrow 00:01:19.949$ time you've come to the last

NOTE Confidence: 0.86898927

 $00:01:19.949 \longrightarrow 00:01:20.549$ minute award ceremony.

NOTE Confidence: 0.86898927

00:01:20.550 --> 00:01:23.392 You may not know the history of

NOTE Confidence: 0.86898927

 $00{:}01{:}23.392 \to 00{:}01{:}25.378$ Seymour last minum less been served

NOTE Confidence: 0.86898927

 $00:01:25.380 \longrightarrow 00:01:27.256$ in the army in World War Two,

NOTE Confidence: 0.86898927

 $00{:}01{:}27.260 \dashrightarrow 00{:}01{:}29.423$ he obtained his PhD in psychology at

NOTE Confidence: 0.86898927

00:01:29.423 --> 00:01:31.336 the University of Chicago and then

NOTE Confidence: 0.86898927

 $00{:}01{:}31.336 \dashrightarrow 00{:}01{:}33.499$ his MD at the University of Illinois.

NOTE Confidence: 0.86898927

 $00:01:33.500 \longrightarrow 00:01:36.188$ Before coming to Yale for his

NOTE Confidence: 0.86898927

 $00:01:36.188 \longrightarrow 00:01:38.060$ psychiatry residency in 1955.

NOTE Confidence: 0.86898927

 $00:01:38.060 \longrightarrow 00:01:40.140$ During his PhD studies,

 $00:01:40.140 \longrightarrow 00:01:41.252$ before coming to Yale,

NOTE Confidence: 0.86898927

 $00{:}01{:}41.252 \dashrightarrow 00{:}01{:}42.920$ he became very interested in the

NOTE Confidence: 0.86898927

00:01:42.977 --> 00:01:44.697 question of nature versus nurture.

NOTE Confidence: 0.86898927

 $00:01:44.700 \longrightarrow 00:01:46.728$ In the words of the day,

NOTE Confidence: 0.86898927

 $00:01:46.730 \longrightarrow 00:01:51.315$ we might call it environment versus genetics.

NOTE Confidence: 0.86898927

 $00:01:51.320 \longrightarrow 00:01:54.050$ And he he continued with that interest

NOTE Confidence: 0.86898927

 $00:01:54.050 \longrightarrow 00:01:56.550$ throughout his clinical and research career.

NOTE Confidence: 0.86898927

00:01:56.550 --> 00:01:58.128 After completing his psychiatry

NOTE Confidence: 0.86898927

00:01:58.128 --> 00:01:59.948 residency in his child Fellowship,

NOTE Confidence: 0.86898927

 $00:01:59.950 \longrightarrow 00:02:03.207$ he joined the faculty in 1962 and a

NOTE Confidence: 0.86898927

 $00:02:03.207 \longrightarrow 00:02:05.440$ grand total of two years later was

NOTE Confidence: 0.86898927

 $00:02:05.513 \longrightarrow 00:02:08.278$ promoted to the rank of full professor,

NOTE Confidence: 0.86898927

 $00{:}02{:}08.280 \dashrightarrow 00{:}02{:}10.020$ which is a rather impressive

NOTE Confidence: 0.86898927

00:02:10.020 --> 00:02:11.680 trajectory that speaks, I think,

NOTE Confidence: 0.86898927

00:02:11.680 --> 00:02:13.205 to how evidenced his his,

 $00:02:13.210 \longrightarrow 00:02:15.315$ his excellence in his contributions

NOTE Confidence: 0.86898927

 $00:02:15.315 \longrightarrow 00:02:16.999$ to the department were.

NOTE Confidence: 0.86898927

 $00:02:17.000 \longrightarrow 00:02:18.540$ He was a dedicated teacher,

NOTE Confidence: 0.86898927

 $00:02:18.540 \longrightarrow 00:02:21.053$ gifted clinician and a very

NOTE Confidence: 0.86898927

 $00{:}02{:}21.053 \dashrightarrow 00{:}02{:}22.617$ careful and creative scientist.

NOTE Confidence: 0.86898927

 $00:02:22.620 \longrightarrow 00:02:25.108$ One thing he did.

NOTE Confidence: 0.86898927

 $00{:}02{:}25.110 \dashrightarrow 00{:}02{:}27.862$ That that story I've heard told many times

NOTE Confidence: 0.86898927

 $00:02:27.862 \longrightarrow 00:02:30.789$ this was after the polio was on the wane,

NOTE Confidence: 0.86898927

 $00{:}02{:}30.790 \longrightarrow 00{:}02{:}32.630$ and there were all of these iron lungs.

NOTE Confidence: 0.86898927

00:02:32.630 --> 00:02:35.108 These big breathing machines that were used,

NOTE Confidence: 0.86898927

 $00{:}02{:}35.110 \longrightarrow 00{:}02{:}37.396$ but it had been used to keep people alive

NOTE Confidence: 0.86898927

 $00:02:37.396 \longrightarrow 00:02:39.429$ when they had severe cases of polio,

NOTE Confidence: 0.86898927

00:02:39.430 --> 00:02:41.150 and they weren't needed anymore,

NOTE Confidence: 0.86898927

 $00{:}02{:}41.150 \dashrightarrow 00{:}02{:}43.195$ and less man repurpose them into

NOTE Confidence: 0.86898927

00:02:43.195 --> 00:02:45.748 basically laboratories to study to study

NOTE Confidence: 0.86898927

 $00:02:45.748 \longrightarrow 00:02:47.968$ children in a controlled environment,

 $00:02:47.970 \longrightarrow 00:02:50.360$ and shows both creativity and

NOTE Confidence: 0.86898927

 $00:02:50.360 \longrightarrow 00:02:53.303$ dedication to to advancing the

NOTE Confidence: 0.86898927

 $00{:}02{:}53.303 \dashrightarrow 00{:}02{:}55.286$ understanding of children.

NOTE Confidence: 0.86898927

 $00:02:55.290 \longrightarrow 00:02:56.220$ And child development.

NOTE Confidence: 0.86898927

 $00:02:56.220 \longrightarrow 00:02:58.810$ And that became the touch stone of his career.

NOTE Confidence: 0.86898927

 $00:02:58.810 \longrightarrow 00:03:00.844$ He was particularly well known for

NOTE Confidence: 0.86898927

 $00:03:00.844 \longrightarrow 00:03:02.556$ working together with other luminaries

NOTE Confidence: 0.86898927

 $00{:}03{:}02.556 \dashrightarrow 00{:}03{:}04.940$ of our department of end of our field.

NOTE Confidence: 0.86898927

 $00:03:04.940 \longrightarrow 00:03:07.824$ Also let Anna Freud and JoJo Goldstein,

NOTE Confidence: 0.894491630769231

 $00{:}03{:}07.830 \dashrightarrow 00{:}03{:}09.860$ and they wrote a text called Beyond

NOTE Confidence: 0.894491630769231

 $00:03:09.860 \longrightarrow 00:03:11.489$ the best interests of the child,

NOTE Confidence: 0.894491630769231

 $00:03:11.490 \longrightarrow 00:03:14.148$ which is really a landmark in

NOTE Confidence: 0.894491630769231

 $00{:}03{:}14.150 \dashrightarrow 00{:}03{:}16.445$ in the development of child

NOTE Confidence: 0.894491630769231

 $00{:}03{:}16.445 {\:\dashrightarrow\:} 00{:}03{:}18.740$ psychiatry in the last century.

NOTE Confidence: 0.894491630769231

 $00:03:18.740 \longrightarrow 00:03:19.900$ Seymour, less than tragically,

 $00:03:19.900 \longrightarrow 00:03:21.636$ died at the age of I believe,

NOTE Confidence: 0.894491630769231

00:03:21.640 --> 00:03:25.399 51 in 1971 in a boating accident,

NOTE Confidence: 0.894491630769231

 $00:03:25.400 \longrightarrow 00:03:27.278$ and our department was robbed of

NOTE Confidence: 0.894491630769231

 $00:03:27.278 \longrightarrow 00:03:29.800$ one of its one of its luminaries.

NOTE Confidence: 0.894491630769231

00:03:29.800 --> 00:03:32.770 And shortly thereafter, in 1973,

NOTE Confidence: 0.894491630769231

 $00:03:32.770 \longrightarrow 00:03:35.380$ his family began their support of

NOTE Confidence: 0.894491630769231

 $00:03:35.380 \longrightarrow 00:03:38.386$ this award to honor his legacy and

NOTE Confidence: 0.894491630769231

 $00:03:38.386 \longrightarrow 00:03:41.144$ to honor the causes of science in

NOTE Confidence: 0.894491630769231

 $00{:}03{:}41.228 \dashrightarrow 00{:}03{:}44.018$ the service of great clinical care

NOTE Confidence: 0.894491630769231

 $00:03:44.020 \longrightarrow 00:03:45.880$ that we continue to celebrate.

NOTE Confidence: 0.88204783

 $00{:}03{:}47.900 \dashrightarrow 00{:}03{:}49.788$ So, remembering and celebrating

NOTE Confidence: 0.881005435294118

 $00:03:49.800 \longrightarrow 00:03:50.830$ that history of our department

NOTE Confidence: 0.881005435294118

 $00{:}03{:}50.830 \to 00{:}03{:}52.382$ is one reason that I think this

NOTE Confidence: 0.881005435294118

 $00:03:52.382 \longrightarrow 00:03:53.567$ is a particularly special day.

NOTE Confidence: 0.756742466666667

 $00:03:53.890 \longrightarrow 00:03:54.700$ A second reason

NOTE Confidence: 0.92832107

 $00:03:54.710 \longrightarrow 00:03:55.730$ is that it's a chance to

 $00:03:55.740 \longrightarrow 00:03:57.500$ honor and celebrate our commitment

NOTE Confidence: 0.86971968

 $00{:}03{:}57.500 \dashrightarrow 00{:}04{:}00.048$ to our trainings. I think that, uh.

NOTE Confidence: 0.86971968

00:04:00.048 --> 00:04:01.536 A particular characteristic of

NOTE Confidence: 0.86971968

 $00:04:01.536 \longrightarrow 00:04:03.401$ the department and one that I

NOTE Confidence: 0.86971968

 $00:04:03.401 \longrightarrow 00:04:04.991$ think through many of us here,

NOTE Confidence: 0.86971968

 $00:04:05.000 \longrightarrow 00:04:06.638$ is that our dedication to supporting

NOTE Confidence: 0.86971968

 $00:04:06.638 \longrightarrow 00:04:08.270$ the young people in our field.

NOTE Confidence: 0.86971968

 $00{:}04{:}08.270 \longrightarrow 00{:}04{:}09.966$ The people who are going to bring new,

NOTE Confidence: 0.86971968

 $00:04:09.970 \longrightarrow 00:04:12.460$ exciting ideas and move us forward.

NOTE Confidence: 0.86971968

 $00:04:12.460 \longrightarrow 00:04:14.640$ And that's what we do today.

NOTE Confidence: 0.86971968

00:04:14.640 --> 00:04:15.309 And of course,

NOTE Confidence: 0.86971968

 $00:04:15.310 \longrightarrow 00:04:17.192$ we also celebrate great science,

NOTE Confidence: 0.86971968

 $00:04:17.192 \longrightarrow 00:04:19.538$ and you're going to hear some

NOTE Confidence: 0.86971968

 $00:04:19.538 \longrightarrow 00:04:22.027$ wonderful science across a wide range

NOTE Confidence: 0.86971968

 $00:04:22.027 \longrightarrow 00:04:23.667$ of clinical translational areas.

 $00:04:23.670 \longrightarrow 00:04:26.480$ In the five presentations today.

NOTE Confidence: 0.86971968

 $00{:}04{:}26.480 \dashrightarrow 00{:}04{:}28.131$ And the final thing that we honor in

NOTE Confidence: 0.86971968

 $00:04:28.131 \longrightarrow 00:04:31.358$ this in this grand rounds is mentorship.

NOTE Confidence: 0.86971968

00:04:31.360 --> 00:04:32.860 The you know, bringing new,

NOTE Confidence: 0.86971968

00:04:32.860 --> 00:04:35.998 bringing new scientists into the field,

NOTE Confidence: 0.86971968

00:04:36.000 --> 00:04:36.924 training new clinicians,

NOTE Confidence: 0.86971968

 $00:04:36.924 \longrightarrow 00:04:38.464$ and advancing the careers of

NOTE Confidence: 0.86971968

 $00:04:38.464 \longrightarrow 00:04:40.378$ those who are going to lead us

NOTE Confidence: 0.86971968

 $00{:}04{:}40.378 \longrightarrow 00{:}04{:}41.623$ forward in the coming decades,

NOTE Confidence: 0.86971968

00:04:41.630 --> 00:04:44.116 only happens with the dedication of

NOTE Confidence: 0.86971968

00:04:44.116 --> 00:04:46.495 mentors who are willing to give of

NOTE Confidence: 0.86971968

 $00:04:46.495 \longrightarrow 00:04:48.349$ their time their energy and their

NOTE Confidence: 0.86971968

 $00:04:48.349 \longrightarrow 00:04:50.580$ caring to the to the young people

NOTE Confidence: 0.86971968

 $00:04:50.580 \longrightarrow 00:04:52.434$ who are entering our field and

NOTE Confidence: 0.86971968

 $00:04:52.440 \longrightarrow 00:04:54.673$ and so each of our honorees today

NOTE Confidence: 0.86971968

00:04:54.673 --> 00:04:56.500 will be introduced very briefly.

00:04:56.500 --> 00:04:57.864 Because of our schedule,

NOTE Confidence: 0.86971968

 $00{:}04{:}57.864 \dashrightarrow 00{:}04{:}59.569$ but very importantly by mentor

NOTE Confidence: 0.86971968

 $00:04:59.569 \longrightarrow 00:05:01.388$ who they've selected and who's

NOTE Confidence: 0.86971968

00:05:01.388 --> 00:05:03.506 been important to them and guiding

NOTE Confidence: 0.86971968

 $00:05:03.568 \longrightarrow 00:05:05.675$ their work and and in addition to

NOTE Confidence: 0.86971968

 $00:05:05.675 \longrightarrow 00:05:07.368$ honoring the award is we want to,

NOTE Confidence: 0.86971968

 $00:05:07.368 \longrightarrow 00:05:08.550$ we want to honor their mentors.

NOTE Confidence: 0.86971968

 $00:05:08.550 \longrightarrow 00:05:09.020$ Thank you.

NOTE Confidence: 0.86971968

 $00{:}05{:}09.020 \dashrightarrow 00{:}05{:}10.430$ Thank you to those mentors for

NOTE Confidence: 0.94170117

 $00:05:10.440 \longrightarrow 00:05:11.488$ being with us today.

NOTE Confidence: 0.9103648

 $00:05:13.370 \longrightarrow 00:05:15.297$ The last man selection committee,

NOTE Confidence: 0.9103648

00:05:15.297 --> 00:05:16.839 who I really want to thank,

NOTE Confidence: 0.9103648

 $00{:}05{:}16.840 --> 00{:}05{:}18.270$ consists of my Co chair,

NOTE Confidence: 0.9103648

00:05:18.270 --> 00:05:20.262 young Sunchoke, Kristen Brennan,

NOTE Confidence: 0.9103648

 $00:05:20.262 \longrightarrow 00:05:21.798$ Marina Picciotto, Khushoo,

00:05:21.798 --> 00:05:23.950 Mark Potenze, Tom Fernandez,

NOTE Confidence: 0.9103648

 $00:05:23.950 \longrightarrow 00:05:26.092$ Jerry Santa Cora with the support in

NOTE Confidence: 0.9103648

 $00:05:26.092 \longrightarrow 00:05:27.779$ the background from John Crystal.

NOTE Confidence: 0.9103648

 $00:05:27.780 \longrightarrow 00:05:29.444$ So I want to thank them for the

NOTE Confidence: 0.9103648

 $00:05:29.444 \longrightarrow 00:05:31.190$ time that went into this selection.

NOTE Confidence: 0.9103648

 $00:05:31.190 \longrightarrow 00:05:33.574$ We had an unusually difficult job this year.

NOTE Confidence: 0.9103648

 $00:05:33.580 \longrightarrow 00:05:35.656$ We had really a an unusually

NOTE Confidence: 0.9103648

 $00:05:35.656 \longrightarrow 00:05:37.534$ large group of really excellent

NOTE Confidence: 0.9103648

 $00{:}05{:}37.534 \dashrightarrow 00{:}05{:}39.614$ presentations and we're able to

NOTE Confidence: 0.9103648

 $00:05:39.614 \longrightarrow 00:05:42.543$ honor five of them today too with

NOTE Confidence: 0.9103648

 $00{:}05{:}42.543 \dashrightarrow 00{:}05{:}44.533$ first price first place awards.

NOTE Confidence: 0.9103648

 $00:05:44.540 \longrightarrow 00:05:45.800$ And three with one runner up

NOTE Confidence: 0.9103648

 $00{:}05{:}45.800 \dashrightarrow 00{:}05{:}46.955$ awards and we'll hear brief

NOTE Confidence: 0.9103648

 $00:05:46.955 \longrightarrow 00:05:48.285$ presentations from each of them.

NOTE Confidence: 0.8986194

 $00:05:49.510 \longrightarrow 00:05:50.590$ But the last thing I want to say

NOTE Confidence: 0.826499665833333

 $00{:}05{:}50.600 \dashrightarrow 00{:}05{:}51.940$ in introducing here is you

 $00:05:51.940 \longrightarrow 00:05:54.250$ know a bit of a poignant note.

NOTE Confidence: 0.826499665833333

 $00:05:54.250 \longrightarrow 00:05:58.324$ It's bittersweet to me to introduce this

NOTE Confidence: 0.826499665833333

 $00:05:58.324 \longrightarrow 00:06:01.706$ award today and to manage these ceremonies,

NOTE Confidence: 0.826499665833333

 $00:06:01.710 \longrightarrow 00:06:03.520$ because that's previously been done

NOTE Confidence: 0.826499665833333

00:06:03.520 --> 00:06:06.276 by one of my mentors, Bob Malison,

NOTE Confidence: 0.826499665833333

 $00:06:06.276 \longrightarrow 00:06:09.114$ who shepherded this process for the

NOTE Confidence: 0.826499665833333

 $00:06:09.114 \longrightarrow 00:06:13.230$ last 20 years and was taken from us.

NOTE Confidence: 0.826499665833333

00:06:13.230 --> 00:06:15.215 Like Seymour last man far

NOTE Confidence: 0.826499665833333

 $00:06:15.215 \longrightarrow 00:06:16.670$ too soon last summer.

NOTE Confidence: 0.826499665833333

 $00{:}06{:}16.670 \dashrightarrow 00{:}06{:}19.731$ And so I I regret that Bob is not

NOTE Confidence: 0.826499665833333

 $00:06:19.731 \longrightarrow 00:06:22.097$ with us today and I honor his

NOTE Confidence: 0.826499665833333

 $00:06:22.097 \longrightarrow 00:06:24.899$ memory in this in this presentation.

NOTE Confidence: 0.88412726

 $00{:}06{:}27.370 \dashrightarrow 00{:}06{:}29.640$ So with that introduction, let's

NOTE Confidence: 0.872006036363636

 $00:06:29.650 \longrightarrow 00:06:32.650$ move on. We have 5 great talks to

NOTE Confidence: 0.872006036363636

 $00:06:32.650 \longrightarrow 00:06:35.445$ here today and 1st is by ARCO.

00:06:35.445 --> 00:06:37.170 First place winner Zach Harvin.

NOTE Confidence: 0.872006036363636

 $00{:}06{:}37.170 \dashrightarrow 00{:}06{:}39.546$ And I'm going to invite his mentor Riggi

NOTE Confidence: 0.872006036363636

 $00:06:39.546 \longrightarrow 00:06:41.957$ to Sinha to give a brief introduction.

NOTE Confidence: 0.872006036363636

00:06:41.960 --> 00:06:42.944 Oh, I'm sorry.

NOTE Confidence: 0.872006036363636

 $00:06:42.944 \longrightarrow 00:06:44.256$ Just one brigitta sorry.

NOTE Confidence: 0.872006036363636

 $00:06:44.260 \longrightarrow 00:06:46.756$ One logistical thing which is about

NOTE Confidence: 0.872006036363636

00:06:46.756 --> 00:06:49.688 question since we do have five talks,

NOTE Confidence: 0.872006036363636

 $00:06:49.690 \longrightarrow 00:06:50.560$ we're going to go through today.

NOTE Confidence: 0.872006036363636

 $00{:}06{:}50.560 {\:\dashrightarrow\:} 00{:}06{:}52.051$ It's going to be a little tight

NOTE Confidence: 0.872006036363636

 $00:06:52.051 \longrightarrow 00:06:53.776$ so we're going to limit it to one

NOTE Confidence: 0.872006036363636

 $00{:}06{:}53.776 \dashrightarrow 00{:}06{:}55.096$ or two questions after the longer

NOTE Confidence: 0.872006036363636

 $00:06:55.096 \longrightarrow 00:06:56.713$ Co first place talks and we're not

NOTE Confidence: 0.872006036363636

00:06:56.713 --> 00:06:58.430 going to be able to have questions

NOTE Confidence: 0.872006036363636

 $00:06:58.430 \longrightarrow 00:07:00.010$ after the shorter runner up talks.

NOTE Confidence: 0.872006036363636

 $00:07:00.010 \longrightarrow 00:07:01.535$ If we stayed remarkably on

NOTE Confidence: 0.872006036363636

 $00:07:01.535 \longrightarrow 00:07:03.639$ time and have time at the end,

 $00:07:03.640 \longrightarrow 00:07:05.383$ then we can perhaps have a little

NOTE Confidence: 0.872006036363636

 $00:07:05.383 \longrightarrow 00:07:06.673$ time for questions and discussion

NOTE Confidence: 0.872006036363636

 $00:07:06.673 \longrightarrow 00:07:08.137$ for any of the presenters at

NOTE Confidence: 0.872006036363636

 $00:07:08.137 \longrightarrow 00:07:09.450$ the end if time permits.

NOTE Confidence: 0.872006036363636

00:07:09.450 --> 00:07:10.488 So with that,

NOTE Confidence: 0.872006036363636

 $00:07:10.488 \longrightarrow 00:07:11.180$ Regina, please.

NOTE Confidence: 0.865723695

00:07:12.180 --> 00:07:15.840 Thank you Chris. It's my real pleasure.

NOTE Confidence: 0.865723695

00:07:15.840 --> 00:07:17.880 An honor to introduce Doctor Zachary.

NOTE Confidence: 0.865723695

 $00{:}07{:}17.880 \to 00{:}07{:}21.653$ Have our neck and congratulations Zach.

NOTE Confidence: 0.865723695

 $00:07:21.653 \longrightarrow 00:07:24.477$ Let me give you a little quick background.

NOTE Confidence: 0.865723695

 $00{:}07{:}24.480 \dashrightarrow 00{:}07{:}27.808$ Zach grew up in Boulder, Co and attended Duke

NOTE Confidence: 0.860402983333333

 $00:07:27.820 \longrightarrow 00:07:30.090$ University for undergraduate work, where

NOTE Confidence: 0.888533950666667

 $00{:}07{:}30.100 \dashrightarrow 00{:}07{:}31.984$ he studied biomedical engineering

NOTE Confidence: 0.888533950666667

 $00:07:31.984 \longrightarrow 00:07:34.339$ and biology and then became

NOTE Confidence: 0.888533950666667

 $00:07:34.339 \longrightarrow 00:07:36.449$ interested in the biology of aging.

 $00:07:36.450 \longrightarrow 00:07:39.145$ He went on to the University of

NOTE Confidence: 0.888533950666667

00:07:39.145 --> 00:07:41.350 Michigan for his MD and pH D.

NOTE Confidence: 0.888533950666667

00:07:41.350 --> 00:07:43.234 His dissertation was focused

NOTE Confidence: 0.888533950666667

 $00:07:43.234 \longrightarrow 00:07:44.647$ on neurobiological mechanisms

NOTE Confidence: 0.8272593975

00:07:44.660 --> 00:07:46.980 through which social stressors

NOTE Confidence: 0.866349392

 $00:07:46.990 \longrightarrow 00:07:49.500$ influenced aging and fruit flies.

NOTE Confidence: 0.866349392

 $00:07:49.500 \longrightarrow 00:07:51.070$ That was really fruitful.

NOTE Confidence: 0.866349392

 $00:07:51.070 \longrightarrow 00:07:54.430$ It led to four top notch

NOTE Confidence: 0.866349392

00:07:54.430 --> 00:07:55.790 publications in science,

NOTE Confidence: 0.866349392

 $00:07:55.790 \longrightarrow 00:07:58.190$ nature, ecology and so on.

NOTE Confidence: 0.866349392

00:07:58.190 --> 00:08:00.486 And then after returning to medical school,

NOTE Confidence: 0.866349392

 $00:08:00.490 \longrightarrow 00:08:02.880$ he developed an interest in

NOTE Confidence: 0.866349392

 $00:08:02.880 \longrightarrow 00:08:05.220$ psychiatry and the mechanisms through

NOTE Confidence: 0.866349392

 $00:08:05.220 \longrightarrow 00:08:06.670$ which stress and mental illness

NOTE Confidence: 0.866349392

 $00:08:06.670 \longrightarrow 00:08:08.804$ influence physical health and aging.

NOTE Confidence: 0.866349392

 $00:08:08.804 \longrightarrow 00:08:12.008$ He came to Yale and we were thrilled

 $00:08:12.010 \longrightarrow 00:08:14.894$ to have him and has worked with

NOTE Confidence: 0.866349392

 $00{:}08{:}14.894 \dashrightarrow 00{:}08{:}16.370$ myself at the Yale Stress Center

NOTE Confidence: 0.866349392

 $00:08:16.370 \longrightarrow 00:08:18.918$ as well as with Doctor Kasshu.

NOTE Confidence: 0.866349392

00:08:18.920 --> 00:08:20.656 Really in, I guess,

NOTE Confidence: 0.866349392

 $00:08:20.656 \longrightarrow 00:08:22.826$ very apropos for this award.

NOTE Confidence: 0.866349392

00:08:22.830 --> 00:08:25.378 An in the legacy of Seymour Lessman,

NOTE Confidence: 0.866349392

 $00:08:25.380 \longrightarrow 00:08:26.835$ a sort of bringing nature

NOTE Confidence: 0.866349392

 $00:08:26.835 \longrightarrow 00:08:28.095$ and nurture together,

NOTE Confidence: 0.866349392

 $00{:}08{:}28.095 \dashrightarrow 00{:}08{:}30.528$ looking at epigenetic mechanisms

NOTE Confidence: 0.866349392

 $00:08:30.530 \longrightarrow 00:08:32.890$ by which stress may influence

NOTE Confidence: 0.866349392

 $00:08:32.890 \longrightarrow 00:08:34.083$ the process of aging.

NOTE Confidence: 0.866349392

 $00{:}08{:}34.083 \dashrightarrow 00{:}08{:}36.441$ He's also collaborating with the sickle

NOTE Confidence: 0.866349392

 $00:08:36.441 \longrightarrow 00:08:38.830$ cell program to examine psychological

NOTE Confidence: 0.866349392

 $00:08:38.830 \longrightarrow 00:08:41.030$ resilience influences on pain,

NOTE Confidence: 0.866349392

 $00:08:41.030 \longrightarrow 00:08:42.419$ an overall health.

00:08:42.420 --> 00:08:44.270 It's been a real pleasure

NOTE Confidence: 0.801790105

 $00:08:44.340 \longrightarrow 00:08:46.948$ for Doctor Kosu and myself to work closely

NOTE Confidence: 0.851915884

 $00:08:46.960 \longrightarrow 00:08:48.900$ with Zach. He's been wonderful.

NOTE Confidence: 0.851915884

 $00:08:48.900 \longrightarrow 00:08:50.444$ His optimism and positive

NOTE Confidence: 0.851915884

 $00:08:50.444 \longrightarrow 00:08:53.263$ energy and an just burst of new

NOTE Confidence: 0.851915884

00:08:53.263 --> 00:08:55.388 ideas has been very refreshing,

NOTE Confidence: 0.851915884

 $00:08:55.390 \longrightarrow 00:09:00.290$ and so I'm thrilled to.

NOTE Confidence: 0.851915884

 $00:09:00.290 \longrightarrow 00:09:02.478$ Congratulate him and would invite

NOTE Confidence: 0.851915884

 $00:09:02.478 \dashrightarrow 00:09:05.936$ you to join me in in wishing him and

NOTE Confidence: 0.851915884

 $00:09:05.936 \longrightarrow 00:09:09.800$ in hearing what he has to say. Zach

NOTE Confidence: 0.857937652222222

 $00:09:10.740 \longrightarrow 00:09:12.825$ thank you for that overly

NOTE Confidence: 0.857937652222222

 $00:09:12.825 \longrightarrow 00:09:14.493$ kind introduction for beta.

NOTE Confidence: 0.856069037272727

00:09:17.570 --> 00:09:19.621 So, uh, today I'll be talking to

NOTE Confidence: 0.856069037272727

00:09:19.621 --> 00:09:21.492 you about how psychological and

NOTE Confidence: 0.856069037272727

 $00:09:21.492 \longrightarrow 00:09:23.348$ biological resilience modulate the

NOTE Confidence: 0.856069037272727

 $00:09:23.348 \longrightarrow 00:09:26.359$ effects of stress on epigenetic aging.

 $00:09:26.360 \longrightarrow 00:09:30.400$ Next slide. 1st, I have no relevant

NOTE Confidence: 0.856069037272727

 $00:09:30.400 \longrightarrow 00:09:32.390$ disclosures or conflicts of interest,

NOTE Confidence: 0.856069037272727

 $00{:}09{:}32.390 \dashrightarrow 00{:}09{:}34.466$ but now, as Regina mentioned before,

NOTE Confidence: 0.856069037272727

00:09:34.470 --> 00:09:36.962 coming to GAIL, I studied the mechanisms

NOTE Confidence: 0.856069037272727

 $00{:}09{:}36.962 \dashrightarrow 00{:}09{:}38.504$ through which social stressors

NOTE Confidence: 0.856069037272727

00:09:38.504 --> 00:09:40.549 regulate aging and intra Sofala,

NOTE Confidence: 0.856069037272727

 $00:09:40.550 \longrightarrow 00:09:41.663$ and I'm not going to bore you

NOTE Confidence: 0.856069037272727

00:09:41.663 --> 00:09:42.610 with talk about fruit flies.

NOTE Confidence: 0.856069037272727

 $00:09:42.610 \longrightarrow 00:09:45.190$ But fundamentally what we found was

NOTE Confidence: 0.856069037272727

 $00:09:45.190 \longrightarrow 00:09:47.950$ that perception of the opposite sex,

NOTE Confidence: 0.856069037272727

 $00:09:47.950 \longrightarrow 00:09:50.680$ basically a social stress led to changes

NOTE Confidence: 0.856069037272727

 $00:09:50.680 \longrightarrow 00:09:52.550$ in neuropeptide ergic signaling,

NOTE Confidence: 0.856069037272727

00:09:52.550 --> 00:09:53.576 downstream Physiology,

NOTE Confidence: 0.856069037272727

 $00:09:53.576 \longrightarrow 00:09:56.654$ and ultimately accelerated aging and death.

NOTE Confidence: 0.856069037272727

 $00:09:56.660 \longrightarrow 00:09:59.156$ However, these negative outcomes could be

 $00:09:59.156 \longrightarrow 00:10:01.330$ minimized by specific protective factors.

NOTE Confidence: 0.856069037272727

 $00:10:01.330 \longrightarrow 00:10:02.956$ In this case it was successful

NOTE Confidence: 0.856069037272727

 $00:10:02.956 \longrightarrow 00:10:03.769$ made next slide.

NOTE Confidence: 0.867585632222222

 $00:10:07.660 \longrightarrow 00:10:09.070$ After finishing my PhD in

NOTE Confidence: 0.867585632222222

00:10:09.070 --> 00:10:10.198 returning to medical school,

NOTE Confidence: 0.867585632222222

00:10:10.200 --> 00:10:12.349 I learned what psychiatry and I suspect

NOTE Confidence: 0.867585632222222

00:10:12.349 --> 00:10:14.776 most people here have known for a long

NOTE Confidence: 0.867585632222222

 $00:10:14.776 \longrightarrow 00:10:16.647$ time that patients with serious mental

NOTE Confidence: 0.867585632222222

 $00:10:16.647 \dashrightarrow 00:10:18.807$ illness die earlier than those without.

NOTE Confidence: 0.867585632222222

 $00:10:18.810 \longrightarrow 00:10:20.619$ The plot on the left is from a study

NOTE Confidence: 0.867585632222222

 $00{:}10{:}20.619 \dashrightarrow 00{:}10{:}22.065$ out of Denmark demonstrating that

NOTE Confidence: 0.867585632222222

00:10:22.065 --> 00:10:24.244 for pretty much all causes of death

NOTE Confidence: 0.867585632222222

 $00:10:24.244 \longrightarrow 00:10:25.612$ and mortality rates significantly

NOTE Confidence: 0.867585632222222

 $00:10:25.612 \longrightarrow 00:10:27.818$ higher in patients with mood disorders

NOTE Confidence: 0.867585632222222

00:10:27.818 --> 00:10:29.770 compared to healthy controls,

NOTE Confidence: 0.867585632222222

 $00:10:29.770 \longrightarrow 00:10:31.840$ and they've gone to demonstrate similar

 $00:10:31.840 \longrightarrow 00:10:33.770$ findings for other psychiatric disorders.

NOTE Confidence: 0.867585632222222

00:10:33.770 --> 00:10:34.236 Now, notably,

NOTE Confidence: 0.867585632222222

 $00:10:34.236 \longrightarrow 00:10:35.867$ this isn't just true for things we

NOTE Confidence: 0.867585632222222

 $00:10:35.867 \longrightarrow 00:10:37.570$ might expect to be psychiatric related.

NOTE Confidence: 0.867585632222222

00:10:37.570 --> 00:10:40.002 Also infections, cardiovascular disease,

NOTE Confidence: 0.867585632222222

 $00:10:40.002 \longrightarrow 00:10:41.826$ other similar costs.

NOTE Confidence: 0.867585632222222

 $00:10:41.830 \longrightarrow 00:10:43.060$ And this is a pattern similar

NOTE Confidence: 0.867585632222222

 $00:10:43.060 \longrightarrow 00:10:44.589$ to what we see as people age.

NOTE Confidence: 0.867585632222222

 $00:10:44.590 \longrightarrow 00:10:46.837$ You are risk for many different diseases

NOTE Confidence: 0.867585632222222

00:10:46.837 --> 00:10:49.041 increases as we get older and on the

NOTE Confidence: 0.867585632222222

00:10:49.041 --> 00:10:51.158 right is a data from a meta analysis

NOTE Confidence: 0.867585632222222

 $00:10:51.158 \longrightarrow 00:10:53.209$ showing that across a wide range of

NOTE Confidence: 0.867585632222222

 $00{:}10{:}53.210 {\:{\circ}{\circ}{\circ}}>00{:}10{:}55.650$ different categories of mental disorders,

NOTE Confidence: 0.867585632222222

 $00{:}10{:}55.650 \dashrightarrow 00{:}10{:}57.732$ patients tend to have shorter TILA

NOTE Confidence: 0.867585632222222

00:10:57.732 --> 00:10:59.456 mirrors than healthy controls and

00:10:59.456 --> 00:11:01.388 shorten telem ears are thought to be

NOTE Confidence: 0.867585632222222

 $00:11:01.388 \longrightarrow 00:11:03.468$ related to accelerated rates of aging.

NOTE Confidence: 0.867585632222222 00:11:03.470 --> 00:11:05.710 Next slide. NOTE Confidence: 0.867585632222222

00:11:05.710 --> 00:11:07.798 Now one uncertain thing about our

NOTE Confidence: 0.867585632222222

 $00:11:07.798 \longrightarrow 00:11:09.640$ mental health diagnostics is that

NOTE Confidence: 0.867585632222222

 $00:11:09.640 \longrightarrow 00:11:11.605$ the difference between pathologic and

NOTE Confidence: 0.867585632222222

 $00:11:11.605 \longrightarrow 00:11:13.570$ non pathologic isn't always clear.

NOTE Confidence: 0.867585632222222

 $00{:}11{:}13.570 \dashrightarrow 00{:}11{:}15.655$ Other researches has shown that

NOTE Confidence: 0.8675856322222222

 $00{:}11{:}15.655 --> 00{:}11{:}16.489 \ {\rm certain \ stressors},$

NOTE Confidence: 0.867585632222222

 $00:11:16.490 \longrightarrow 00:11:18.527$ even in the absence of a diagnosed

NOTE Confidence: 0.8675856322222222

 $00{:}11{:}18.527 --> 00{:}11{:}19.109 \ \mathrm{mental \ illness},$

NOTE Confidence: 0.867585632222222

 $00:11:19.110 \longrightarrow 00:11:20.998$ can cause similar patterns

NOTE Confidence: 0.867585632222222

 $00:11:20.998 \longrightarrow 00:11:22.414$ of accelerated aging.

NOTE Confidence: 0.867585632222222

00:11:22.420 --> 00:11:24.232 A range of studies have examined

NOTE Confidence: 0.867585632222222

 $00:11:24.232 \longrightarrow 00:11:25.440$ physical health outcomes and

NOTE Confidence: 0.867585632222222

 $00{:}11{:}25.492 \dashrightarrow 00{:}11{:}27.297$ stealing your length associated with

00:11:27.297 --> 00:11:29.102 stressors like early life adversity,

NOTE Confidence: 0.867585632222222

00:11:29.110 --> 00:11:31.118 associate economic status, discrimination,

NOTE Confidence: 0.867585632222222

 $00:11:31.118 \longrightarrow 00:11:33.126$ or even medical internship.

NOTE Confidence: 0.867585632222222

 $00:11:33.130 \longrightarrow 00:11:34.621$ The plot here is from the intern

NOTE Confidence: 0.867585632222222

00:11:34.621 --> 00:11:35.047 Health study,

NOTE Confidence: 0.867585632222222

00:11:35.050 --> 00:11:37.154 which is actually being run by a yell

NOTE Confidence: 0.867585632222222

 $00:11:37.154 \longrightarrow 00:11:39.112$ cyka lump doctor Sen and shows the

NOTE Confidence: 0.867585632222222

 $00{:}11{:}39.112 \dashrightarrow 00{:}11{:}41.049$ more hours per week into his work.

NOTE Confidence: 0.867585632222222

 $00:11:41.050 \longrightarrow 00:11:42.038$ The more their tillers,

NOTE Confidence: 0.867585632222222

 $00:11:42.038 \longrightarrow 00:11:44.050$ short and over the course of that year.

NOTE Confidence: 0.867585632222222

00:11:44.050 --> 00:11:46.710 Now this idea brought me back to my PhD work.

NOTE Confidence: 0.867585632222222

 $00:11:46.710 \longrightarrow 00:11:48.696$ If a stress like this can

NOTE Confidence: 0.8675856322222222

00:11:48.696 --> 00:11:49.689 cause accelerated aging,

NOTE Confidence: 0.867585632222222

 $00{:}11{:}49.690 \dashrightarrow 00{:}11{:}51.384$ we should be able to identify the

NOTE Confidence: 0.867585632222222

00:11:51.384 --> 00:11:52.539 physiologic pathways to which is.

00:11:52.540 --> 00:11:54.820 Happening and hopefully we can identify

NOTE Confidence: 0.867585632222222

 $00{:}11{:}54.820 \dashrightarrow 00{:}11{:}58.400$ ways to protect against next slide.

NOTE Confidence: 0.867585632222222

00:11:58.400 --> 00:11:59.688 And while psychiatric diagnosis

NOTE Confidence: 0.867585632222222

 $00:11:59.688 \longrightarrow 00:12:00.654$ can be unclear,

NOTE Confidence: 0.867585632222222

 $00:12:00.660 \longrightarrow 00:12:03.076$ the term stress can times be equally vague.

NOTE Confidence: 0.867585632222222

 $00:12:03.080 \longrightarrow 00:12:04.556$ So I want to provide a

NOTE Confidence: 0.867585632222222

 $00:12:04.556 \longrightarrow 00:12:05.980$ definition that can be helped.

NOTE Confidence: 0.867585632222222

 $00:12:05.980 \longrightarrow 00:12:08.437$ We can define stress as a process,

NOTE Confidence: 0.867585632222222

 $00{:}12{:}08.440 \dashrightarrow 00{:}12{:}10.066$ and it's the process of identifying,

NOTE Confidence: 0.867585632222222

 $00:12:10.070 \longrightarrow 00:12:11.429$ interpreting, responding to,

NOTE Confidence: 0.8675856322222222

 $00:12:11.429 \longrightarrow 00:12:13.241$ and adapting to potential

NOTE Confidence: 0.867585632222222

 $00:12:13.241 \longrightarrow 00:12:14.600$ threats or challenges.

NOTE Confidence: 0.867585632222222

 $00:12:14.600 \longrightarrow 00:12:16.875$ Now first it involves individuals

NOTE Confidence: 0.867585632222222

00:12:16.875 --> 00:12:18.240 identifying these stressors,

NOTE Confidence: 0.867585632222222 00:12:18.240 --> 00:12:18.902 and second, NOTE Confidence: 0.867585632222222

 $00{:}12{:}18.902 \dashrightarrow 00{:}12{:}20.557$ it involves their interpretation of

 $00{:}12{:}20.557 \dashrightarrow 00{:}12{:}22.705$ the stressor which could include both

NOTE Confidence: 0.867585632222222

 $00{:}12{:}22.705 \dashrightarrow 00{:}12{:}24.169$ societal and individual factors.

NOTE Confidence: 0.867585632222222

 $00{:}12{:}24.170 \dashrightarrow 00{:}12{:}26.798$ And this interpretation could lead to

NOTE Confidence: 0.867585632222222

 $00:12:26.798 \longrightarrow 00:12:29.310$ either amplifying or suppressing the stress.

NOTE Confidence: 0.867585632222222

 $00:12:29.310 \longrightarrow 00:12:30.542$ Depending on these factors,

NOTE Confidence: 0.8675856322222222

00:12:30.542 --> 00:12:30.850 next,

NOTE Confidence: 0.867585632222222

 $00:12:30.850 \longrightarrow 00:12:32.270$ there's an acute response and

NOTE Confidence: 0.867585632222222

 $00:12:32.270 \longrightarrow 00:12:34.136$ it can be behavioral but also

NOTE Confidence: 0.867585632222222

 $00{:}12{:}34.136 \to 00{:}12{:}36.066$ physiological heart rate might elevate.

NOTE Confidence: 0.867585632222222

 $00{:}12{:}36.070 \dashrightarrow 00{:}12{:}37.514$ Specific circuits might fire

NOTE Confidence: 0.867585632222222

 $00{:}12{:}37.514 \dashrightarrow 00{:}12{:}39.319$ changes might occur in cortisol

NOTE Confidence: 0.867585632222222

 $00:12:39.319 \longrightarrow 00:12:40.830$ or metabolic pathways,

NOTE Confidence: 0.867585632222222

 $00{:}12{:}40.830 \mathrel{--}{>} 00{:}12{:}43.110$ and ultimately these short term

NOTE Confidence: 0.867585632222222

 $00:12:43.110 \longrightarrow 00:12:46.500$ responses can lead to long term effects.

NOTE Confidence: 0.867585632222222 $00:12:46.500 \longrightarrow 00:12:48.680$ Next slide. NOTE Confidence: 0.867585632222222 00:12:48.680 --> 00:12:49.016 Now,

NOTE Confidence: 0.867585632222222

 $00:12:49.016 \longrightarrow 00:12:50.696$ using this definition of stress,

NOTE Confidence: 0.867585632222222

 $00:12:50.700 \longrightarrow 00:12:53.458$ we can return to this overall hypothesis

NOTE Confidence: 0.867585632222222

 $00{:}12{:}53.458 \dashrightarrow 00{:}12{:}55.518$ that stress accelerates aging via

NOTE Confidence: 0.867585632222222

 $00:12:55.518 \longrightarrow 00:12:57.142$ physiologic changes moderated by

NOTE Confidence: 0.867585632222222

00:12:57.142 --> 00:12:59.760 protective factors in a systematic way.

NOTE Confidence: 0.867585632222222 00:12:59.760 --> 00:13:00.081 First,

NOTE Confidence: 0.867585632222222

 $00{:}13{:}00.081 \dashrightarrow 00{:}13{:}01.686$ we can ask whether cumulative

NOTE Confidence: 0.8675856322222222

 $00:13:01.686 \longrightarrow 00:13:02.970$ lifetime stress leads to

NOTE Confidence: 0.82803308

 $00:13:03.031 \longrightarrow 00:13:05.243$ accelerated aging as a long term consequence

NOTE Confidence: 0.82803308

 $00:13:05.243 \longrightarrow 00:13:07.780$ in an otherwise healthy Community population.

NOTE Confidence: 0.82803308

00:13:07.780 --> 00:13:09.436 Even in the absence of diagnosis,

NOTE Confidence: 0.82803308

 $00:13:09.440 \longrightarrow 00:13:12.744$ mental illness. If so, we can ask whether

NOTE Confidence: 0.82803308

00:13:12.744 --> 00:13:14.998 stress related Physiology like changes

NOTE Confidence: 0.82803308

00:13:14.998 --> 00:13:18.148 in the HPA axis and insulin signaling

NOTE Confidence: 0.82803308

 $00{:}13{:}18.225 \dashrightarrow 00{:}13{:}21.027$ are also related to accelerated aging.

00:13:21.030 --> 00:13:24.174 And finally, we can ask if an individual

NOTE Confidence: 0.82803308

 $00{:}13{:}24.174 \longrightarrow 00{:}13{:}26.532$ psychological resilience can serve as a

NOTE Confidence: 0.82803308

 $00:13:26.532 \longrightarrow 00:13:28.412$ protective factor in these relationships.

NOTE Confidence: 0.82803308

00:13:28.420 --> 00:13:31.464 Next slide. Now to measure aging,

NOTE Confidence: 0.82803308

 $00{:}13{:}31.464 \dashrightarrow 00{:}13{:}33.460$ a lot of the previous studies I've

NOTE Confidence: 0.82803308

00:13:33.460 --> 00:13:35.335 mentioned utilized stealing their life,

NOTE Confidence: 0.82803308

00:13:35.340 --> 00:13:37.108 particularly in young populations.

NOTE Confidence: 0.82803308

 $00:13:37.108 \longrightarrow 00:13:40.260$ We we don't have more obvious indications

NOTE Confidence: 0.82803308

 $00:13:40.260 \longrightarrow 00:13:42.804$ of aging like frailty or death,

NOTE Confidence: 0.82803308

 $00:13:42.810 \longrightarrow 00:13:45.029$ but telomere length is really only weakly

NOTE Confidence: 0.82803308

 $00{:}13{:}45.029 \dashrightarrow 00{:}13{:}46.920$ correlated to aging related outcomes.

NOTE Confidence: 0.82803308

00:13:46.920 --> 00:13:48.890 Might care about like morbidity

NOTE Confidence: 0.82803308

 $00{:}13{:}48.890 \dashrightarrow 00{:}13{:}49.678$ and mortality.

NOTE Confidence: 0.82803308

00:13:49.680 --> 00:13:51.918 Luckily, recent advances in epigenetics have

NOTE Confidence: 0.82803308

 $00:13:51.918 \longrightarrow 00:13:54.640$ led to the development of epigenetic clocks,

 $00:13:54.640 \longrightarrow 00:13:56.690$ and these are based on

NOTE Confidence: 0.82803308

00:13:56.690 --> 00:13:57.920 DNA methylation patterns,

NOTE Confidence: 0.82803308

 $00:13:57.920 \longrightarrow 00:13:59.880$ and these clocks have really been a leap

NOTE Confidence: 0.82803308

00:13:59.880 --> 00:14:01.956 forward in terms of predictions of frailty,

NOTE Confidence: 0.82803308

 $00:14:01.960 \longrightarrow 00:14:03.820$ morbidity and mortality when

NOTE Confidence: 0.82803308

 $00:14:03.820 \longrightarrow 00:14:06.145$ compared to tumor based studies.

NOTE Confidence: 0.82803308

 $00:14:06.150 \longrightarrow 00:14:07.595$ Here we're going to utilize

NOTE Confidence: 0.82803308

00:14:07.595 --> 00:14:09.040 one of these epigenetic clocks,

NOTE Confidence: 0.82803308

00:14:09.040 --> 00:14:11.330 rimage, to address our hypothesis,

NOTE Confidence: 0.82803308

00:14:11.330 --> 00:14:13.090 but I think we might actually hear more

NOTE Confidence: 0.82803308

 $00{:}14{:}13.090 \dashrightarrow 00{:}14{:}15.010$ about how we can continue to improve our

NOTE Confidence: 0.82803308

00:14:15.010 --> 00:14:16.690 measures of aging from Albert Higgins,

NOTE Confidence: 0.82803308

 $00:14:16.690 \longrightarrow 00:14:17.214$ Chanina.

NOTE Confidence: 0.82803308

00:14:17.214 --> 00:14:18.262 Later presentation,

NOTE Confidence: 0.82803308

00:14:18.262 --> 00:14:19.310 next slide. NOTE Confidence: 0.852387335714286

 $00:14:21.820 \longrightarrow 00:14:24.793$ Our study population was a group of 444

00:14:24.793 --> 00:14:27.297 individuals between the ages of 18 and 50,

NOTE Confidence: 0.852387335714286

 $00:14:27.300 \longrightarrow 00:14:29.440$ taking no prescription medications with

NOTE Confidence: 0.852387335714286

 $00:14:29.440 \longrightarrow 00:14:32.400$ no chronic health issues and with no DSM.

NOTE Confidence: 0.852387335714286

00:14:32.400 --> 00:14:34.020 Four diagnosis of their

NOTE Confidence: 0.852387335714286

 $00:14:34.020 \longrightarrow 00:14:35.235$ indicati use disorder.

NOTE Confidence: 0.852387335714286

00:14:35.240 --> 00:14:37.470 So healthy in this group,

NOTE Confidence: 0.852387335714286

 $00:14:37.470 \longrightarrow 00:14:39.295$ we obtained survey measurements of

NOTE Confidence: 0.852387335714286

 $00:14:39.295 \longrightarrow 00:14:40.755$ stress and psychological resilience,

NOTE Confidence: 0.852387335714286

 $00{:}14{:}40.760 \dashrightarrow 00{:}14{:}43.616$ as well as physiological epigeic measure.

NOTE Confidence: 0.852387335714286

00:14:43.620 --> 00:14:47.158 Next slide. Now in this population

NOTE Confidence: 0.852387335714286

00:14:47.158 --> 00:14:49.150 we see a positive correlation between

NOTE Confidence: 0.852387335714286

 $00:14:49.205 \longrightarrow 00:14:51.725$ cumulative stress levels as measured by

NOTE Confidence: 0.852387335714286

 $00:14:51.725 \longrightarrow 00:14:53.405$ interview and grammage acceleration.

NOTE Confidence: 0.852387335714286

00:14:53.410 --> 00:14:55.082 I want to take a moment to discuss

NOTE Confidence: 0.852387335714286

 $00:14:55.082 \longrightarrow 00:14:56.282$ these measures because we're going

00:14:56.282 --> 00:14:57.990 to be using them throughout the talk.

NOTE Confidence: 0.852387335714286

 $00{:}14{:}57.990 \dashrightarrow 00{:}14{:}59.358$ The cumulative adversity index

NOTE Confidence: 0.852387335714286

00:14:59.358 --> 00:15:01.969 or CE AI is on the X axis,

NOTE Confidence: 0.852387335714286

00:15:01.970 --> 00:15:04.133 and it's an interview based measure of

NOTE Confidence: 0.852387335714286

 $00:15:04.133 \longrightarrow 00:15:05.690$ cumulative stress taking into account

NOTE Confidence: 0.852387335714286

00:15:05.690 --> 00:15:07.713 of a multitude of different types of

NOTE Confidence: 0.852387335714286

00:15:07.713 --> 00:15:09.293 stressors across the lifespan and

NOTE Confidence: 0.852387335714286

00:15:09.293 --> 00:15:11.147 higher lifetime stress leads to higher.

NOTE Confidence: 0.852387335714286

 $00:15:11.150 \longrightarrow 00:15:11.939$ See AI score.

NOTE Confidence: 0.852387335714286

00:15:11.939 --> 00:15:14.491 You can see that see AI is positively

NOTE Confidence: 0.852387335714286

 $00:15:14.491 \longrightarrow 00:15:17.067$ correlated with brimmage acceleration,

NOTE Confidence: 0.852387335714286

 $00:15:17.070 \longrightarrow 00:15:19.110$ which fundamentally represents the

NOTE Confidence: 0.852387335714286

 $00{:}15{:}19.110 \dashrightarrow 00{:}15{:}20.640$ difference between individuals.

NOTE Confidence: 0.852387335714286

 $00:15:20.640 \longrightarrow 00:15:22.670$ Epigenetic age and chronological age

NOTE Confidence: 0.852387335714286

00:15:22.670 --> 00:15:24.700 with a positive number indicating

NOTE Confidence: 0.852387335714286

 $00:15:24.765 \longrightarrow 00:15:27.099$ that they are biologically older than

 $00:15:27.099 \longrightarrow 00:15:29.180$ their chronological age would suggest.

NOTE Confidence: 0.852387335714286 00:15:29.180 --> 00:15:31.240 Next slide. NOTE Confidence: 0.852387335714286

 $00:15:31.240 \longrightarrow 00:15:32.968$ The one potential explanation for these

NOTE Confidence: 0.852387335714286

00:15:32.968 --> 00:15:34.716 findings is that stress might result

NOTE Confidence: 0.852387335714286

00:15:34.716 --> 00:15:36.348 in substance use, behavior changes,

NOTE Confidence: 0.852387335714286

 $00:15:36.348 \longrightarrow 00:15:39.176$ or be due to different demographic factors.

NOTE Confidence: 0.852387335714286

 $00:15:39.180 \longrightarrow 00:15:42.316$ But even when we take into account

NOTE Confidence: 0.852387335714286

 $00{:}15{:}42.316 \rightarrow 00{:}15{:}45.280$ a smoking BMI, alcohol use, race,

NOTE Confidence: 0.852387335714286

00:15:45.280 --> 00:15:47.378 sex, marital status, income,

NOTE Confidence: 0.852387335714286 00:15:47.378 --> 00:15:47.856 education, NOTE Confidence: 0.852387335714286

 $00:15:47.856 \longrightarrow 00:15:50.724$ when we do that via multivariate

NOTE Confidence: 0.852387335714286

 $00:15:50.724 \longrightarrow 00:15:51.680$ linear regression,

NOTE Confidence: 0.852387335714286

 $00{:}15{:}51.680 \dashrightarrow 00{:}15{:}54.145$ there's still a significant independent

NOTE Confidence: 0.852387335714286

 $00:15:54.145 \longrightarrow 00:15:56.610$ effective stress on grammage acceleration.

NOTE Confidence: 0.852387335714286

00:15:56.610 --> 00:15:58.368 Now, except where I mentioned otherwise,

 $00:15:58.370 \longrightarrow 00:16:00.085$ all the rest of our analysis will

NOTE Confidence: 0.852387335714286

 $00{:}16{:}00.085 \dashrightarrow 00{:}16{:}01.869$ account for all of these covariates,

NOTE Confidence: 0.85238733571428600:16:01.870 --> 00:16:02.820 and notably,

NOTE Confidence: 0.852387335714286

 $00:16:02.820 \longrightarrow 00:16:06.145$ these covers are related to to aging,

NOTE Confidence: 0.852387335714286

 $00:16:06.150 \longrightarrow 00:16:08.621$ as demonstrated by the large change in

NOTE Confidence: 0.852387335714286

00:16:08.621 --> 00:16:11.179 the R-squared you get from the simple

NOTE Confidence: 0.852387335714286

 $00:16:11.179 \longrightarrow 00:16:12.974$ going from the simple relationship

NOTE Confidence: 0.852387335714286

 $00:16:12.974 \longrightarrow 00:16:15.480$ on the plot to the full models R 2 .

NOTE Confidence: 0.852387335714286 00:16:15.480 --> 00:16:17.070 Next slide. NOTE Confidence: 0.852387335714286

00:16:17.070 --> 00:16:18.870 So going back to our hypothesis,

NOTE Confidence: 0.852387335714286

 $00{:}16{:}18.870 --> 00{:}16{:}19.490 \ cumulative \ stress,$

NOTE Confidence: 0.852387335714286

 $00:16:19.490 \longrightarrow 00:16:21.660$ even in the absence of mental illness,

NOTE Confidence: 0.852387335714286

 $00:16:21.660 \longrightarrow 00:16:23.820$ is associated with accelerated aging.

NOTE Confidence: 0.852387335714286

00:16:23.820 --> 00:16:25.950 Even were accounting for all of

NOTE Confidence: 0.852387335714286

 $00:16:25.950 \longrightarrow 00:16:26.660$ those covariates.

NOTE Confidence: 0.852387335714286

 $00:16:26.660 \longrightarrow 00:16:28.788$ So next we decided to look at measures

00:16:28.788 --> 00:16:30.440 of stress related Physiology,

NOTE Confidence: 0.852387335714286

 $00:16:30.440 \longrightarrow 00:16:32.720$ including both metabolic and hormonal

NOTE Confidence: 0.852387335714286 00:16:32.720 --> 00:16:33.176 factors, NOTE Confidence: 0.852387335714286

 $00:16:33.180 \longrightarrow 00:16:35.539$ and see if they are also related

NOTE Confidence: 0.852387335714286

 $00:16:35.539 \longrightarrow 00:16:36.960$ to accelerated biological aging.

NOTE Confidence: 0.852387335714286 00:16:36.960 --> 00:16:38.790 Next slide. NOTE Confidence: 0.852387335714286

 $00:16:38.790 \longrightarrow 00:16:41.130$ So we first assess the relationship

NOTE Confidence: 0.852387335714286

 $00{:}16{:}41.130 \dashrightarrow 00{:}16{:}42.870$ between grim age acceleration in

NOTE Confidence: 0.852387335714286

 $00:16:42.870 \longrightarrow 00:16:44.688$ HP I8HP axis via the cortisol.

NOTE Confidence: 0.852387335714286

00:16:44.690 --> 00:16:45.935 The ACTH ratio,

NOTE Confidence: 0.852387335714286

 $00:16:45.935 \longrightarrow 00:16:48.840$ which is a measure of adrenal sensitivity.

NOTE Confidence: 0.852387335714286

 $00:16:48.840 \longrightarrow 00:16:51.016$ Now in this plot you can see a

NOTE Confidence: 0.852387335714286

 $00{:}16{:}51.016 \dashrightarrow 00{:}16{:}51.945$ significant negative correlation

NOTE Confidence: 0.852387335714286

 $00:16:51.945 \longrightarrow 00:16:53.925$ between adrenal sensitivity on the X

NOTE Confidence: 0.852387335714286

 $00:16:53.925 \longrightarrow 00:16:56.626$ axis and grim age acceleration on the Y axis,

 $00:16:56.630 \longrightarrow 00:16:58.810$ But this relationship becomes nonsignificant

NOTE Confidence: 0.852387335714286

 $00{:}16{:}58.810 \dashrightarrow 00{:}17{:}00.990$ when we account for covariates.

NOTE Confidence: 0.852387335714286

 $00:17:00.990 \longrightarrow 00:17:02.390$ This does seem to be appeared to be.

NOTE Confidence: 0.852387335714286

 $00:17:02.390 \longrightarrow 00:17:05.360$ This appears to be driven in part by a

NOTE Confidence: 0.852387335714286

 $00:17:05.360 \longrightarrow 00:17:07.281$ differential responses in men and women's.

NOTE Confidence: 0.852387335714286

 $00{:}17{:}07.281 \dashrightarrow 00{:}17{:}09.129$ When we remove sex is a covariant that.

NOTE Confidence: 0.852387335714286

 $00:17:09.130 \longrightarrow 00:17:11.680$ Their relationship is once again significant.

NOTE Confidence: 0.852387335714286 00:17:11.680 --> 00:17:13.830 Next slide. NOTE Confidence: 0.852387335714286

 $00:17:13.830 \longrightarrow 00:17:16.030$ We next examined insulin resistance,

NOTE Confidence: 0.852387335714286

00:17:16.030 --> 00:17:18.010 which is another physiologic process,

NOTE Confidence: 0.852387335714286

 $00{:}17{:}18.010 \dashrightarrow 00{:}17{:}19.711$ are related to stress and to do

NOTE Confidence: 0.852387335714286

00:17:19.711 --> 00:17:21.748 this we use the measurement Houma,

NOTE Confidence: 0.852387335714286

 $00:17:21.750 \longrightarrow 00:17:23.685$ which is calculated based on

NOTE Confidence: 0.852387335714286

00:17:23.685 --> 00:17:24.780 individuals glucose, insulin.

NOTE Confidence: 0.852387335714286

00:17:24.780 --> 00:17:27.660 Now in this plot you can see that Houma,

NOTE Confidence: 0.8497675

 $00{:}17{:}27.660 \dashrightarrow 00{:}17{:}30.065$ which increases as an individual's

00:17:30.065 --> 00:17:31.508 insulin resistance increases,

NOTE Confidence: 0.8497675

00:17:31.510 --> 00:17:32.911 is positively correlated

NOTE Confidence: 0.8497675

 $00:17:32.911 \longrightarrow 00:17:34.779$ with cream age acceleration.

NOTE Confidence: 0.8497675

00:17:34.780 --> 00:17:36.928 Now, unlike the cortisol ACTH ratio,

NOTE Confidence: 0.8497675

 $00{:}17{:}36.930 \dashrightarrow 00{:}17{:}38.666$ this relationship remains significant

NOTE Confidence: 0.8497675

00:17:38.666 --> 00:17:42.890 after accounting for ARCO very next slide.

NOTE Confidence: 0.8497675

00:17:42.890 --> 00:17:45.151 So thus far we've identified at least

NOTE Confidence: 0.8497675

 $00{:}17{:}45.151 \dashrightarrow 00{:}17{:}46.546$ one potential physiologic mechanism

NOTE Confidence: 0.8497675

 $00:17:46.546 \longrightarrow 00:17:48.736$ through which stress might influence aging

NOTE Confidence: 0.8497675

00:17:48.736 --> 00:17:50.859 through changes in insulin resistance,

NOTE Confidence: 0.8497675

 $00{:}17{:}50.860 \dashrightarrow 00{:}17{:}52.981$ as well as a potentially more complex

NOTE Confidence: 0.8497675

 $00:17:52.981 \longrightarrow 00:17:54.979$ story with adrenal sensitivity in sex.

NOTE Confidence: 0.8497675

00:17:54.980 --> 00:17:56.756 It's also worth noting that while

NOTE Confidence: 0.8497675

 $00:17:56.756 \longrightarrow 00:17:58.320$ accounting for both of these,

NOTE Confidence: 0.8497675

 $00:17:58.320 \longrightarrow 00:18:00.798$ we continue to see an independent

 $00:18:00.798 \longrightarrow 00:18:03.618$ effect of stress on aging as well.

NOTE Confidence: 0.8497675

 $00{:}18{:}03.620 \dashrightarrow 00{:}18{:}05.020$ Now, as I mentioned earlier,

NOTE Confidence: 0.8497675

 $00:18:05.020 \longrightarrow 00:18:06.640$ we don't just want to find

NOTE Confidence: 0.8497675

 $00:18:06.640 \longrightarrow 00:18:08.180$ ways how we're aging faster,

NOTE Confidence: 0.8497675

00:18:08.180 --> 00:18:09.938 but ways to protect against it,

NOTE Confidence: 0.8497675

 $00:18:09.940 \longrightarrow 00:18:11.975$ and one potential counter distress

NOTE Confidence: 0.8497675

 $00{:}18{:}11.975 \dashrightarrow 00{:}18{:}13.603$ would be psychological resilience.

NOTE Confidence: 0.8497675

 $00{:}18{:}13.610 \dashrightarrow 00{:}18{:}15.716$ So we next asked whether characteristics

NOTE Confidence: 0.8497675

 $00{:}18{:}15.716 \longrightarrow 00{:}18{:}18.244$ such as emotion regulation and self-control

NOTE Confidence: 0.8497675

00:18:18.244 --> 00:18:20.374 might alter the relationship between stress,

NOTE Confidence: 0.8497675

00:18:20.380 --> 00:18:22.996 Physiology, and aging.

NOTE Confidence: 0.8497675

 $00:18:22.996 \longrightarrow 00:18:24.740$ Next slide.

NOTE Confidence: 0.8497675

 $00{:}18{:}24.740 \dashrightarrow 00{:}18{:}26.581$ Well when we assess self control we

NOTE Confidence: 0.8497675

 $00{:}18{:}26.581 \dashrightarrow 00{:}18{:}28.794$ see that it moderates the relationship

NOTE Confidence: 0.8497675

 $00:18:28.794 \longrightarrow 00:18:31.069$ between stress and insulin resistance.

NOTE Confidence: 0.8497675

 $00:18:31.070 \longrightarrow 00:18:33.398$ In this plot you can see three lines

 $00:18:33.398 \longrightarrow 00:18:34.624$ representing the relationship between

NOTE Confidence: 0.8497675

 $00:18:34.624 \longrightarrow 00:18:37.304$ stress on the X axis in Houma on the

NOTE Confidence: 0.8497675

 $00{:}18{:}37.304 \rightarrow 00{:}18{:}39.044$ Y axis for individuals with good,

NOTE Confidence: 0.8497675

 $00:18:39.050 \longrightarrow 00:18:41.490$ fair or poor self control.

NOTE Confidence: 0.8497675

 $00:18:41.490 \longrightarrow 00:18:42.890$ You'll notice that the individual

NOTE Confidence: 0.8497675

 $00:18:42.890 \longrightarrow 00:18:43.730$ good self control.

NOTE Confidence: 0.8497675

00:18:43.730 --> 00:18:46.005 There's little effect of stress

NOTE Confidence: 0.8497675

00:18:46.005 --> 00:18:47.370 on insulin resistance,

NOTE Confidence: 0.8497675

 $00:18:47.370 \longrightarrow 00:18:48.945$ but in those with poor self control,

NOTE Confidence: 0.8497675

 $00:18:48.950 \longrightarrow 00:18:50.422$ there's a large effect,

NOTE Confidence: 0.8497675

 $00{:}18{:}50.422 \dashrightarrow 00{:}18{:}51.894$ and this moderating effective

NOTE Confidence: 0.8497675

00:18:51.894 --> 00:18:53.530 self control is significant.

NOTE Confidence: 0.8497675

 $00{:}18{:}53.530 \dashrightarrow 00{:}18{:}56.020$ Even we were accounting for covariance.

NOTE Confidence: 0.8497675

 $00:18:56.020 \longrightarrow 00:18:59.244$ Now one cover it. I do want to.

NOTE Confidence: 0.8497675

 $00:18:59.250 \longrightarrow 00:19:00.142$ A1 covariate.

 $00:19:00.142 \longrightarrow 00:19:03.710$ I'd like to point out there is BMI.

NOTE Confidence: 0.8497675

 $00:19:03.710 \longrightarrow 00:19:04.022$ Hey,

NOTE Confidence: 0.8497675

 $00:19:04.022 \longrightarrow 00:19:05.894$ well BMI is related to both

NOTE Confidence: 0.8497675

 $00:19:05.894 \longrightarrow 00:19:07.570$ stress and insulin resistance.

NOTE Confidence: 0.8497675

 $00:19:07.570 \longrightarrow 00:19:09.490$ I want to emphasize that this

NOTE Confidence: 0.8497675

 $00{:}19{:}09.490 \dashrightarrow 00{:}19{:}10.450$ relationship between stress,

NOTE Confidence: 0.8497675

 $00:19:10.450 \longrightarrow 00:19:12.405$ self control and insulin resistance

NOTE Confidence: 0.8497675

00:19:12.405 --> 00:19:14.360 is still significant after accounting

NOTE Confidence: 0.8497675

 $00:19:14.419 \longrightarrow 00:19:16.855$ for BMI and that self control is

NOTE Confidence: 0.8497675

00:19:16.855 --> 00:19:17.899 actually specifically moderating

NOTE Confidence: 0.8497675

 $00{:}19{:}17.960 \dashrightarrow 00{:}19{:}20.065$ the relationship between stress and

NOTE Confidence: 0.8497675

 $00:19:20.065 \longrightarrow 00:19:23.742$ insulin resistance, not stress and BMI.

NOTE Confidence: 0.8497675

00:19:23.742 --> 00:19:25.158 Next slide.

NOTE Confidence: 0.8497675

 $00:19:25.160 \longrightarrow 00:19:27.320$ So now we've identified at least

NOTE Confidence: 0.8497675

 $00:19:27.320 \longrightarrow 00:19:28.760$ one psychological resilience factor,

NOTE Confidence: 0.8497675

 $00:19:28.760 \longrightarrow 00:19:31.640$ self control that can influence stress

 $00:19:31.640 \longrightarrow 00:19:34.070$ related Physiology and thus aging.

NOTE Confidence: 0.8497675

 $00{:}19{:}34.070 \dashrightarrow 00{:}19{:}35.990$ But it was striking to us that there

NOTE Confidence: 0.8497675

 $00{:}19{:}35.990 \dashrightarrow 00{:}19{:}37.528$ still remains a significant independent

NOTE Confidence: 0.8497675

00:19:37.528 --> 00:19:39.208 effects of stress on grammage,

NOTE Confidence: 0.8497675

 $00:19:39.210 \longrightarrow 00:19:41.100$ so we access with their other

NOTE Confidence: 0.8497675

 $00:19:41.100 \longrightarrow 00:19:42.045$ psychological resilience factors.

NOTE Confidence: 0.8497675

00:19:42.050 --> 00:19:43.950 Might moderate this seemingly

NOTE Confidence: 0.8497675

 $00:19:43.950 \dashrightarrow 00:19:46.325$ independent effective stress on H.

NOTE Confidence: 0.8497675

00:19:46.330 --> 00:19:46.810 Excite NOTE Confidence: 0.861093793846154

 $00:19:49.070 \longrightarrow 00:19:51.485$ so next we asked whether emotion regulation

NOTE Confidence: 0.861093793846154

 $00:19:51.485 \dashrightarrow 00:19:53.810$ might be important for this relationship.

NOTE Confidence: 0.861093793846154

 $00:19:53.810 \longrightarrow 00:19:55.791$ When we examine the effects of emotion

NOTE Confidence: 0.861093793846154

 $00{:}19{:}55.791 \dashrightarrow 00{:}19{:}57.606$ regulation on the relationship between

NOTE Confidence: 0.861093793846154

00:19:57.606 --> 00:19:59.426 stress and grammage acceleration,

NOTE Confidence: 0.861093793846154

 $00:19:59.430 \longrightarrow 00:20:01.590$ we see a strong moderating effect.

 $00:20:01.590 \longrightarrow 00:20:03.830$ As you can see in the plot.

NOTE Confidence: 0.861093793846154

 $00{:}20{:}03.830 \to 00{:}20{:}05.690$ People with better emotion regulation

NOTE Confidence: 0.861093793846154

 $00:20:05.690 \longrightarrow 00:20:08.524$ as represented by the blue line leads

NOTE Confidence: 0.861093793846154

 $00:20:08.524 \longrightarrow 00:20:10.340$ have blunted relationship between

NOTE Confidence: 0.861093793846154

 $00:20:10.340 \longrightarrow 00:20:12.156$ stress and grammage acceleration,

NOTE Confidence: 0.861093793846154

00:20:12.160 --> 00:20:14.155 whereas poor emotion regulation is

NOTE Confidence: 0.861093793846154

 $00:20:14.155 \longrightarrow 00:20:16.818$ represented by the red line amplifies

NOTE Confidence: 0.861093793846154

 $00:20:16.818 \longrightarrow 00:20:20.290$ that relationship. Next slide.

NOTE Confidence: 0.861093793846154

00:20:20.290 --> 00:20:21.496 So going back to our model,

NOTE Confidence: 0.861093793846154

 $00:20:21.500 \longrightarrow 00:20:24.041$ we can think of stress is directly

NOTE Confidence: 0.861093793846154

 $00:20:24.041 \longrightarrow 00:20:26.359$ impacting age acceleration in a fashion

NOTE Confidence: 0.861093793846154

 $00:20:26.359 \longrightarrow 00:20:28.294$ that's moderated by emotion regulation.

NOTE Confidence: 0.861093793846154

00:20:28.300 --> 00:20:29.790 And after adding emotion regulation,

NOTE Confidence: 0.861093793846154

00:20:29.790 --> 00:20:31.920 stress does continue to impact aging

NOTE Confidence: 0.861093793846154

00:20:31.920 --> 00:20:33.340 through elevated insulin resistance,

NOTE Confidence: 0.861093793846154

 $00:20:33.340 \longrightarrow 00:20:35.937$ which again is influenced by self control.

 $00:20:35.940 \longrightarrow 00:20:38.140$ Next slide.

NOTE Confidence: 0.861093793846154

 $00:20:38.140 \longrightarrow 00:20:40.006$ So to bring these results together,

NOTE Confidence: 0.861093793846154

 $00:20:40.010 \longrightarrow 00:20:42.500$ we wanted to compare the

NOTE Confidence: 0.861093793846154

 $00:20:42.500 \longrightarrow 00:20:43.994$ contributions of stress,

NOTE Confidence: 0.861093793846154

 $00:20:44.000 \longrightarrow 00:20:45.776$ emotion regulation and insulin

NOTE Confidence: 0.861093793846154

 $00:20:45.776 \longrightarrow 00:20:48.440$ resistance to aging in the context

NOTE Confidence: 0.861093793846154

00:20:48.513 --> 00:20:50.768 of other more familiar variables.

NOTE Confidence: 0.861093793846154

 $00:20:50.770 \longrightarrow 00:20:51.550$ To do this,

NOTE Confidence: 0.861093793846154

 $00:20:51.550 \longrightarrow 00:20:52.590$ we used estimated marginal

NOTE Confidence: 0.861093793846154

 $00:20:52.590 \longrightarrow 00:20:54.090$ means in the linear model,

NOTE Confidence: 0.861093793846154

 $00:20:54.090 \longrightarrow 00:20:56.790$ incorporating all of our covariates are

NOTE Confidence: 0.861093793846154

 $00:20:56.790 \longrightarrow 00:20:59.165$ stress related Physiology factors and

NOTE Confidence: 0.861093793846154

 $00{:}20{:}59.165 \dashrightarrow 00{:}21{:}01.189$ our psychological resilience factors.

NOTE Confidence: 0.861093793846154

 $00:21:01.190 \longrightarrow 00:21:02.480$ And when we do this,

NOTE Confidence: 0.861093793846154

 $00:21:02.480 \longrightarrow 00:21:04.696$ we see that stress continues to have a

 $00:21:04.696 \longrightarrow 00:21:05.829$ significant relationship to grammage.

NOTE Confidence: 0.861093793846154

 $00:21:05.830 \longrightarrow 00:21:08.360$ That's moderated by emotion regulation.

NOTE Confidence: 0.861093793846154

 $00:21:08.360 \longrightarrow 00:21:10.800$ And it's worth noting that when we assess

NOTE Confidence: 0.861093793846154

 $00:21:10.800 \longrightarrow 00:21:13.089$ our model at poor emotion regulation.

NOTE Confidence: 0.861093793846154

 $00:21:13.090 \longrightarrow 00:21:15.040$ And there's a highly significant

NOTE Confidence: 0.861093793846154

 $00:21:15.040 \longrightarrow 00:21:16.990$ effect of stress on crymych.

NOTE Confidence: 0.861093793846154

00:21:16.990 --> 00:21:19.080 In these individuals, stress alone,

NOTE Confidence: 0.861093793846154

00:21:19.080 --> 00:21:21.168 independent of our covariates,

NOTE Confidence: 0.861093793846154

 $00:21:21.168 \longrightarrow 00:21:25.290$ has a strong impact on Grim Age's BMI.

NOTE Confidence: 0.861093793846154 00:21:25.290 --> 00:21:25.518 However, NOTE Confidence: 0.861093793846154

 $00{:}21{:}25.518 \dashrightarrow 00{:}21{:}27.114$ when we assess our models in those

NOTE Confidence: 0.861093793846154

00:21:27.114 --> 00:21:28.520 with good emotional regulation,

NOTE Confidence: 0.861093793846154

 $00{:}21{:}28.520 \dashrightarrow 00{:}21{:}30.806$ this relationship becomes

NOTE Confidence: 0.861093793846154

 $00:21:30.806 \longrightarrow 00:21:32.330$ entirely nonsignificant.

NOTE Confidence: 0.861093793846154

00:21:32.330 --> 00:21:33.102 Insulin resistance,

NOTE Confidence: 0.861093793846154

 $00:21:33.102 \longrightarrow 00:21:34.646$ which, as we've shown,

00:21:34.650 --> 00:21:36.365 is related to stress via self control,

NOTE Confidence: 0.861093793846154

 $00:21:36.370 \longrightarrow 00:21:39.110$ predicts a further increase in

NOTE Confidence: 0.861093793846154

 $00{:}21{:}39.110 \dashrightarrow 00{:}21{:}40.690$ inflammation cellarage next slide.

NOTE Confidence: 0.761497820555556

00:21:42.730 --> 00:21:45.490 So in summary, today I've shown you that

NOTE Confidence: 0.76149782055556

 $00:21:45.490 \longrightarrow 00:21:47.009$ cumulative stress predicts biological

NOTE Confidence: 0.761497820555556

 $00:21:47.009 \longrightarrow 00:21:49.289$ aging is measured by cream age,

NOTE Confidence: 0.761497820555556

 $00:21:49.290 \longrightarrow 00:21:51.789$ and this is not accounted for by

NOTE Confidence: 0.761497820555556

 $00:21:51.789 \longrightarrow 00:21:53.610$ demographic or behavioral covariates.

NOTE Confidence: 0.761497820555556

 $00:21:53.610 \longrightarrow 00:21:56.234$ We see that these interactions are at part

NOTE Confidence: 0.76149782055556

 $00:21:56.234 \longrightarrow 00:21:58.350$ mediated through insulin resistance and that

NOTE Confidence: 0.761497820555556

00:21:58.350 --> 00:22:00.790 adrenal sensitivity may also play a role.

NOTE Confidence: 0.761497820555556

 $00:22:00.790 \longrightarrow 00:22:02.975$ Remarkably, these interactions are highly

NOTE Confidence: 0.761497820555556

 $00{:}22{:}02.975 \dashrightarrow 00{:}22{:}04.723$ dependent on psychological resilience.

NOTE Confidence: 0.761497820555556

 $00:22:04.730 \longrightarrow 00:22:06.785$ Factors with strong self control

NOTE Confidence: 0.761497820555556

 $00:22:06.785 \longrightarrow 00:22:08.429$ blunting the relationship between

00:22:08.429 --> 00:22:10.545 stress and insulin resistance and

NOTE Confidence: 0.761497820555556

 $00:22:10.545 \longrightarrow 00:22:12.600$ strong emotion regulation dampening the

NOTE Confidence: 0.761497820555556

 $00:22:12.600 \longrightarrow 00:22:14.818$ direct effects of stress on scrimmage.

NOTE Confidence: 0.761497820555556

00:22:14.820 --> 00:22:18.183 Next slide. So looking forward,

NOTE Confidence: 0.761497820555556

 $00:22:18.183 \longrightarrow 00:22:20.150$ I would like to use the theoretical

NOTE Confidence: 0.761497820555556

00:22:20.205 --> 00:22:22.197 model we built the highlight potential,

NOTE Confidence: 0.761497820555556

00:22:22.200 --> 00:22:22.880 future directions,

NOTE Confidence: 0.761497820555556

 $00:22:22.880 \longrightarrow 00:22:24.240$ and possible interventions that

NOTE Confidence: 0.76149782055556

 $00:22:24.240 \longrightarrow 00:22:25.260$ could decrease age,

NOTE Confidence: 0.761497820555556

 $00:22:25.260 \longrightarrow 00:22:28.480$ acceleration in highly stressed populations.

NOTE Confidence: 0.761497820555556

 $00:22:28.480 \longrightarrow 00:22:30.300$ Looking at our biological factors,

NOTE Confidence: 0.761497820555556

 $00:22:30.300 \longrightarrow 00:22:33.036$ an obvious place to intervene is

NOTE Confidence: 0.76149782055556

 $00:22:33.036 \longrightarrow 00:22:34.404$ on insulin resistance.

NOTE Confidence: 0.761497820555556

00:22:34.410 --> 00:22:36.180 Metformin is actually being investigated

NOTE Confidence: 0.761497820555556

00:22:36.180 --> 00:22:38.739 as an anti-aging broke now is part of

NOTE Confidence: 0.761497820555556

00:22:38.739 --> 00:22:40.440 the team trial in future work could

00:22:40.498 --> 00:22:42.613 determine if it's effective specifically

NOTE Confidence: 0.761497820555556

00:22:42.613 --> 00:22:44.305 in highly stressed populations.

NOTE Confidence: 0.761497820555556

00:22:44.310 --> 00:22:46.000 Further studies might also clarify

NOTE Confidence: 0.761497820555556

00:22:46.000 --> 00:22:47.690 both the rollup adrenal sensitivity,

NOTE Confidence: 0.76149782055556

00:22:47.690 --> 00:22:49.910 but also potentially new neural for

NOTE Confidence: 0.761497820555556

 $00:22:49.910 \longrightarrow 00:22:51.917$ modal or cellular pathways that

NOTE Confidence: 0.761497820555556

 $00:22:51.917 \longrightarrow 00:22:53.777$ mediate this seemingly independent

NOTE Confidence: 0.761497820555556

 $00:22:53.777 \longrightarrow 00:22:56.102$ relationship between stress and aging,

NOTE Confidence: 0.761497820555556

00:22:56.110 --> 00:22:58.162 as well as the mechanisms through

NOTE Confidence: 0.76149782055556

 $00:22:58.162 \longrightarrow 00:22:59.530$ which emotion regulation is

NOTE Confidence: 0.761497820555556

 $00:22:59.592 \longrightarrow 00:23:01.218$ moderating this relationship.

NOTE Confidence: 0.761497820555556

 $00:23:01.220 \longrightarrow 00:23:03.400$ There's also the potential for

NOTE Confidence: 0.761497820555556

 $00{:}23{:}03.400 \dashrightarrow 00{:}23{:}04.708$ psychotherapeutic interventions that

NOTE Confidence: 0.761497820555556

 $00{:}23{:}04.708 \dashrightarrow 00{:}23{:}06.232$ prove psychological resilience to

NOTE Confidence: 0.76149782055556

 $00:23:06.232 \longrightarrow 00:23:08.332$ decrease the effects of stress on aging.

 $00:23:08.340 \longrightarrow 00:23:09.188$ For example,

NOTE Confidence: 0.76149782055556

 $00{:}23{:}09.188 \dashrightarrow 00{:}23{:}11.308$ there's evidence that mindfulness based

NOTE Confidence: 0.761497820555556

 $00:23:11.308 \longrightarrow 00:23:13.618$ interventions may improve emotion right now.

NOTE Confidence: 0.761497820555556

00:23:13.620 --> 00:23:16.604 And finally we can work for social changes

NOTE Confidence: 0.761497820555556

 $00:23:16.604 \longrightarrow 00:23:19.079$ to decrease environmental stressors.

NOTE Confidence: 0.761497820555556

00:23:19.080 --> 00:23:21.340 Societal changes that address poverty,

NOTE Confidence: 0.761497820555556 00:23:21.340 --> 00:23:21.688 racism, NOTE Confidence: 0.761497820555556

 $00{:}23{:}21.688 \rightarrow 00{:}23{:}23.776$ and other sources of trauma could

NOTE Confidence: 0.761497820555556

 $00{:}23{:}23.776 \dashrightarrow 00{:}23{:}25.661$ ultimately lead to decreases in

NOTE Confidence: 0.761497820555556

 $00:23:25.661 \longrightarrow 00:23:27.217$ lifetime stress and improvements

NOTE Confidence: 0.761497820555556

 $00:23:27.217 \longrightarrow 00:23:29.180$ in overall health and aging.

NOTE Confidence: 0.761497820555556

00:23:29.180 --> 00:23:29.818 And ultimately,

NOTE Confidence: 0.76149782055556

 $00:23:29.818 \longrightarrow 00:23:32.051$ this work could be extended beyond the

NOTE Confidence: 0.761497820555556

 $00:23:32.051 \longrightarrow 00:23:33.820$ healthy population to other groups,

NOTE Confidence: 0.761497820555556

00:23:33.820 --> 00:23:35.872 such as those with serious mental

NOTE Confidence: 0.761497820555556

 $00:23:35.872 \longrightarrow 00:23:38.543$ illness in whom stress and adversity are

00:23:38.543 --> 00:23:40.603 obviously a significant risk factor,

NOTE Confidence: 0.761497820555556

 $00:23:40.610 \longrightarrow 00:23:43.074$ and that might allow us to address that

NOTE Confidence: 0.761497820555556

00:23:43.074 --> 00:23:44.909 mortality gap I mentioned earlier.

NOTE Confidence: 0.761497820555556 00:23:44.910 --> 00:23:45.400 Next slide.

NOTE Confidence: 0.850858968

 $00{:}23{:}47.500 \dashrightarrow 00{:}23{:}48.984$ So finally I just like to thank

NOTE Confidence: 0.850858968

 $00:23:48.984 \longrightarrow 00:23:50.933$ the last Min family and the Lessman

NOTE Confidence: 0.850858968

 $00:23:50.933 \longrightarrow 00:23:52.503$ Foundation and the selection Committee

NOTE Confidence: 0.850858968

00:23:52.503 --> 00:23:54.292 for giving me the opportunity to talk

NOTE Confidence: 0.850858968

 $00:23:54.292 \longrightarrow 00:23:56.756$ to you about my work today. My mentors,

NOTE Confidence: 0.850858968

00:23:56.756 --> 00:23:59.030 including Ira Cheetos and Hypo Shoe,

NOTE Confidence: 0.850858968

00:23:59.030 --> 00:24:01.368 as well as neofolk woman who helped

NOTE Confidence: 0.850858968

 $00{:}24{:}01.368 \dashrightarrow 00{:}24{:}03.020$ tremen dously with stats and Albert

NOTE Confidence: 0.850858968

 $00{:}24{:}03.020 \dashrightarrow 00{:}24{:}05.372$ Higgins Chen who provided a lot of fruit.

NOTE Confidence: 0.850858968

 $00:24:05.380 \longrightarrow 00:24:08.005$ Early guidance on using aperture that clocks.

NOTE Confidence: 0.850858968

 $00:24:08.010 \longrightarrow 00:24:09.602$ But also like to thank the yell at

00:24:09.602 --> 00:24:11.290 RTP and residency and our funding,

NOTE Confidence: 0.850858968

 $00:24:11.290 \longrightarrow 00:24:13.036$ I'm happy to take any questions.

NOTE Confidence: 0.7438214

00:24:17.810 --> 00:24:19.944 Create. Thank you Zack,

NOTE Confidence: 0.7438214

 $00:24:19.944 \longrightarrow 00:24:21.470$ and I applaud both the quality of

NOTE Confidence: 0.851762478947368

 $00:24:21.520 \longrightarrow 00:24:22.755$ your science and the quality

NOTE Confidence: 0.851762478947368

 $00:24:22.755 \longrightarrow 00:24:23.743$ of your time control.

NOTE Confidence: 0.851762478947368

 $00:24:23.750 \longrightarrow 00:24:26.150$ That was 14 minutes 57 seconds,

NOTE Confidence: 0.851762478947368

 $00:24:26.150 \longrightarrow 00:24:27.956$ which is about as spot on

NOTE Confidence: 0.8615791825

 $00:24:27.970 \longrightarrow 00:24:28.938$ as I've ever seen.

NOTE Confidence: 0.896888041428571

 $00:24:30.480 \longrightarrow 00:24:32.520$ We do have time for a question or two

NOTE Confidence: 0.896888041428571

 $00:24:32.520 \longrightarrow 00:24:36.720$ before moving on to our next presentation.

NOTE Confidence: 0.896888041428571

 $00:24:36.720 \longrightarrow 00:24:38.216$ But how am I going to see if

NOTE Confidence: 0.896888041428571

00:24:38.216 --> 00:24:39.459 people are asking questions?

NOTE Confidence: 0.896888041428571

 $00:24:39.460 \longrightarrow 00:24:41.266$ Please raise your hand, use the zoom.

NOTE Confidence: 0.75846835

 $00:24:43.790 \longrightarrow 00:24:45.402$ Button to raise your hand. If you

NOTE Confidence: 0.75846835

 $00{:}24{:}45.402 \dashrightarrow 00{:}24{:}47.820$ have any questions for Zach. Debbie

 $00:24:48.620 \longrightarrow 00:24:51.170$ I I just want to say

NOTE Confidence: 0.899316912

 $00:24:51.170 \longrightarrow 00:24:52.870$ that was extremely clear.

NOTE Confidence: 0.899316912

 $00:24:52.870 \longrightarrow 00:24:54.970$ An in the world of epigenetics.

NOTE Confidence: 0.899316912

00:24:54.970 --> 00:24:56.602 I'm practically layperson Anzac.

NOTE Confidence: 0.899316912

 $00{:}24{:}56.602 \dashrightarrow 00{:}24{:}58.642$ You made this absolutely understandable

NOTE Confidence: 0.899316912

 $00:24:58.642 \longrightarrow 00:25:00.848$ an as a proponent of psychotherapy,

NOTE Confidence: 0.899316912

 $00:25:00.850 \longrightarrow 00:25:02.954$ I'd love to see how that fit in,

NOTE Confidence: 0.899316912

 $00:25:02.960 \longrightarrow 00:25:04.704$ and I found that this is the kind

NOTE Confidence: 0.899316912

 $00:25:04.704 \longrightarrow 00:25:05.988$ of research that our department

NOTE Confidence: 0.899316912

 $00:25:05.988 \longrightarrow 00:25:07.582$ is very proud to sponsor.

NOTE Confidence: 0.899316912

 $00:25:07.582 \longrightarrow 00:25:09.470$ So well done and well presented.

NOTE Confidence: 0.899316912

 $00:25:09.470 \longrightarrow 00:25:10.030$ Thank you.

NOTE Confidence: 0.915197714

 $00{:}25{:}12.310 \dashrightarrow 00{:}25{:}15.030$ Thank you and I am I.

NOTE Confidence: 0.915197714

 $00:25:15.030 \longrightarrow 00:25:16.950$ I'm looking forward to looking into

NOTE Confidence: 0.915197714

 $00:25:16.950 \longrightarrow 00:25:18.491$ these sort of psychotherapeutic

 $00:25:18.491 \longrightarrow 00:25:20.559$ interventions as to how.

NOTE Confidence: 0.915197714

 $00{:}25{:}20.560 {\:{\mbox{--}}}{\:{\mbox{>}}}\ 00{:}25{:}22.964$ We can use psychotherapeutic

NOTE Confidence: 0.915197714

 $00:25:22.964 \longrightarrow 00:25:24.166$ interventions too.

NOTE Confidence: 0.915197714

00:25:24.170 --> 00:25:25.322 Improve both physical health,

NOTE Confidence: 0.915197714

 $00:25:25.322 \longrightarrow 00:25:27.050$ as in addition to mental health.

NOTE Confidence: 0.9005376

00:25:35.990 --> 00:25:38.970 OK. Seeing no further questions right now,

NOTE Confidence: 0.862355242

 $00:25:38.970 \longrightarrow 00:25:40.657$ so we'll go on to our next

NOTE Confidence: 0.862355242

00:25:40.657 --> 00:25:41.845 presentation and perhaps have a

NOTE Confidence: 0.862355242

 $00:25:41.845 \longrightarrow 00:25:44.050$ little time for discussion at the end.

NOTE Confidence: 0.862355242

00:25:44.050 --> 00:25:47.287 So our second Co first place winner of

NOTE Confidence: 0.862355242

00:25:47.287 --> 00:25:50.052 the last minute work is Emily Olson,

NOTE Confidence: 0.862355242

 $00:25:50.060 \longrightarrow 00:25:53.138$ and to invite her I'm sorry to introduce her.

NOTE Confidence: 0.862355242

 $00:25:53.138 \longrightarrow 00:25:54.674$ I'm going to remember Tom Fernandez.

NOTE Confidence: 0.88532317

00:25:56.620 --> 00:25:57.996 Good morning everyone. I'm so

NOTE Confidence: 0.88532317

 $00:25:57.996 \longrightarrow 00:25:59.730$ happy for all the Lessmann

NOTE Confidence: 0.790711831538462

 $00:25:59.791 \longrightarrow 00:26:00.739$ award winners this

 $00:26:00.750 \longrightarrow 00:26:02.150$ year. So my Congrats to

NOTE Confidence: 0.7817538325

 $00:26:02.150 \longrightarrow 00:26:04.440$ you all. I'm in especially

NOTE Confidence: 0.84998075

 $00:26:04.450 \longrightarrow 00:26:08.560$ happy to introduce Doctor Emily often.

NOTE Confidence: 0.84998075

 $00:26:08.560 \longrightarrow 00:26:10.156$ There's a little bit of background.

NOTE Confidence: 0.84998075

 $00{:}26{:}10.160 \dashrightarrow 00{:}26{:}13.638$ Emily earned her MD and PhD in Human

NOTE Confidence: 0.84998075

 $00{:}26{:}13.638 \dashrightarrow 00{:}26{:}16.670$ and statistical genetics in 2016

NOTE Confidence: 0.893749286666667

00:26:16.680 --> 00:26:18.039 from Washington University

NOTE Confidence: 0.893749286666667

 $00:26:18.040 \longrightarrow 00:26:20.410$ in Saint Louis. We're

NOTE Confidence: 0.804724783333333

00:26:20.420 --> 00:26:22.660 very fortunate that that Emily

NOTE Confidence: 0.804724783333333

 $00:26:22.660 \longrightarrow 00:26:24.462$ matched into our Solnit integrated

NOTE Confidence: 0.804724783333333

00:26:24.462 --> 00:26:26.360 training program at that time,

NOTE Confidence: 0.804724783333333

 $00{:}26{:}26{:}26{:}360 \dashrightarrow 00{:}26{:}29{:}284$ and since then I have to say she

NOTE Confidence: 0.804724783333333

 $00:26:29.284 \longrightarrow 00:26:31.622$ is proven on every level to be

NOTE Confidence: 0.804724783333333

 $00{:}26{:}31.622 \dashrightarrow 00{:}26{:}34.250$ really a model clinician scientist.

NOTE Confidence: 0.804724783333333

 $00:26:34.250 \longrightarrow 00:26:36.351$ She's been incredibly productive

 $00:26:36.351 \longrightarrow 00:26:38.659$ with research during residency.

NOTE Confidence: 0.804724783333333

00:26:38.660 --> 00:26:39.998 She's leading several

NOTE Confidence: 0.804724783333333

00:26:40.000 --> 00:26:41.203 gene discovery projects,

NOTE Confidence: 0.804724783333333

00:26:41.203 --> 00:26:44.010 including the one you'll hear about today,

NOTE Confidence: 0.804724783333333

 $00:26:44.010 \longrightarrow 00:26:44.694$ but also others,

NOTE Confidence: 0.804724783333333

00:26:44.694 --> 00:26:47.100 but I hope you'll hear about in the future,

NOTE Confidence: 0.804724783333333

 $00:26:47.100 \longrightarrow 00:26:50.160$ and those include projects discovering

NOTE Confidence: 0.804724783333333

00:26:50.160 --> 00:26:52.204 new risk, genes for hair pulling,

NOTE Confidence: 0.804724783333333

 $00{:}26{:}52.204 \dashrightarrow 00{:}26{:}53.740$ and skin picking disorders.

NOTE Confidence: 0.804724783333333

00:26:53.740 --> 00:26:57.443 And ADHD. And just in summary,

NOTE Confidence: 0.8047247833333333

 $00:26:57.443 \longrightarrow 00:26:59.846$ you know Emily continues to amaze me with

NOTE Confidence: 0.804724783333333

 $00:26:59.846 \longrightarrow 00:27:01.646$ their ongoing research accomplishments.

NOTE Confidence: 0.804724783333333

 $00{:}27{:}01.650 \dashrightarrow 00{:}27{:}03.958$ Despite her busy clinical schedule.

NOTE Confidence: 0.804724783333333

 $00:27:03.960 \longrightarrow 00:27:05.074$ And I should also mention

NOTE Confidence: 0.804724783333333

 $00:27:05.074 \longrightarrow 00:27:06.610$ a busy family schedule.

NOTE Confidence: 0.804724783333333

 $00:27:06.610 \longrightarrow 00:27:08.350$ Emily and her husband have welcomed

 $00:27:08.350 \longrightarrow 00:27:10.640$ 2 new additions to their family

NOTE Confidence: 0.804724783333333

 $00{:}27{:}10.640 \dashrightarrow 00{:}27{:}13.982$ during her training and what a way to

NOTE Confidence: 0.804724783333333

 $00:27:13.982 \longrightarrow 00:27:15.770$ welcome me back from maternity leave.

NOTE Confidence: 0.804724783333333

 $00:27:15.770 \longrightarrow 00:27:17.240$ With this award today.

NOTE Confidence: 0.92045784

00:27:19.310 --> 00:27:21.716 I am certain that Emily, as a researcher,

NOTE Confidence: 0.896696033333333

 $00:27:21.720 \longrightarrow 00:27:24.175$ will continue to advance the field of

NOTE Confidence: 0.896696033333333

00:27:24.175 --> 00:27:25.891 psychiatric genetics for years to come,

NOTE Confidence: 0.896696033333333

 $00{:}27{:}25.891 \to 00{:}27{:}27.258$ and I really look forward to

NOTE Confidence: 0.896696033333333

 $00:27:27.258 \longrightarrow 00:27:29.238$ continuing to work with her as a

NOTE Confidence: 0.896696033333333

 $00{:}27{:}29.238 \dashrightarrow 00{:}27{:}30.686$ clinical and research colleague.

NOTE Confidence: 0.8966960333333333

 $00:27:30.686 \longrightarrow 00:27:33.640$ So thank you Emily. The floor is yours.

NOTE Confidence: 0.862712684285714

 $00:27:36.230 \longrightarrow 00:27:38.402$ Thank you Tom for that very

NOTE Confidence: 0.862712684285714

 $00:27:38.402 \longrightarrow 00:27:42.960$ kind introduction slide.

NOTE Confidence: 0.862712684285714

 $00:27:42.960 \longrightarrow 00:27:47.190$ So I don't have any disclosures today, fine.

NOTE Confidence: 0.862712684285714

 $00:27:47.190 \longrightarrow 00:27:49.621$ So before I get started, I just wanted

00:27:49.621 --> 00:27:52.063 to thank the Seymour Lessman award,

NOTE Confidence: 0.862712684285714

 $00{:}27{:}52.070 \dashrightarrow 00{:}27{:}54.482$ and although I never had the chance to meet

NOTE Confidence: 0.862712684285714

 $00:27:54.482 \longrightarrow 00:27:56.519$ Doctor Glassman from reading about him,

NOTE Confidence: 0.862712684285714

 $00:27:56.520 \longrightarrow 00:27:58.998$ I feel that his legacy has really

NOTE Confidence: 0.862712684285714

 $00:27:58.998 \longrightarrow 00:28:00.949$ influenced my training here at Yale.

NOTE Confidence: 0.862712684285714

00:28:00.950 --> 00:28:02.990 And I thought I would just

NOTE Confidence: 0.862712684285714

 $00:28:02.990 \longrightarrow 00:28:05.144$ highlight this quote written by the

NOTE Confidence: 0.862712684285714

 $00:28:05.144 \longrightarrow 00:28:06.954$ namesake of the residency program.

NOTE Confidence: 0.862712684285714

00:28:06.960 --> 00:28:08.236 I'm in Doctor Schoolnet,

NOTE Confidence: 0.862712684285714

 $00:28:08.236 \longrightarrow 00:28:10.721$ and so he writes in a scholarly

NOTE Confidence: 0.862712684285714

 $00:28:10.721 \longrightarrow 00:28:12.227$ and courageous way.

NOTE Confidence: 0.862712684285714

 $00{:}28{:}12.230 \to 00{:}28{:}14.696$ Dr Lessman repeatedly wrote about the

NOTE Confidence: 0.862712684285714

 $00:28:14.696 \longrightarrow 00:28:17.280$ importance of basic research and spoke out.

NOTE Confidence: 0.862712684285714

 $00:28:17.280 \longrightarrow 00:28:19.310$ For conditions that would promote

NOTE Confidence: 0.862712684285714

 $00:28:19.310 \longrightarrow 00:28:21.352$ opportunities for young investigators to

NOTE Confidence: 0.862712684285714

00:28:21.352 --> 00:28:23.554 develop their interests and capacities,

 $00:28:23.554 \longrightarrow 00:28:26.446$ and I'm so grateful to doctor

NOTE Confidence: 0.862712684285714

00:28:26.446 --> 00:28:28.060 Lessman doctor Solnit,

NOTE Confidence: 0.862712684285714

 $00:28:28.060 \longrightarrow 00:28:29.776$ my mentors and the many others

NOTE Confidence: 0.862712684285714

 $00:28:29.776 \longrightarrow 00:28:31.993$ who paved the way for me to be

NOTE Confidence: 0.862712684285714

 $00:28:31.993 \longrightarrow 00:28:33.499$ able to work on the research.

NOTE Confidence: 0.862712684285714

 $00:28:33.500 \longrightarrow 00:28:34.460$ Then going to present with

NOTE Confidence: 0.862712684285714

 $00:28:34.460 \longrightarrow 00:28:38.330$ you to you today slide.

NOTE Confidence: 0.862712684285714

 $00:28:38.330 \longrightarrow 00:28:40.328$ So today in the next I

NOTE Confidence: 0.862712684285714

00:28:40.328 --> 00:28:41.980 guess 14 minutes or so,

NOTE Confidence: 0.862712684285714

 $00:28:41.980 \longrightarrow 00:28:44.104$ I'm going to talk to you a little bit

NOTE Confidence: 0.862712684285714

00:28:44.104 --> 00:28:46.097 about our genomics work of childhood,

NOTE Confidence: 0.862712684285714

 $00:28:46.100 \longrightarrow 00:28:48.730$ anxiety disorders,

NOTE Confidence: 0.862712684285714

 $00:28:48.730 \longrightarrow 00:28:51.330$ and this makes up the most common class

NOTE Confidence: 0.862712684285714

 $00{:}28{:}51.330 \dashrightarrow 00{:}28{:}53.479$ of childhood psychiatric conditions.

NOTE Confidence: 0.862712684285714

 $00:28:53.480 \longrightarrow 00:28:55.016$ And for a long time we've known that

 $00:28:55.016 \longrightarrow 00:28:56.592$ genetic factors are important than

NOTE Confidence: 0.862712684285714

 $00:28:56.592 \longrightarrow 00:28:58.224$ we know this from family studies,

NOTE Confidence: 0.862712684285714

 $00:28:58.230 \longrightarrow 00:29:00.470$ and we know this from twin studies.

NOTE Confidence: 0.862712684285714

 $00:29:00.470 \longrightarrow 00:29:02.325$ And he studies show us that there

NOTE Confidence: 0.862712684285714

 $00:29:02.325 \longrightarrow 00:29:04.006$ is a genetic overlap between

NOTE Confidence: 0.862712684285714

 $00:29:04.006 \longrightarrow 00:29:05.662$ different anxiety disorders and

NOTE Confidence: 0.862712684285714

 $00{:}29{:}05.662 \dashrightarrow 00{:}29{:}07.706$ that the contribution of genetic

NOTE Confidence: 0.862712684285714

00:29:07.706 --> 00:29:09.316 factors to anxiety may change

NOTE Confidence: 0.862712684285714

 $00{:}29{:}09.316 \dashrightarrow 00{:}29{:}11.117$ over the course of development.

NOTE Confidence: 0.862712684285714

00:29:11.117 --> 00:29:12.131 And specifically,

NOTE Confidence: 0.862712684285714

 $00{:}29{:}12.131 \dashrightarrow 00{:}29{:}14.666$ there's some evidence there's a.

NOTE Confidence: 0.862712684285714

 $00:29:14.670 \longrightarrow 00:29:17.592$ There's a larger genetic contribution to

NOTE Confidence: 0.862712684285714

00:29:17.592 --> 00:29:20.820 anxiety that develops in early childhood,

NOTE Confidence: 0.862712684285714

 $00:29:20.820 \longrightarrow 00:29:23.880$ and so this highlights the discovery

NOTE Confidence: 0.862712684285714

 $00:29:23.880 \longrightarrow 00:29:26.485$ potential of genomic investigations that

NOTE Confidence: 0.862712684285714

 $00:29:26.485 \longrightarrow 00:29:31.270$ focus on childhood anxiety disorders slide.

 $00:29:31.270 \longrightarrow 00:29:32.890$ And so we know that genetic

NOTE Confidence: 0.862712684285714

 $00:29:32.890 \longrightarrow 00:29:33.700$ factors are important,

NOTE Confidence: 0.862712684285714

 $00:29:33.700 \longrightarrow 00:29:35.905$ but something that's been harder

NOTE Confidence: 0.862712684285714

 $00:29:35.905 \longrightarrow 00:29:37.669$ for scientists until recently

NOTE Confidence: 0.862712684285714

00:29:37.669 --> 00:29:40.080 is finding specific risk genes.

NOTE Confidence: 0.862712684285714

 $00:29:40.080 \longrightarrow 00:29:41.960$ And when we think about

NOTE Confidence: 0.862712684285714

00:29:41.960 --> 00:29:43.088 identifying druggable targets,

NOTE Confidence: 0.862712684285714

00:29:43.090 --> 00:29:44.950 this process of finding risk

NOTE Confidence: 0.862712684285714

 $00:29:44.950 \longrightarrow 00:29:46.438$ streams is really important.

NOTE Confidence: 0.862712684285714

00:29:46.440 --> 00:29:48.192 And it's only been in the

NOTE Confidence: 0.862712684285714

 $00:29:48.192 \longrightarrow 00:29:49.630$ last five years or so.

NOTE Confidence: 0.862712684285714

 $00:29:49.630 \longrightarrow 00:29:51.590$ With Genome wide association studies

NOTE Confidence: 0.862712684285714

 $00{:}29{:}51.590 \dashrightarrow 00{:}29{:}54.464$ that a few common variants have been

NOTE Confidence: 0.862712684285714

 $00{:}29{:}54.464 {\:\dashrightarrow\:} 00{:}29{:}56.579$ associated with anxiety disorders and

NOTE Confidence: 0.862712684285714

00:29:56.579 --> 00:29:58.983 actually the largest of these studies

 $00:29:58.983 \longrightarrow 00:30:01.468$ was led by Daniel Levy Angelica learner.

NOTE Confidence: 0.862712684285714

 $00:30:01.470 \longrightarrow 00:30:02.790$ Here at Yale.

NOTE Confidence: 0.862712684285714

00:30:02.790 --> 00:30:05.430 But in addition to common variance,

NOTE Confidence: 0.862712684285714

 $00:30:05.430 \longrightarrow 00:30:08.052$ it's also likely that rare variants

NOTE Confidence: 0.862712684285714

00:30:08.052 --> 00:30:11.259 influence the risk of anxiety disorders,

NOTE Confidence: 0.862712684285714

 $00:30:11.260 \longrightarrow 00:30:15.070$ and to study these we need

NOTE Confidence: 0.862712684285714

 $00:30:15.070 \longrightarrow 00:30:17.610$ DNA sequencing studies slide.

NOTE Confidence: 0.862712684285714

00:30:17.610 --> 00:30:19.455 And one approach that's been

NOTE Confidence: 0.862712684285714

 $00{:}30{:}19.455 \dashrightarrow 00{:}30{:}21.750$ especially fruitful in the field of

NOTE Confidence: 0.862712684285714

00:30:21.750 --> 00:30:23.665 child psychiatry is DNA sequencing.

NOTE Confidence: 0.862712684285714

 $00{:}30{:}23.670 \dashrightarrow 00{:}30{:}25.950$ Studies of parent child trios,

NOTE Confidence: 0.862712684285714

 $00:30:25.950 \longrightarrow 00:30:28.926$ where the child is impacted by the disorder.

NOTE Confidence: 0.862712684285714

00:30:28.930 --> 00:30:32.090 So since all of us inherit half of our DNA,

NOTE Confidence: 0.862712684285714

 $00:30:32.090 \longrightarrow 00:30:34.190$ in theory from our parents,

NOTE Confidence: 0.862712684285714

 $00:30:34.190 \longrightarrow 00:30:37.095$ this process can allow us to identify

NOTE Confidence: 0.862712684285714

 $00:30:37.095 \longrightarrow 00:30:38.837$ rare variants associated with

 $00:30:38.837 \longrightarrow 00:30:40.957$ the condition that are inherited,

NOTE Confidence: 0.862712684285714

00:30:40.960 --> 00:30:44.308 but also new or DENOVO mutations

NOTE Confidence: 0.862712684285714

 $00:30:44.308 \longrightarrow 00:30:46.550$ that are specific only found

NOTE Confidence: 0.862712684285714

 $00:30:46.550 \longrightarrow 00:30:48.230$ in the child and not found.

NOTE Confidence: 0.862712684285714

 $00:30:48.230 \longrightarrow 00:30:49.370$ In the parents.

NOTE Confidence: 0.862712684285714

 $00:30:49.370 \longrightarrow 00:30:52.955$ And all of us have about 50 to 100

NOTE Confidence: 0.862712684285714

 $00:30:52.955 \longrightarrow 00:30:55.040$ knew or de Novo mutations.

NOTE Confidence: 0.862712684285714

 $00:30:55.040 \longrightarrow 00:30:57.950$ And when these occur within genes,

NOTE Confidence: 0.86865248375

 $00{:}30{:}57.950 \dashrightarrow 00{:}31{:}00.572$ they can actually impact the resulting

NOTE Confidence: 0.86865248375

 $00:31:00.572 \longrightarrow 00:31:02.578$ protein function. Anhava impact

NOTE Confidence: 0.86865248375

 $00:31:02.578 \longrightarrow 00:31:07.470$ on brain function as well slide.

NOTE Confidence: 0.86865248375

00:31:07.470 --> 00:31:09.619 And so this approach of sequencing parent,

NOTE Confidence: 0.86865248375

 $00{:}31{:}09.620 \dashrightarrow 00{:}31{:}12.284$ child trios an looking for these de Novo

NOTE Confidence: 0.86865248375

 $00:31:12.284 \longrightarrow 00:31:15.128$ variants in order to find risk genes was

NOTE Confidence: 0.86865248375

 $00:31:15.128 \longrightarrow 00:31:17.628$ initially pioneered in the field of autism.

 $00:31:17.630 \longrightarrow 00:31:19.058$ In the first study,

NOTE Confidence: 0.86865248375

 $00:31:19.058 \longrightarrow 00:31:22.525$ only had about 200 trios and they were able

NOTE Confidence: 0.86865248375

00:31:22.525 --> 00:31:25.319 to find a high confidence Christine SCN,

NOTE Confidence: 0.86865248375

00:31:25.319 --> 00:31:28.100 2A, which is now continues to be one of

NOTE Confidence: 0.86865248375

 $00:31:28.175 \longrightarrow 00:31:31.263$ the most well studied risk genes for autism.

NOTE Confidence: 0.86865248375

00:31:31.270 --> 00:31:32.710 But since that time,

NOTE Confidence: 0.86865248375

 $00:31:32.710 \longrightarrow 00:31:34.510$ now they've sequenced thousands of

NOTE Confidence: 0.86865248375

 $00:31:34.510 \longrightarrow 00:31:36.866$ it trees an I'm highlighting here.

NOTE Confidence: 0.86865248375

00:31:36.870 --> 00:31:39.446 A recent paper where they've now found over

NOTE Confidence: 0.86865248375

 $00:31:39.446 \longrightarrow 00:31:42.200$ 100 high confidence risk genes for autism.

NOTE Confidence: 0.86865248375 00:31:42.200 --> 00:31:44.490 Slide. NOTE Confidence: 0.86865248375

00:31:44.490 --> 00:31:47.227 And this is important because these risk

NOTE Confidence: 0.86865248375

 $00:31:47.227 \longrightarrow 00:31:50.088$ genes are already impacting clinical care.

NOTE Confidence: 0.86865248375

00:31:50.090 --> 00:31:53.060 So for families just knowing why

NOTE Confidence: 0.86865248375

00:31:53.060 --> 00:31:56.250 their child has autism is important,

NOTE Confidence: 0.86865248375

 $00:31:56.250 \longrightarrow 00:31:58.475$ understanding the likelihood of other

 $00:31:58.475 \longrightarrow 00:32:01.130$ family members being impacted and some

NOTE Confidence: 0.86865248375

 $00:32:01.130 \dashrightarrow 00:32:03.308$ of these risk genes are associated

NOTE Confidence: 0.86865248375

 $00:32:03.308 \longrightarrow 00:32:04.899$ with other medical comorbidities

NOTE Confidence: 0.86865248375

 $00:32:04.899 \longrightarrow 00:32:07.479$ that impact clinical care as well,

NOTE Confidence: 0.86865248375

 $00:32:07.480 \longrightarrow 00:32:10.896$ and so this approach was pioneered in autism,

NOTE Confidence: 0.86865248375

 $00:32:10.900 \longrightarrow 00:32:14.080$ but more recently it's been shown.

NOTE Confidence: 0.86865248375

 $00:32:14.080 \longrightarrow 00:32:17.674$ To have discovery potential and several

NOTE Confidence: 0.86865248375

 $00:32:17.674 \longrightarrow 00:32:20.070$ other psychiatric conditions slide.

NOTE Confidence: 0.86865248375

00:32:20.070 --> 00:32:21.780 And so here I'm just highlighting

NOTE Confidence: 0.86865248375

 $00:32:21.780 \longrightarrow 00:32:23.768$ two papers led by my mentor Tom,

NOTE Confidence: 0.86865248375

00:32:23.770 --> 00:32:25.826 that use this approach to find risk genes,

NOTE Confidence: 0.86865248375

 $00{:}32{:}25.830 \dashrightarrow 00{:}32{:}30.348$ interet disorder and OC D sign.

NOTE Confidence: 0.86865248375

 $00{:}32{:}30.350 \dashrightarrow 00{:}32{:}32.317$ And so our goal really was trying

NOTE Confidence: 0.86865248375

 $00:32:32.317 \longrightarrow 00:32:34.881$ to use this approach to see if we

NOTE Confidence: 0.86865248375

00:32:34.881 --> 00:32:36.536 could similarly find risk genes

00:32:36.602 --> 00:32:38.538 in childhood anxiety disorders.

NOTE Confidence: 0.86865248375

 $00:32:38.540 \longrightarrow 00:32:41.109$ So we collaborated with the program for

NOTE Confidence: 0.86865248375

00:32:41.109 --> 00:32:43.688 Anxiety disorders at the Child Study Center,

NOTE Confidence: 0.86865248375

 $00:32:43.690 \longrightarrow 00:32:45.394$ and I want to give a big thank

NOTE Confidence: 0.86865248375

 $00:32:45.394 \longrightarrow 00:32:46.810$ you to Wendy Silverman.

NOTE Confidence: 0.86865248375

00:32:46.810 --> 00:32:48.690 Annelie Liebowitz,

NOTE Confidence: 0.86865248375

 $00:32:48.690 \longrightarrow 00:32:51.195$ who let our recruitment and clinical

NOTE Confidence: 0.86865248375

 $00:32:51.195 \longrightarrow 00:32:53.290$ assessments and gave me the opportunity

NOTE Confidence: 0.938355074

 $00:32:53.300 \longrightarrow 00:32:54.780$ to work on this project.

NOTE Confidence: 0.880598765384615

 $00:32:55.580 \longrightarrow 00:32:58.240$ So we recruited children who were presenting

NOTE Confidence: 0.880598765384615

 $00:32:58.240 \longrightarrow 00:33:00.670$ with a primary concern of anxiety.

NOTE Confidence: 0.880598765384615

 $00:33:00.670 \longrightarrow 00:33:02.180$ In both of their parents,

NOTE Confidence: 0.880598765384615

 $00:33:02.180 \longrightarrow 00:33:05.240$ we collected saliva for DNA analysis

NOTE Confidence: 0.880598765384615

 $00:33:05.240 \longrightarrow 00:33:08.179$ and all families completed the aidas.

NOTE Confidence: 0.880598765384615

 $00:33:08.180 \longrightarrow 00:33:11.130$ The anxiety disorder interview schedule

NOTE Confidence: 0.880598765384615

 $00{:}33{:}11.130 \dashrightarrow 00{:}33{:}14.080$ that assesses for anxiety disorders

00:33:14.159 --> 00:33:17.069 and commonly Co occurring conditions.

NOTE Confidence: 0.880598765384615

 $00:33:17.070 \longrightarrow 00:33:19.560$ We then conducted high coverage

NOTE Confidence: 0.880598765384615

 $00:33:19.560 \longrightarrow 00:33:22.510$ whole exome sequencing of at 76

NOTE Confidence: 0.880598765384615

00:33:22.510 --> 00:33:24.810 parent child trios with anxiety,

NOTE Confidence: 0.880598765384615

 $00:33:24.810 \longrightarrow 00:33:28.532$ and we compared this to 225 controls

NOTE Confidence: 0.880598765384615

 $00:33:28.532 \longrightarrow 00:33:30.779$ and we did a variety of quality

NOTE Confidence: 0.880598765384615

 $00:33:30.779 \longrightarrow 00:33:33.016$ control checks on our sequencing data.

NOTE Confidence: 0.880598765384615

 $00{:}33{:}33.020 \dashrightarrow 00{:}33{:}37.840$ And we ended up comparing 65 trios to

NOTE Confidence: 0.880598765384615

00:33:37.840 --> 00:33:41.490 222 previously sequence control trios.

NOTE Confidence: 0.880598765384615 00:33:41.490 --> 00:33:43.680 Next slide.

NOTE Confidence: 0.880598765384615

 $00:33:43.680 \longrightarrow 00:33:46.760$ So here are the characteristics of the

NOTE Confidence: 0.880598765384615

 $00:33:46.760 \longrightarrow 00:33:49.178$ 68 children with anxiety disorders that

NOTE Confidence: 0.880598765384615

 $00{:}33{:}49.178 \dashrightarrow 00{:}33{:}51.963$ we ended up including in our Dinovo

NOTE Confidence: 0.880598765384615

 $00:33:51.963 \longrightarrow 00:33:54.177$ analysis and what I'm highlighting here

NOTE Confidence: 0.880598765384615

 $00:33:54.177 \longrightarrow 00:33:57.045$ in the red box is that many of these

00:33:57.045 --> 00:34:00.429 children met criteria for several.

NOTE Confidence: 0.880598765384615

00:34:00.430 --> 00:34:02.860 Anxiety disorders and this is really

NOTE Confidence: 0.880598765384615

 $00:34:02.860 \longrightarrow 00:34:05.309$ typical of clinical samples and anxiety.

NOTE Confidence: 0.880598765384615

 $00:34:05.310 \longrightarrow 00:34:07.932$ So the most common disorders were

NOTE Confidence: 0.880598765384615

00:34:07.932 --> 00:34:09.243 generalized anxiety disorder,

NOTE Confidence: 0.880598765384615

00:34:09.250 --> 00:34:10.780 social phobia,

NOTE Confidence: 0.880598765384615 00:34:10.780 --> 00:34:11.545 separation,

NOTE Confidence: 0.880598765384615

00:34:11.545 --> 00:34:16.135 anxiety disorder and specific phobia slide.

NOTE Confidence: 0.880598765384615

00:34:16.140 --> 00:34:18.230 And in our genomic analysis,

NOTE Confidence: 0.880598765384615

 $00:34:18.230 \longrightarrow 00:34:20.715$ we focused on rare de Novo variants

NOTE Confidence: 0.880598765384615

 $00:34:20.715 \longrightarrow 00:34:22.615$ that were thought to influence

NOTE Confidence: 0.880598765384615

 $00:34:22.615 \longrightarrow 00:34:24.530$ the coding region of genes,

NOTE Confidence: 0.880598765384615

 $00:34:24.530 \longrightarrow 00:34:26.846$ and our hypothesis was based on

NOTE Confidence: 0.880598765384615

 $00:34:26.846 \longrightarrow 00:34:28.886$ studies of other neuro psychiatric

NOTE Confidence: 0.880598765384615

 $00:34:28.886 \longrightarrow 00:34:31.418$ conditions and that we thought we

NOTE Confidence: 0.880598765384615

 $00{:}34{:}31.418 \dashrightarrow 00{:}34{:}34.084$ would find an enrichment of these

00:34:34.084 --> 00:34:36.299 Sonoma de Novo damaging mutations

NOTE Confidence: 0.880598765384615

 $00{:}34{:}36.299 \dashrightarrow 00{:}34{:}39.869$ in cases versus controls slide.

NOTE Confidence: 0.889142758333333

 $00:34:42.510 \longrightarrow 00:34:45.258$ And So what we found here

NOTE Confidence: 0.889142758333333

 $00:34:45.258 \longrightarrow 00:34:47.870$ what I'm showing here in red.

NOTE Confidence: 0.889142758333333

 $00:34:47.870 \longrightarrow 00:34:50.089$ Are the anxiety cases an in blue?

NOTE Confidence: 0.889142758333333

 $00{:}34{:}50.090 \dashrightarrow 00{:}34{:}52.274$ Are the controls and I'm showing you

NOTE Confidence: 0.889142758333333

 $00:34:52.274 \longrightarrow 00:34:54.682$ that there is an enrichment of these

NOTE Confidence: 0.889142758333333

 $00{:}34{:}54.682 \dashrightarrow 00{:}34{:}56.782$ damaging de Novo mutations and so

NOTE Confidence: 0.889142758333333

 $00:34:56.848 \longrightarrow 00:34:59.081$ this shows for the first time that

NOTE Confidence: 0.889142758333333

00:34:59.081 --> 00:35:01.668 this approach of focusing on de Novo

NOTE Confidence: 0.889142758333333

00:35:01.668 --> 00:35:04.110 variants in anxiety has the potential

NOTE Confidence: 0.889142758333333

 $00:35:04.186 \longrightarrow 00:35:06.712$ to identify risk genes and these

NOTE Confidence: 0.889142758333333

 $00{:}35{:}06.712 \dashrightarrow 00{:}35{:}08.882$ damaging variants that are enriched

NOTE Confidence: 0.889142758333333

 $00:35:08.882 \longrightarrow 00:35:11.731$ in cases compared to the controls are

NOTE Confidence: 0.889142758333333

 $00:35:11.731 \longrightarrow 00:35:13.945$ thought to alter protein functions.

 $00:35:13.945 \longrightarrow 00:35:15.805$ So specifically here we're

NOTE Confidence: 0.889142758333333

 $00:35:15.805 \longrightarrow 00:35:17.850$ focusing on damaging variance.

NOTE Confidence: 0.889142758333333

 $00:35:17.850 \longrightarrow 00:35:19.680$ That are likely Jinja struct?

NOTE Confidence: 0.889142758333333

 $00:35:19.680 \longrightarrow 00:35:22.564$ If so, these may introduce a stop

NOTE Confidence: 0.889142758333333

 $00:35:22.564 \longrightarrow 00:35:25.258$ codon early in the gene cause a

NOTE Confidence: 0.889142758333333

 $00:35:25.260 \longrightarrow 00:35:27.512$ frameshift insertion or deletion

NOTE Confidence: 0.889142758333333

 $00:35:27.512 \longrightarrow 00:35:30.890$ or alter a critical splice site.

NOTE Confidence: 0.889142758333333

 $00{:}35{:}30.890 \dashrightarrow 00{:}35{:}33.310$ We also included missense variants

NOTE Confidence: 0.889142758333333

 $00:35:33.310 \longrightarrow 00:35:37.122$ that may change in amino acid that is

NOTE Confidence: 0.889142758333333

 $00:35:37.122 \longrightarrow 00:35:40.167$ predicted to be damaging of the protein.

NOTE Confidence: 0.889142758333333 00:35:40.170 --> 00:35:40.522 Uhm? NOTE Confidence: 0.889142758333333

00:35:40.522 --> 00:35:40.874 So.

NOTE Confidence: 0.889142758333333

 $00:35:40.874 \longrightarrow 00:35:43.338$ I guess I just want to highlight

NOTE Confidence: 0.889142758333333

 $00:35:43.338 \longrightarrow 00:35:45.529$ that this was very exciting,

NOTE Confidence: 0.889142758333333

 $00:35:45.530 \longrightarrow 00:35:48.127$ that even for this common class of

NOTE Confidence: 0.889142758333333

 $00{:}35{:}48.127 \dashrightarrow 00{:}35{:}49.960$ conditions for anxiety disorders,

 $00:35:49.960 \longrightarrow 00:35:52.445$ we still see this enrichment

NOTE Confidence: 0.889142758333333

00:35:52.445 --> 00:35:54.930 of de Novo variance slide.

NOTE Confidence: 0.862579132333333

 $00:35:57.180 \longrightarrow 00:35:59.600$ And what we can do is we can look at

NOTE Confidence: 0.862579132333333

00:35:59.674 --> 00:36:02.264 the list of genes that have damaging

NOTE Confidence: 0.862579132333333

 $00:36:02.264 \longrightarrow 00:36:04.519$ mutations in these anxiety cases and

NOTE Confidence: 0.862579132333333

00:36:04.519 --> 00:36:07.018 see if they overlap with risk genes

NOTE Confidence: 0.862579132333333

 $00:36:07.020 \longrightarrow 00:36:09.670$ for other nuro psychiatric conditions.

NOTE Confidence: 0.862579132333333

 $00{:}36{:}09.670 \dashrightarrow 00{:}36{:}11.340$ And I'm highlighting here the

NOTE Confidence: 0.862579132333333

 $00:36:11.340 \longrightarrow 00:36:13.260$ two dream jeans that did so.

NOTE Confidence: 0.862579132333333

 $00{:}36{:}13.260 \dashrightarrow 00{:}36{:}16.050$ The first gene CAC N A1A in codes of

NOTE Confidence: 0.862579132333333

00:36:16.050 --> 00:36:17.979 voltage gated calcium channel and

NOTE Confidence: 0.862579132333333

 $00{:}36{:}17.979 \dashrightarrow 00{:}36{:}20.864$ this has been identified as a risk

NOTE Confidence: 0.8625791323333333

 $00{:}36{:}20.864 \dashrightarrow 00{:}36{:}22.808$ gene for developmental disorders

NOTE Confidence: 0.862579132333333

 $00:36:22.808 \longrightarrow 00:36:25.238$ in denovo sequencing studies as

NOTE Confidence: 0.862579132333333

 $00:36:25.240 \longrightarrow 00:36:27.900$ well as epileptic encephalopathies.

 $00:36:27.900 \longrightarrow 00:36:31.655$ The second gene is a regulatory

NOTE Confidence: 0.862579132333333

00:36:31.655 --> 00:36:34.580 subunit of protein phosphatase 2A,

NOTE Confidence: 0.862579132333333

 $00:36:34.580 \longrightarrow 00:36:38.000$ and this has been associated with

NOTE Confidence: 0.862579132333333

 $00:36:38.000 \longrightarrow 00:36:40.280$ developmental disorders as well

NOTE Confidence: 0.862579132333333

 $00:36:40.369 \longrightarrow 00:36:44.830$ as intellectual disability slide.

NOTE Confidence: 0.862579132333333

00:36:44.830 --> 00:36:47.798 But here in our cohort we're finding

NOTE Confidence: 0.862579132333333

 $00:36:47.798 \longrightarrow 00:36:50.087$ damaging mutations in these genes

NOTE Confidence: 0.862579132333333

 $00:36:50.087 \longrightarrow 00:36:52.437$ and individuals who have anxiety.

NOTE Confidence: 0.862579132333333

00:36:52.440 --> 00:36:55.030 They don't have any known history of

NOTE Confidence: 0.862579132333333

00:36:55.030 --> 00:36:56.790 neurologic or neurodevelopmental conditions,

NOTE Confidence: 0.8625791323333333

 $00{:}36{:}56.790 \dashrightarrow 00{:}36{:}58.582$ and so this really gets at this

NOTE Confidence: 0.862579132333333

00:36:58.582 --> 00:37:00.831 idea of Pleo tropi wear jeans with

NOTE Confidence: 0.862579132333333

 $00:37:00.831 \longrightarrow 00:37:02.961$ damaging variants may lead to different

NOTE Confidence: 0.862579132333333

 $00:37:03.022 \longrightarrow 00:37:04.766$ clinical manifestations in different

NOTE Confidence: 0.862579132333333

 $00:37:04.766 \longrightarrow 00:37:06.946$ individuals and this is something

NOTE Confidence: 0.862579132333333

 $00:37:06.946 \longrightarrow 00:37:11.318$ that we're continuing to explore.

00:37:11.318 --> 00:37:12.300 Slide.

NOTE Confidence: 0.862579132333333

 $00:37:12.300 \longrightarrow 00:37:14.604$ So we can also use this list of

NOTE Confidence: 0.862579132333333

 $00:37:14.604 \longrightarrow 00:37:16.734$ genes with damaging mutations to

NOTE Confidence: 0.862579132333333

00:37:16.734 --> 00:37:18.710 conduct exploratory pathway analysis

NOTE Confidence: 0.862579132333333

 $00:37:18.710 \longrightarrow 00:37:21.560$ by looking at whether these genes

NOTE Confidence: 0.862579132333333

 $00:37:21.560 \longrightarrow 00:37:23.720$ cluster in certain pathways more

NOTE Confidence: 0.862579132333333

 $00:37:23.720 \longrightarrow 00:37:26.078$ than might be expected by chance.

NOTE Confidence: 0.862579132333333

 $00:37:26.078 \longrightarrow 00:37:29.099$ And here I'm showing all of the gene

NOTE Confidence: 0.862579132333333

 $00{:}37{:}29.099 \dashrightarrow 00{:}37{:}31.598$ ontology based sets that have a Q

NOTE Confidence: 0.862579132333333

 $00{:}37{:}31.598 \dashrightarrow 00{:}37{:}34.140$ value less than .05 and the darker

NOTE Confidence: 0.862579132333333

 $00:37:34.140 \longrightarrow 00:37:35.940$ red indicates more significance and

NOTE Confidence: 0.862579132333333

 $00:37:35.940 \longrightarrow 00:37:37.951$ the bigger circle indicates that

NOTE Confidence: 0.862579132333333

 $00{:}37{:}37.951 \dashrightarrow 00{:}37{:}40.333$ more genes are contributing and you

NOTE Confidence: 0.862579132333333

 $00:37:40.399 \longrightarrow 00:37:42.506$ can see here that the top pathway,

NOTE Confidence: 0.862579132333333

 $00:37:42.510 \longrightarrow 00:37:44.897$ which is the darkest red in terms

 $00:37:44.897 \longrightarrow 00:37:46.378$ of significance is glutamatergic

NOTE Confidence: 0.862579132333333

 $00:37:46.378 \longrightarrow 00:37:49.129$ synapse and so this is kind of

NOTE Confidence: 0.862579132333333

 $00{:}37{:}49.129 \dashrightarrow 00{:}37{:}51.380$ consistent with the potential role

NOTE Confidence: 0.862579132333333

 $00:37:51.380 \longrightarrow 00:37:53.240$ of glutamate neurotransmission in

NOTE Confidence: 0.862579132333333

 $00:37:53.240 \longrightarrow 00:37:57.240$ the development of anxiety.

NOTE Confidence: 0.862579132333333 00:37:57.240 --> 00:37:57.938 Uhm? NOTE Confidence: 0.862579132333333

 $00:37:57.938 \longrightarrow 00:38:02.126$ And so this further highlights the

NOTE Confidence: 0.862579132333333

 $00:38:02.126 \longrightarrow 00:38:04.917$ significant discovery potential of

NOTE Confidence: 0.862579132333333

 $00{:}38{:}04.917 \dashrightarrow 00{:}38{:}08.511$ using this approach to understand the

NOTE Confidence: 0.862579132333333

 $00:38:08.511 \longrightarrow 00:38:11.790$ pathways involved in anxiety slide.

NOTE Confidence: 0.8625791323333333

 $00{:}38{:}11.790 \dashrightarrow 00{:}38{:}13.568$ So at the beginning of this talk,

NOTE Confidence: 0.862579132333333

00:38:13.570 --> 00:38:15.676 I discussed how this approach of

NOTE Confidence: 0.862579132333333

 $00:38:15.676 \longrightarrow 00:38:17.416$ sequencing parent child trios had

NOTE Confidence: 0.8625791323333333

00:38:17.416 --> 00:38:19.348 led to the discovery of risk genes,

NOTE Confidence: 0.862579132333333

 $00:38:19.350 \longrightarrow 00:38:20.922$ first in autism,

NOTE Confidence: 0.862579132333333

 $00:38:20.922 \longrightarrow 00:38:24.066$ and now many other psychiatric conditions.

 $00:38:24.070 \longrightarrow 00:38:26.420$ Sign.

NOTE Confidence: 0.862579132333333

00:38:26.420 --> 00:38:28.802 And today I'm showed you new

NOTE Confidence: 0.862579132333333

 $00{:}38{:}28.802 \dashrightarrow 00{:}38{:}31.038$ evidence that this approach also

NOTE Confidence: 0.862579132333333

00:38:31.038 --> 00:38:33.338 has significant discovery potential

NOTE Confidence: 0.862579132333333

 $00:38:33.338 \longrightarrow 00:38:35.638$ in childhood anxiety conditions.

NOTE Confidence: 0.862579132333333

 $00:38:35.640 \longrightarrow 00:38:38.020$ And as Tom mentioned,

NOTE Confidence: 0.862579132333333

 $00:38:38.020 \longrightarrow 00:38:42.540$ we also have promising data looking at ADHD,

NOTE Confidence: 0.862579132333333

 $00{:}38{:}42.540 \dashrightarrow 00{:}38{:}44.004$ trichotillomania and excoriation

NOTE Confidence: 0.862579132333333

 $00:38:44.004 \longrightarrow 00:38:45.468$ disorder as well,

NOTE Confidence: 0.862579132333333

 $00:38:45.470 \longrightarrow 00:38:47.882$ and it's likely that many other

NOTE Confidence: 0.862579132333333

00:38:47.882 --> 00:38:49.088 psychiatric conditions could

NOTE Confidence: 0.862579132333333

 $00:38:49.088 \longrightarrow 00:38:51.322$ benefit from this approach for

NOTE Confidence: 0.862579132333333

00:38:51.322 --> 00:38:54.130 finding risk genes side.

NOTE Confidence: 0.862579132333333

00:38:54.130 --> 00:38:55.588 So in terms of next steps,

NOTE Confidence: 0.8625791323333333

00:38:55.590 --> 00:38:57.566 given our promising data,

 $00:38:57.566 \longrightarrow 00:39:00.036$ we're continuing to recruit and

NOTE Confidence: 0.862579132333333

 $00{:}39{:}00.036 \dashrightarrow 00{:}39{:}02.403$ sequence parent child trios to

NOTE Confidence: 0.862579132333333

00:39:02.403 --> 00:39:04.648 find high confidence risk genes.

NOTE Confidence: 0.862579132333333

00:39:04.650 --> 00:39:06.498 As I mentioned previously,

NOTE Confidence: 0.862579132333333

 $00:39:06.498 \longrightarrow 00:39:09.270$ usually studies have needed about a

NOTE Confidence: 0.862579132333333

00:39:09.351 --> 00:39:12.575 few 100 trios to find these first high

NOTE Confidence: 0.862579132333333

 $00:39:12.575 \longrightarrow 00:39:15.470$ competence risk genes due to rare variants.

NOTE Confidence: 0.862579132333333

00:39:15.470 --> 00:39:19.718 And then I also want to highlight that

NOTE Confidence: 0.862579132333333

00:39:19.720 --> 00:39:21.995 you know my talk today focused really

NOTE Confidence: 0.862579132333333

00:39:21.995 --> 00:39:24.410 on the process of finding risk genes,

NOTE Confidence: 0.862579132333333

 $00:39:24.410 \longrightarrow 00:39:26.097$ and I think it's important to highlight

NOTE Confidence: 0.862579132333333

 $00:39:26.097 \longrightarrow 00:39:27.610$ that that's really just a first step.

NOTE Confidence: 0.862579132333333

00:39:27.610 --> 00:39:29.200 It's an important first step,

NOTE Confidence: 0.8625791323333333

 $00:39:29.200 \longrightarrow 00:39:31.650$ but once we find these risk genes

NOTE Confidence: 0.862579132333333

 $00:39:31.650 \longrightarrow 00:39:33.810$ understanding the pathways that are involved,

NOTE Confidence: 0.862579132333333

 $00:39:33.810 \longrightarrow 00:39:35.665$ the mechanisms for which they

00:39:35.665 --> 00:39:37.520 contribute to anxiety and other

NOTE Confidence: 0.894576832222222

 $00:39:37.583 \longrightarrow 00:39:38.813$ psychiatric conditions is

NOTE Confidence: 0.894576832222222

 $00:39:38.813 \longrightarrow 00:39:40.863$ really a critical next step.

NOTE Confidence: 0.894576832222222

 $00:39:40.870 \longrightarrow 00:39:43.516$ In turn, when we think of

NOTE Confidence: 0.894576832222222

00:39:43.516 --> 00:39:44.839 developing better treatments.

NOTE Confidence: 0.894576832222222

 $00:39:44.840 \longrightarrow 00:39:47.668$ Sign. So with that,

NOTE Confidence: 0.894576832222222

00:39:47.668 --> 00:39:50.376 I first want to thank all of the family

NOTE Confidence: 0.894576832222222

 $00:39:50.376 \dashrightarrow 00:39:52.704$ members who participated in the study.

NOTE Confidence: 0.894576832222222

 $00:39:52.710 \longrightarrow 00:39:55.076$ It wouldn't have been possible without them.

NOTE Confidence: 0.894576832222222

 $00:39:55.080 \longrightarrow 00:39:57.810$ I want to thank my mentor,

NOTE Confidence: 0.894576832222222

 $00:39:57.810 \longrightarrow 00:39:59.950$ Tom Fernandez, who's been

NOTE Confidence: 0.894576832222222

 $00:39:59.950 \longrightarrow 00:40:02.090$ incredibly supportive and generous,

NOTE Confidence: 0.894576832222222

 $00{:}40{:}02.090 \dashrightarrow 00{:}40{:}03.818$ and I've just learned so much

NOTE Confidence: 0.894576832222222

 $00{:}40{:}03.818 \dashrightarrow 00{:}40{:}05.984$ working in his lab and I'm looking

NOTE Confidence: 0.894576832222222

 $00:40:05.984 \longrightarrow 00:40:07.892$ forward to continuing to work on

 $00:40:07.892 \longrightarrow 00:40:09.770$ this and other projects slide.

NOTE Confidence: 0.894576832222222

00:40:09.770 --> 00:40:12.530 I also want to take Wendy Silverman Eli

NOTE Confidence: 0.894576832222222

 $00:40:12.530 \longrightarrow 00:40:14.699$ Lebowitz for giving me the opportunity

NOTE Confidence: 0.894576832222222

00:40:14.699 --> 00:40:17.885 to work on this project and for leading

NOTE Confidence: 0.894576832222222

 $00:40:17.885 \longrightarrow 00:40:20.400$ our recruitment and clinical assessments.

NOTE Confidence: 0.894576832222222

 $00{:}40{:}20.400 \dashrightarrow 00{:}40{:}22.829$ I also want to thank Michael Block.

NOTE Confidence: 0.894576832222222

 $00:40:22.830 \longrightarrow 00:40:25.062$ He wasn't directly involved in this

NOTE Confidence: 0.894576832222222

 $00:40:25.062 \longrightarrow 00:40:27.681$ project but has mentored me on several

NOTE Confidence: 0.894576832222222

00:40:27.681 --> 00:40:30.159 projects during my time in residency slide.

NOTE Confidence: 0.894576832222222

00:40:30.160 --> 00:40:32.544 And I want to thank everyone who's part

NOTE Confidence: 0.8945768322222222

 $00{:}40{:}32.544 \dashrightarrow 00{:}40{:}34.790$ of all their groups have contributed

NOTE Confidence: 0.894576832222222

 $00:40:34.790 \longrightarrow 00:40:37.160$ to this Ain other projects that

NOTE Confidence: 0.894576832222222

 $00:40:37.233 \longrightarrow 00:40:39.308$ neurogenetics group here at Yale

NOTE Confidence: 0.894576832222222

00:40:39.310 --> 00:40:41.018 the Psychiatry residency program,

NOTE Confidence: 0.894576832222222

00:40:41.018 --> 00:40:42.299 the NRT PHE,

NOTE Confidence: 0.894576832222222

 $00{:}40{:}42.300 \dashrightarrow 00{:}40{:}44.020$ and especially this moment program.

 $00:40:44.020 \longrightarrow 00:40:46.987$ The work I presented today was funded by

NOTE Confidence: 0.894576832222222

00:40:46.987 --> 00:40:49.603 the Yale Child Study Center and the NIH.

NOTE Confidence: 0.894576832222222

00:40:49.610 --> 00:40:52.976 I also want to give a big thank you

NOTE Confidence: 0.894576832222222

 $00{:}40{:}52.976 \dashrightarrow 00{:}40{:}55.750$ for to the Seaman Lessman award in

NOTE Confidence: 0.894576832222222

 $00:40:55.750 \longrightarrow 00:40:57.780$ the selection committee as well.

NOTE Confidence: 0.894576832222222 00:40:57.780 --> 00:40:58.143 Fine.

NOTE Confidence: 0.894576832222222

00:40:58.143 --> 00:41:01.047 And I want to thank this is my

NOTE Confidence: 0.894576832222222

 $00:41:01.047 \longrightarrow 00:41:03.896$ village so all my family and friends.

NOTE Confidence: 0.894576832222222

 $00:41:03.900 \longrightarrow 00:41:05.832$ These are my Co residents both

NOTE Confidence: 0.894576832222222

 $00:41:05.832 \longrightarrow 00:41:07.120$ in the adult program.

NOTE Confidence: 0.894576832222222

00:41:07.120 --> 00:41:08.860 The sole net program, my parents,

NOTE Confidence: 0.894576832222222

00:41:08.860 --> 00:41:11.626 my sister, my husband and I

NOTE Confidence: 0.894576832222222

 $00{:}41{:}11.626 \dashrightarrow 00{:}41{:}14.559$ couldn't not mention my two kiddos.

NOTE Confidence: 0.894576832222222

 $00:41:14.560 \longrightarrow 00:41:16.114$ So next slide.

NOTE Confidence: 0.894576832222222

00:41:16.114 --> 00:41:17.668 So with that,

 $00:41:17.670 \longrightarrow 00:41:21.960$ I'm happy to take any questions.

NOTE Confidence: 0.894576832222222

 $00{:}41{:}21.960 \dashrightarrow 00{:}41{:}24.464$ I guess I'm only allowed a few questions.

NOTE Confidence: 0.894576832222222 00:41:24.470 --> 00:41:25.020 Yeah couple

NOTE Confidence: 0.88657859

00:41:25.030 --> 00:41:25.750 questions though. Again,

NOTE Confidence: 0.88657859

 $00:41:25.750 \longrightarrow 00:41:28.209$ you were right on time and I appreciate that.

NOTE Confidence: 0.88657859

 $00:41:28.210 \longrightarrow 00:41:30.186$ So yeah, we have time for a couple

NOTE Confidence: 0.88657859

 $00:41:30.186 \longrightarrow 00:41:31.790$ questions for Emily on that wonderful

NOTE Confidence: 0.88657859

 $00:41:31.790 \longrightarrow 00:41:33.647$ talk in the data she showed us

NOTE Confidence: 0.88657859

 $00{:}41{:}33.647 --> 00{:}41{:}35.009$ I did miss a couple questions

NOTE Confidence: 0.88657859

00:41:35.009 --> 00:41:36.410 in the chat after Zacks talks.

NOTE Confidence: 0.88657859

 $00:41:36.410 \longrightarrow 00:41:37.850$ I'll keep an eye on that,

NOTE Confidence: 0.88657859

 $00:41:37.850 \longrightarrow 00:41:38.840$ so please raise your hand

NOTE Confidence: 0.88657859

 $00:41:38.840 \longrightarrow 00:41:40.084$ or put something in the chat

NOTE Confidence: 0.88657859

 $00:41:40.084 \longrightarrow 00:41:41.129$ if you have any questions.

NOTE Confidence: 0.88657859

 $00:41:41.130 \longrightarrow 00:41:42.058$ Family at this time.

NOTE Confidence: 0.58428305

 $00{:}41{:}50.460 \dashrightarrow 00{:}41{:}52.659$ Emily, I have a question if I may.

 $00:41:52.660 \longrightarrow 00:41:55.412$ So in this study with the 70 ish

NOTE Confidence: 0.58428305

 $00{:}41{:}55.412 \dashrightarrow 00{:}41{:}57.690$ trios you found a bunch of hits,

NOTE Confidence: 0.58428305

00:41:57.690 --> 00:42:00.620 but you didn't find any duplicates, right?

NOTE Confidence: 0.58428305

00:42:00.620 --> 00:42:03.224 And then you compared to the existing,

NOTE Confidence: 0.58428305

 $00{:}42{:}03.230 \to 00{:}42{:}04.750$ you know the data that's already out there,

NOTE Confidence: 0.58428305

 $00:42:04.750 \longrightarrow 00:42:06.346$ and I know that in the original

NOTE Confidence: 0.58428305

 $00:42:06.346 \longrightarrow 00:42:07.831$ studies that are looking at this

NOTE Confidence: 0.58428305

 $00{:}42{:}07.831 \dashrightarrow 00{:}42{:}09.337$ kind of exome sequence that hits

NOTE Confidence: 0.58428305

 $00:42:09.337 \longrightarrow 00:42:11.027$ were continued considered real when

NOTE Confidence: 0.58428305

 $00:42:11.027 \longrightarrow 00:42:12.757$ you have duplicates because that

NOTE Confidence: 0.58428305

 $00:42:12.757 \longrightarrow 00:42:14.055$ increases your statistical confidence.

NOTE Confidence: 0.58428305

 $00:42:14.055 \longrightarrow 00:42:15.945$ But I think it's really interesting

NOTE Confidence: 0.58428305

 $00{:}42{:}15.945 \dashrightarrow 00{:}42{:}17.626$ what you did now that we're getting

NOTE Confidence: 0.58428305

 $00:42:17.626 \longrightarrow 00:42:19.010$ more hits in more disorders.

NOTE Confidence: 0.58428305

00:42:19.010 --> 00:42:20.320 That kind of you know,

 $00:42:20.320 \longrightarrow 00:42:22.348$ overlap with existing with findings from

NOTE Confidence: 0.58428305

 $00:42:22.348 \longrightarrow 00:42:24.510$ other disorders is a really interesting

NOTE Confidence: 0.58428305

 $00:42:24.510 \longrightarrow 00:42:26.754$ alternative way to find valid hits,

NOTE Confidence: 0.58428305

00:42:26.760 --> 00:42:27.621 and I wonder if you can speak

NOTE Confidence: 0.58428305

 $00:42:27.621 \longrightarrow 00:42:28.180$ a little to that.

NOTE Confidence: 0.58428305

 $00{:}42{:}28.180 \dashrightarrow 00{:}42{:}30.259$ Do you consider these proven hits or

NOTE Confidence: 0.58428305

 $00:42:30.259 \longrightarrow 00:42:31.811$ do you consider these provisional

NOTE Confidence: 0.58428305

 $00:42:31.811 \longrightarrow 00:42:33.677$ until replicated an you know they

NOTE Confidence: 0.58428305

 $00:42:33.677 \longrightarrow 00:42:35.862$ are the things that we should we

NOTE Confidence: 0.58428305

 $00:42:35.862 \longrightarrow 00:42:37.265$ should run within functional studies?

NOTE Confidence: 0.58428305

 $00:42:37.265 \longrightarrow 00:42:38.152$ Or is this still?

NOTE Confidence: 0.58428305

 $00:42:38.152 \longrightarrow 00:42:40.056$ A little work to do before we

NOTE Confidence: 0.58428305

 $00:42:40.056 \longrightarrow 00:42:41.370$ get to that point,

NOTE Confidence: 0.58428305 00:42:41.370 --> 00:42:41.680 I'd NOTE Confidence: 0.824154758

 $00:42:41.690 \longrightarrow 00:42:43.170$ say they're still provisional. I.

NOTE Confidence: 0.824154758

00:42:43.170 --> 00:42:44.605 I mean, you made a great point,

00:42:44.610 --> 00:42:47.630 so I tried to allude to this a little bit,

NOTE Confidence: 0.824154758

 $00{:}42{:}47.630 \dashrightarrow 00{:}42{:}49.667$ but in autism the first study they

NOTE Confidence: 0.824154758

 $00:42:49.667 \longrightarrow 00:42:51.802$ did 200 trios and they found one

NOTE Confidence: 0.824154758

 $00:42:51.802 \longrightarrow 00:42:54.053$ risk gene in that first study, right?

NOTE Confidence: 0.824154758

 $00:42:54.053 \longrightarrow 00:42:56.368$ They got 1 double hit.

NOTE Confidence: 0.824154758

00:42:56.370 --> 00:42:59.700 So I think you're right, it's like

NOTE Confidence: 0.905713934285714

00:42:59.710 --> 00:43:01.607 lightning striking twice in the same place.

NOTE Confidence: 0.905713934285714

 $00:43:01.610 \longrightarrow 00:43:02.605$ So then you know something

NOTE Confidence: 0.905713934285714

00:43:02.605 --> 00:43:04.060 weird is going on right, right?

NOTE Confidence: 0.892376651111111

 $00{:}43{:}04.070 \dashrightarrow 00{:}43{:}05.470$ So that's really the statistical

NOTE Confidence: 0.8923766511111111

00:43:05.470 --> 00:43:06.891 power of this approach, right?

NOTE Confidence: 0.892376651111111

00:43:06.891 --> 00:43:08.396 Is because these de Novo

NOTE Confidence: 0.8923766511111111

 $00:43:08.396 \longrightarrow 00:43:09.600$ variants are so rare.

NOTE Confidence: 0.892376651111111

 $00:43:09.600 \longrightarrow 00:43:12.288$ If you see them in unrelated individuals,

NOTE Confidence: 0.892376651111111

 $00:43:12.290 \longrightarrow 00:43:14.240$ it's likely that that's very

 $00:43:14.240 \longrightarrow 00:43:17.019$ unlikely to just be due to chance.

NOTE Confidence: 0.892376651111111

00:43:17.020 --> 00:43:18.124 And so you're right,

NOTE Confidence: 0.892376651111111

 $00:43:18.124 \longrightarrow 00:43:20.759$ that's kind of what we were harnessing here.

NOTE Confidence: 0.892376651111111

 $00:43:20.760 \longrightarrow 00:43:23.042$ I think there's a lot of evidence

NOTE Confidence: 0.892376651111111

 $00:43:23.042 \longrightarrow 00:43:24.771$ across different areas of psychiatry

NOTE Confidence: 0.892376651111111

00:43:24.771 --> 00:43:26.823 that instead of thinking of risk,

NOTE Confidence: 0.892376651111111

 $00:43:26.830 \longrightarrow 00:43:28.406$ genes for individual disorders,

NOTE Confidence: 0.892376651111111

 $00:43:28.406 \longrightarrow 00:43:30.376$ we may be thinking more

NOTE Confidence: 0.8923766511111111

 $00{:}43{:}30.376 \dashrightarrow 00{:}43{:}32.318$ about brain genes in general.

NOTE Confidence: 0.892376651111111

00:43:32.320 --> 00:43:33.224 But I think still,

NOTE Confidence: 0.8923766511111111

 $00{:}43{:}33.224 \dashrightarrow 00{:}43{:}34.870$ they're going to be jeans that are

NOTE Confidence: 0.892376651111111

00:43:34.870 --> 00:43:36.334 more common in one disorder versus

NOTE Confidence: 0.892376651111111

 $00{:}43{:}36.334 \dashrightarrow 00{:}43{:}37.900$ more common in another disorder.

NOTE Confidence: 0.8923766511111111

00:43:37.900 --> 00:43:40.875 And I think sorting that out is

NOTE Confidence: 0.892376651111111

 $00:43:40.875 \longrightarrow 00:43:42.400$ important in terms of understanding.

NOTE Confidence: 0.892376651111111

 $00:43:42.400 \longrightarrow 00:43:45.730$ Kind of the circuits involved.

 $00:43:45.730 \longrightarrow 00:43:47.860$ So this that was the approach,

NOTE Confidence: 0.892376651111111

 $00:43:47.860 \longrightarrow 00:43:49.380$ as he said that we took here because

NOTE Confidence: 0.892376651111111

 $00:43:49.380 \longrightarrow 00:43:50.896$ we thought it could give more insight.

NOTE Confidence: 0.892376651111111

 $00:43:50.900 \longrightarrow 00:43:52.580$ I I wouldn't run with these yet.

NOTE Confidence: 0.892376651111111

 $00{:}43{:}52.580 \dashrightarrow 00{:}43{:}54.752$ I I think getting these double

NOTE Confidence: 0.892376651111111

 $00:43:54.752 \longrightarrow 00:43:56.200$ hits will be helpful.

NOTE Confidence: 0.892376651111111

 $00:43:56.200 \longrightarrow 00:43:57.736$ I think the thing here that

NOTE Confidence: 0.892376651111111

00:43:57.736 --> 00:43:59.070 was encouraging is you know,

NOTE Confidence: 0.892376651111111

 $00{:}43{:}59.070 \dashrightarrow 00{:}44{:}00.648$ even though we had this hypothesis,

NOTE Confidence: 0.892376651111111

 $00:44:00.650 \longrightarrow 00:44:02.214$ anxiety is different than

NOTE Confidence: 0.8923766511111111

 $00:44:02.214 \longrightarrow 00:44:03.387$ these other conditions,

NOTE Confidence: 0.892376651111111

 $00:44:03.390 \longrightarrow 00:44:05.613$ so we weren't even sure we would see this

NOTE Confidence: 0.8923766511111111

 $00{:}44{:}05.613 \dashrightarrow 00{:}44{:}07.579$ damn increase in damaging mutations.

NOTE Confidence: 0.892376651111111

 $00:44:07.580 \longrightarrow 00:44:09.278$ But I think this is reassuring

NOTE Confidence: 0.892376651111111

 $00{:}44{:}09.278 \dashrightarrow 00{:}44{:}10.979$ that using this approach in larger

00:44:10.979 --> 00:44:12.778 cohorts may lead us to those double

NOTE Confidence: 0.892376651111111

00:44:12.778 --> 00:44:15.610 hits that you're alluding to.

NOTE Confidence: 0.892376651111111

 $00:44:15.610 \longrightarrow 00:44:18.480$ So that's the whole great stay tuned.

NOTE Confidence: 0.823786119166667

 $00:44:19.790 \longrightarrow 00:44:21.824$ Well, we have a question in

NOTE Confidence: 0.823786119166667

 $00:44:21.824 \longrightarrow 00:44:23.890$ the chat from Zarins in below.

NOTE Confidence: 0.823786119166667

00:44:23.890 --> 00:44:25.110 It says great talk Emily.

NOTE Confidence: 0.823786119166667

 $00:44:25.110 \longrightarrow 00:44:26.540$ Any evidence that the same

NOTE Confidence: 0.823786119166667

00:44:26.540 --> 00:44:28.574 gene like a phosphatase,

NOTE Confidence: 0.823786119166667

00:44:28.574 --> 00:44:31.014 for example with different mutations,

NOTE Confidence: 0.823786119166667

 $00:44:31.020 \longrightarrow 00:44:32.510$ might lead to different disorders.

NOTE Confidence: 0.823786119166667

00:44:32.510 --> 00:44:34.016 It's kind of the converse of

NOTE Confidence: 0.823786119166667

00:44:34.016 --> 00:44:35.940 the point you were just. Making

NOTE Confidence: 0.875171442857143

 $00:44:35.970 \longrightarrow 00:44:40.345$ yeah so that I think as we.

NOTE Confidence: 0.875171442857143

 $00{:}44{:}40.350 \dashrightarrow 00{:}44{:}41.870$ That's something that's really interesting.

NOTE Confidence: 0.875171442857143

00:44:41.870 --> 00:44:44.358 I didn't spend just because of the numbers,

NOTE Confidence: 0.875171442857143

 $00:44:44.360 \longrightarrow 00:44:45.944$ and I only I didn't have a double hit.

00:44:45.950 --> 00:44:47.763 I didn't spend time looking at exactly

NOTE Confidence: 0.875171442857143

 $00:44:47.763 \longrightarrow 00:44:49.479$ where like the point mutation is,

NOTE Confidence: 0.875171442857143

 $00:44:49.480 \longrightarrow 00:44:51.916$ but there are definitely examples in other

NOTE Confidence: 0.875171442857143

 $00:44:51.916 \longrightarrow 00:44:54.505$ areas of genetics where a mutation in one

NOTE Confidence: 0.875171442857143

 $00:44:54.505 \longrightarrow 00:44:56.652$ area predisposes you to one condition in

NOTE Confidence: 0.875171442857143

00:44:56.652 --> 00:44:59.068 a mutation in a different area than the

NOTE Confidence: 0.875171442857143

 $00:44:59.070 \longrightarrow 00:45:03.570$ gene predisposes you to another mutation.

NOTE Confidence: 0.875171442857143

 $00:45:03.570 \longrightarrow 00:45:06.108$ Actually, that SCN 2A mutation that

NOTE Confidence: 0.875171442857143

00:45:06.108 --> 00:45:08.470 gene that I mentioned earlier?

NOTE Confidence: 0.875171442857143

 $00:45:08.470 \longrightarrow 00:45:10.300$ That's an example of that where?

NOTE Confidence: 0.875171442857143

 $00{:}45{:}10.300 \dashrightarrow 00{:}45{:}13.540$ You get epilepsy if the mutations

NOTE Confidence: 0.875171442857143

 $00:45:13.540 \longrightarrow 00:45:16.734$ gain of function and you get autism.

NOTE Confidence: 0.875171442857143

 $00{:}45{:}16.734 \dashrightarrow 00{:}45{:}18.926$ If it's kind of a loss of function.

NOTE Confidence: 0.875171442857143

 $00:45:18.930 \longrightarrow 00:45:21.000$ So so there definitely is something

NOTE Confidence: 0.875171442857143

 $00:45:21.000 \longrightarrow 00:45:23.720$ in that I didn't quite do that here.

00:45:23.720 --> 00:45:25.589 'cause I think it's a little premature,

NOTE Confidence: 0.875171442857143

00:45:25.590 --> 00:45:27.882 but definitely something worth

NOTE Confidence: 0.875171442857143

 $00:45:27.882 \longrightarrow 00:45:29.028$ thinking about.

NOTE Confidence: 0.875171442857143

 $00:45:29.030 \longrightarrow 00:45:32.607$ As more and more genes pop up.

NOTE Confidence: 0.875171442857143

 $00:45:32.610 \longrightarrow 00:45:35.490$ Great, thank you.

NOTE Confidence: 0.83642531625

 $00:45:35.490 \longrightarrow 00:45:36.910$ Alright, another great talk

NOTE Confidence: 0.83642531625

 $00:45:36.910 \longrightarrow 00:45:39.509$ and we'll move on to our three

NOTE Confidence: 0.83642531625

00:45:39.509 --> 00:45:41.867 runners up for this year's award,

NOTE Confidence: 0.83642531625

00:45:41.870 --> 00:45:44.570 beginning with Albert Higgins Chen.

NOTE Confidence: 0.83642531625

00:45:44.570 --> 00:45:46.614 Albert Albert's primary mentor,

NOTE Confidence: 0.83642531625

00:45:46.614 --> 00:45:47.636 Morgan Levine,

NOTE Confidence: 0.83642531625

 $00:45:47.640 \longrightarrow 00:45:49.040$ was unable to be with us today,

NOTE Confidence: 0.83642531625

00:45:49.040 --> 00:45:50.546 but she's written up an introduction,

NOTE Confidence: 0.83642531625

 $00:45:50.550 \longrightarrow 00:45:53.030$ which I will give.

NOTE Confidence: 0.83642531625

00:45:53.030 --> 00:45:54.885 So Morgan says I would like to

NOTE Confidence: 0.83642531625

 $00{:}45{:}54.890 \dashrightarrow 00{:}45{:}56.692$ introduce Doctor Albert Higgins Chen

 $00:45:56.692 \longrightarrow 00:45:58.798$ and congratulate him on being selected

NOTE Confidence: 0.83642531625

00:45:58.798 --> 00:46:01.066 for honorable mention for the 2021 Last

NOTE Confidence: 0.83642531625

00:46:01.066 --> 00:46:02.806 minute award for psychiatric research.

NOTE Confidence: 0.83642531625

 $00:46:02.810 \longrightarrow 00:46:05.015$ Albert was the recipient of the 2020

NOTE Confidence: 0.83642531625

 $00:46:05.015 \longrightarrow 00:46:07.358$ Lustman Award and is being honored again.

NOTE Confidence: 0.83642531625

 $00:46:07.360 \longrightarrow 00:46:10.033$ The only goes to show how remarkable he is.

NOTE Confidence: 0.83642531625

 $00:46:10.040 \longrightarrow 00:46:11.601$ Albert is the embodiment of what it

NOTE Confidence: 0.83642531625

 $00:46:11.601 \longrightarrow 00:46:13.370$ means to be a physician scientist.

NOTE Confidence: 0.83642531625

 $00:46:13.370 \longrightarrow 00:46:15.506$ He is a brilliant independent researcher

NOTE Confidence: 0.83642531625

 $00:46:15.506 \longrightarrow 00:46:16.930$ with deep scientific knowledge,

NOTE Confidence: 0.83642531625

00:46:16.930 --> 00:46:18.334 intellectual curiosity,

NOTE Confidence: 0.83642531625

 $00:46:18.334 \longrightarrow 00:46:20.440$ creativity and compassion.

NOTE Confidence: 0.83642531625

 $00{:}46{:}20.440 \dashrightarrow 00{:}46{:}22.630$ Albert has perfectly melded his research

NOTE Confidence: 0.83642531625

 $00:46:22.630 \longrightarrow 00:46:24.939$ training in genetics and aging biology.

NOTE Confidence: 0.83642531625

00:46:24.940 --> 00:46:27.278 With his work as a psychiatry resident,

 $00:46:27.280 \longrightarrow 00:46:29.074$ as the field continues to delve

NOTE Confidence: 0.83642531625

 $00{:}46{:}29.074 \dashrightarrow 00{:}46{:}30.270$ into the molecular mechanisms

NOTE Confidence: 0.83642531625

00:46:30.321 --> 00:46:31.938 underlying psychiatric disorders,

NOTE Confidence: 0.83642531625

 $00{:}46{:}31.940 \dashrightarrow 00{:}46{:}34.145$ progress will depend on people like Albert,

NOTE Confidence: 0.83642531625

 $00:46:34.150 \longrightarrow 00:46:35.839$ who have interdisciplinary,

NOTE Confidence: 0.83642531625 00:46:35.839 --> 00:46:36.402 clinical, NOTE Confidence: 0.83642531625

 $00{:}46{:}36.402 \dashrightarrow 00{:}46{:}38.091$ molecular and computational

NOTE Confidence: 0.83642531625

 $00:46:38.091 \longrightarrow 00:46:40.833$ expertise to unravel the complex

NOTE Confidence: 0.83642531625

 $00:46:40.833 \longrightarrow 00:46:42.797$ signals of multifactorial traits.

NOTE Confidence: 0.83642531625

00:46:42.800 --> 00:46:44.739 Today he will discuss his recent paper,

NOTE Confidence: 0.83642531625

 $00:46:44.740 \longrightarrow 00:46:46.484$ aimed at dramatically bolstering

NOTE Confidence: 0.83642531625

 $00:46:46.484 \longrightarrow 00:46:48.228$ the reliability of epigenetic

NOTE Confidence: 0.83642531625

 $00:46:48.228 \longrightarrow 00:46:49.510$ biomarkers of aging.

NOTE Confidence: 0.83642531625

 $00:46:49.510 \longrightarrow 00:46:51.770$ While our lab was not the first to show

NOTE Confidence: 0.83642531625

 $00:46:51.770 \longrightarrow 00:46:53.695$ that these measures can be extremely noisy,

NOTE Confidence: 0.83642531625

 $00:46:53.700 \longrightarrow 00:46:55.370$ Albert's paper is the first

 $00:46:55.370 \longrightarrow 00:46:56.706$ to offer a solution.

NOTE Confidence: 0.83642531625

 $00:46:56.710 \longrightarrow 00:46:57.373$ In doing so,

NOTE Confidence: 0.83642531625

00:46:57.373 --> 00:46:58.699 the work he presents today will

NOTE Confidence: 0.83642531625

 $00:46:58.699 \longrightarrow 00:47:00.114$ have far reaching implications

NOTE Confidence: 0.83642531625

 $00:47:00.114 \longrightarrow 00:47:01.638$ for longitudinal an intervention,

NOTE Confidence: 0.83642531625

 $00:47:01.640 \longrightarrow 00:47:04.460$ studies of aging and disease.

NOTE Confidence: 0.83642531625

 $00:47:04.460 \longrightarrow 00:47:07.110$ So with that Albert take it away.

NOTE Confidence: 0.83642531625 00:47:07.110 --> 00:47:07.530 OK,

NOTE Confidence: 0.792656638888889

 $00:47:07.910 \longrightarrow 00:47:10.375$ thank you Chris and thank

NOTE Confidence: 0.792656638888889

00:47:10.375 --> 00:47:12.347 you Morgan by proxy.

NOTE Confidence: 0.792656638888889

 $00:47:12.350 \longrightarrow 00:47:15.890$ Exline for disclosure is the

NOTE Confidence: 0.792656638888889

 $00:47:15.890 \longrightarrow 00:47:17.092$ methodology presented in this

NOTE Confidence: 0.792656638888889

 $00{:}47{:}17.092 \dashrightarrow 00{:}47{:}19.150$ talk is the subject of a pending

NOTE Confidence: 0.792656638888889

 $00{:}47{:}19.217 \dashrightarrow 00{:}47{:}20.913$ patent application and related

NOTE Confidence: 0.792656638888889

 $00:47:20.913 \longrightarrow 00:47:22.609$ technologies have been licensed

00:47:22.609 --> 00:47:24.639 to Alicia Mhealth next slide.

NOTE Confidence: 0.79436129

 $00:47:26.680 \longrightarrow 00:47:27.868$ So a patient comes

NOTE Confidence: 0.885461302857143

 $00:47:27.880 \longrightarrow 00:47:30.855$ to your office 65 year old veteran,

NOTE Confidence: 0.885461302857143

00:47:30.860 --> 00:47:33.005 recently placed in assisted Living

NOTE Confidence: 0.885461302857143

 $00:47:33.005 \longrightarrow 00:47:35.690$ Quick chart review shows that he

NOTE Confidence: 0.885461302857143

00:47:35.690 --> 00:47:37.674 has schizophrenia, PTSD, HIV, A50,

NOTE Confidence: 0.885461302857143

00:47:37.674 --> 00:47:39.772 plus pack year, smoking history,

NOTE Confidence: 0.885461302857143

 $00:47:39.772 \longrightarrow 00:47:41.350$ and multiple comorbidities,

NOTE Confidence: 0.885461302857143

 $00:47:41.350 \longrightarrow 00:47:43.254$ and the first thing you notice about

NOTE Confidence: 0.885461302857143

 $00:47:43.254 \longrightarrow 00:47:45.460$ him when he walks into your office is

NOTE Confidence: 0.885461302857143

 $00:47:45.460 \longrightarrow 00:47:47.760$ that he looks like he is 85 years old.

NOTE Confidence: 0.885461302857143

 $00:47:47.760 \longrightarrow 00:47:49.434$ So all of these conditions along

NOTE Confidence: 0.885461302857143

 $00{:}47{:}49.434 \dashrightarrow 00{:}47{:}50.902$ with the social diversity and

NOTE Confidence: 0.885461302857143

 $00{:}47{:}50.902 \dashrightarrow 00{:}47{:}52.678$ discrimination that goes along with it,

NOTE Confidence: 0.885461302857143

 $00:47:52.680 \longrightarrow 00:47:55.560$ accelerates the biological aging process.

NOTE Confidence: 0.885461302857143

 $00:47:55.560 \longrightarrow 00:47:57.720$ Now this patient has a far higher risk

00:47:57.720 --> 00:47:59.306 of cardiovascular disease, dementia,

NOTE Confidence: 0.885461302857143

 $00{:}47{:}59.306 \dashrightarrow 00{:}48{:}01.730$ and numerous other conditions.

NOTE Confidence: 0.885461302857143

 $00:48:01.730 \longrightarrow 00:48:04.234$ So turns out that we can actually quantify

NOTE Confidence: 0.885461302857143

 $00:48:04.234 \longrightarrow 00:48:06.298$ this accelerated aging with the blood test.

NOTE Confidence: 0.885461302857143

00:48:06.300 --> 00:48:08.264 As Zach is mentioned.

NOTE Confidence: 0.885461302857143 00:48:08.264 --> 00:48:09.246 Next slide. NOTE Confidence: 0.901401721428572

 $00:48:11.430 \longrightarrow 00:48:13.860$ So this is the difference between

NOTE Confidence: 0.901401721428572

 $00{:}48{:}13.860 \to 00{:}48{:}16.129$ chronological age and biological age.

NOTE Confidence: 0.901401721428572

 $00{:}48{:}16.130 \dashrightarrow 00{:}48{:}18.750$ So chronological age is simply time

NOTE Confidence: 0.901401721428572

 $00:48:18.750 \longrightarrow 00:48:20.436$ since birth, it's not modifiable

NOTE Confidence: 0.901401721428572

00:48:20.436 --> 00:48:22.480 and we can't do anything about it.

NOTE Confidence: 0.901401721428572

00:48:22.480 --> 00:48:23.222 But importantly,

NOTE Confidence: 0.901401721428572

 $00{:}48{:}23.222 \dashrightarrow 00{:}48{:}25.077$ it has positive connotations and

NOTE Confidence: 0.901401721428572

00:48:25.077 --> 00:48:27.000 it is something to celebrate.

NOTE Confidence: 0.901401721428572

00:48:27.000 --> 00:48:28.203 Biological age, however,

00:48:28.203 --> 00:48:30.609 quantify is how much ones biology

NOTE Confidence: 0.901401721428572

00:48:30.609 --> 00:48:32.149 actually changes with time,

NOTE Confidence: 0.901401721428572

 $00:48:32.150 \longrightarrow 00:48:34.236$ and this is something that is modifiable,

NOTE Confidence: 0.901401721428572

 $00:48:34.240 \longrightarrow 00:48:37.601$ and it predicts morbidity and mortality.

NOTE Confidence: 0.901401721428572

00:48:37.601 --> 00:48:39.725 Importantly, these are separable

NOTE Confidence: 0.901401721428572

00:48:39.725 --> 00:48:42.380 and people can differ dramatically

NOTE Confidence: 0.901401721428572

 $00:48:42.451 \longrightarrow 00:48:44.629$ in the rate of biological aging,

NOTE Confidence: 0.901401721428572

00:48:44.630 --> 00:48:46.856 and we can measure this using aging

NOTE Confidence: 0.901401721428572

00:48:46.856 --> 00:48:48.656 biomarkers, and if you can measure it,

NOTE Confidence: 0.901401721428572

00:48:48.660 --> 00:48:51.996 you can manage it next line.

NOTE Confidence: 0.901401721428572

 $00{:}48{:}52.000 \dashrightarrow 00{:}48{:}54.806$ So some of the best current biomarkers

NOTE Confidence: 0.901401721428572

 $00:48:54.806 \longrightarrow 00:48:57.743$ of aging are ethnic locks as Zach heads

NOTE Confidence: 0.901401721428572

 $00:48:57.743 \longrightarrow 00:48:59.915$ discussing it in his excellent talk.

NOTE Confidence: 0.901401721428572

 $00:48:59.920 \longrightarrow 00:49:01.516$ These use the insight of that

NOTE Confidence: 0.901401721428572

00:49:01.516 --> 00:49:02.884 millions of DNA metalation sites

NOTE Confidence: 0.901401721428572

 $00{:}49{:}02.884 \dashrightarrow 00{:}49{:}04.648$ change with age and we can use

00:49:04.648 --> 00:49:05.969 machine learning techniques to select

NOTE Confidence: 0.901401721428572

 $00:49:05.969 \longrightarrow 00:49:07.761$ a few hundred that predict age or

NOTE Confidence: 0.901401721428572

 $00{:}49{:}07.770 \dashrightarrow 00{:}49{:}11.070$ mortality risk with high accuracy.

NOTE Confidence: 0.901401721428572

00:49:11.070 --> 00:49:12.858 Now I previously found that these

NOTE Confidence: 0.901401721428572

 $00:49:12.858 \longrightarrow 00:49:14.620$ clocks are the predicted mortality.

NOTE Confidence: 0.901401721428572

 $00:49:14.620 \longrightarrow 00:49:16.245$ Find that people with schizophrenia

NOTE Confidence: 0.901401721428572 00:49:16.245 --> 00:49:16.895 are older, NOTE Confidence: 0.901401721428572

00:49:16.900 --> 00:49:18.855 consistent with him dying 15

NOTE Confidence: 0.901401721428572

 $00:49:18.855 \longrightarrow 00:49:20.810$ years earlier than everyone else.

NOTE Confidence: 0.901401721428572

 $00:49:20.810 \longrightarrow 00:49:22.357$ So there is a ton of interest.

NOTE Confidence: 0.901401721428572

00:49:22.360 --> 00:49:24.604 Then eventually using these biomarkers in

NOTE Confidence: 0.901401721428572

00:49:24.604 --> 00:49:27.019 clinical practice or in clinical trial,

NOTE Confidence: 0.901401721428572 00:49:27.020 --> 00:49:27.461 however, NOTE Confidence: 0.901401721428572

 $00:49:27.461 \longrightarrow 00:49:30.989$ I found that there is a major problem

NOTE Confidence: 0.901401721428572

 $00:49:30.989 \longrightarrow 00:49:33.598$ with these epigenetic clocks.

 $00:49:33.600 \longrightarrow 00:49:34.260$ Next slide.

NOTE Confidence: 0.729820886363637

 $00{:}49{:}36.400 \dashrightarrow 00{:}49{:}38.008$ So I looked at these aging

NOTE Confidence: 0.729820886363637

 $00:49:38.008 \longrightarrow 00:49:39.560$ clocks an ask very simply.

NOTE Confidence: 0.729820886363637

 $00:49:39.560 \longrightarrow 00:49:41.495$ If you measure the same

NOTE Confidence: 0.729820886363637

 $00:49:41.495 \longrightarrow 00:49:42.656$ sample multiple times,

NOTE Confidence: 0.729820886363637

 $00:49:42.660 \longrightarrow 00:49:44.988$ do you get the same answer?

NOTE Confidence: 0.729820886363637

00:49:44.990 --> 00:49:48.278 No, so I looked at 36 blood samples,

NOTE Confidence: 0.729820886363637

 $00{:}49{:}48.280 \dashrightarrow 00{:}49{:}50.170$ each measured twice and I calculated

NOTE Confidence: 0.729820886363637

00:49:50.170 --> 00:49:51.791 the epigenetic clocks and plotted

NOTE Confidence: 0.729820886363637

00:49:51.791 --> 00:49:53.926 in the biological ages of the two

NOTE Confidence: 0.729820886363637

 $00{:}49{:}53.926 \dashrightarrow 00{:}49{:}55.150$ replicates against each other.

NOTE Confidence: 0.729820886363637

 $00:49:55.150 \longrightarrow 00:49:56.254$ Here on the left,

NOTE Confidence: 0.729820886363637

 $00:49:56.254 \longrightarrow 00:49:58.205$ and the correlation is not nearly as

NOTE Confidence: 0.729820886363637

 $00:49:58.205 \longrightarrow 00:50:00.125$ strong as one with like on the right.

NOTE Confidence: 0.729820886363637

 $00:50:00.130 \longrightarrow 00:50:02.716$ Then I plotted the difference between

NOTE Confidence: 0.729820886363637

 $00:50:02.716 \longrightarrow 00:50:05.169$ the two repeated measurements and some

 $00:50:05.169 \longrightarrow 00:50:07.793$ samples differed by as much as nine years.

NOTE Confidence: 0.729820886363637

 $00:50:07.800 \longrightarrow 00:50:08.892$ So in plain English,

NOTE Confidence: 0.729820886363637

 $00:50:08.892 \longrightarrow 00:50:11.102$ what that means is that if I could

NOTE Confidence: 0.729820886363637

00:50:11.102 --> 00:50:13.142 measure your age one day and it says

NOTE Confidence: 0.729820886363637

 $00:50:13.210 \longrightarrow 00:50:15.418$ you're 50 and next day says you're 59.

NOTE Confidence: 0.729820886363637

 $00:50:15.420 \longrightarrow 00:50:18.785$ Oh, so these are not

NOTE Confidence: 0.729820886363637

00:50:18.785 --> 00:50:21.477 particularly reliable next line.

NOTE Confidence: 0.729820886363637

 $00:50:21.480 \longrightarrow 00:50:23.814$ So I tried many methods of improving

NOTE Confidence: 0.729820886363637

 $00:50:23.814 \longrightarrow 00:50:25.659$ the reliability of these clocks

NOTE Confidence: 0.729820886363637

 $00{:}50{:}25.659 \dashrightarrow 00{:}50{:}27.939$ and eventually I found a simple

NOTE Confidence: 0.729820886363637

00:50:27.939 --> 00:50:29.502 solution using principle component

NOTE Confidence: 0.729820886363637

 $00{:}50{:}29.502 \dashrightarrow 00{:}50{:}31.312$ analysis which I won't describe

NOTE Confidence: 0.729820886363637

 $00{:}50{:}31.312 \dashrightarrow 00{:}50{:}33.672$ in detail but just know that it

NOTE Confidence: 0.729820886363637

 $00{:}50{:}33.672 \dashrightarrow 00{:}50{:}35.496$ is a method that can separate

NOTE Confidence: 0.729820886363637

00:50:35.496 --> 00:50:37.385 signal from noise and instead

 $00:50:37.385 \longrightarrow 00:50:39.280$ of using directly the metalation

NOTE Confidence: 0.729820886363637

 $00{:}50{:}39.280 \dashrightarrow 00{:}50{:}41.339$ sites to predict biological age,

NOTE Confidence: 0.729820886363637

 $00:50:41.340 \longrightarrow 00:50:43.725$ I transformed the DNA methylation

NOTE Confidence: 0.729820886363637

00:50:43.725 --> 00:50:45.633 using principle component Alesis

NOTE Confidence: 0.729820886363637

 $00:50:45.633 \longrightarrow 00:50:48.554$ and then used the new variables to

NOTE Confidence: 0.729820886363637

 $00{:}50{:}48.554 \dashrightarrow 00{:}50{:}50.550$ predict biological age next line.

NOTE Confidence: 0.8348008825

 $00{:}50{:}52.610 \dashrightarrow 00{:}50{:}55.274$ And this new clocks are way more reliable.

NOTE Confidence: 0.8348008825

00:50:55.280 --> 00:50:57.422 Here we can look again at 36 samples each,

NOTE Confidence: 0.8348008825

 $00:50:57.430 \longrightarrow 00:51:01.070$ measure twice and blue is our new clocks.

NOTE Confidence: 0.8348008825

 $00:51:01.070 \longrightarrow 00:51:02.390$ And now the replicates.

NOTE Confidence: 0.8348008825

 $00{:}51{:}02.390 \dashrightarrow 00{:}51{:}05.074$ Now agree far more closely and most agree

NOTE Confidence: 0.8348008825

 $00{:}51{:}05.074 \dashrightarrow 00{:}51{:}07.386$ within one year and we applied this

NOTE Confidence: 0.8348008825

 $00:51:07.386 \longrightarrow 00:51:09.851$ method to six commonly used clocks and

NOTE Confidence: 0.8348008825

 $00:51:09.851 \longrightarrow 00:51:13.720$ they all greatly improved next slide.

NOTE Confidence: 0.8348008825

 $00:51:13.720 \longrightarrow 00:51:16.324$ And so does this mean the clocks

NOTE Confidence: 0.8348008825

 $00:51:16.324 \longrightarrow 00:51:17.440$ more clinically relevant?

 $00:51:17.440 \longrightarrow 00:51:19.526$ Yes, so I showed that these have

NOTE Confidence: 0.8348008825

 $00:51:19.526 \longrightarrow 00:51:20.793$ much stronger relationships with

NOTE Confidence: 0.8348008825

00:51:20.793 --> 00:51:22.368 mortality and many other factors,

NOTE Confidence: 0.8348008825

 $00:51:22.370 \longrightarrow 00:51:24.876$ and because they are now less noisy

NOTE Confidence: 0.8348008825

 $00:51:24.876 \longrightarrow 00:51:26.750$ and furthermore we can actually

NOTE Confidence: 0.8348008825

 $00:51:26.750 \longrightarrow 00:51:29.056$ use these clocks to track someones

NOTE Confidence: 0.8348008825

 $00:51:29.056 \longrightarrow 00:51:30.238$ aging process overtime.

NOTE Confidence: 0.8348008825

 $00:51:30.240 \longrightarrow 00:51:32.752$ So I looked at 300 people followed for

NOTE Confidence: 0.8348008825

 $00{:}51{:}32.752 \dashrightarrow 00{:}51{:}35.900$ 20 years and we see that the original

NOTE Confidence: 0.8348008825

 $00:51:35.900 \longrightarrow 00:51:38.269$ clocks actually fluctuate wildly over time.

NOTE Confidence: 0.8348008825

 $00:51:38.270 \longrightarrow 00:51:40.358$ Turns out that this is mostly just noise,

NOTE Confidence: 0.8348008825

 $00:51:40.360 \longrightarrow 00:51:43.496$ so and our new clocks actually show

NOTE Confidence: 0.8348008825

 $00{:}51{:}43.496 {\:{\mbox{--}}}{\:{\mbox{-}}} 00{:}51{:}46.435$ a nice steady aging trend there

NOTE Confidence: 0.8348008825

00:51:46.435 --> 00:51:49.423 on the bottom right next slide.

NOTE Confidence: 0.8348008825

 $00:51:49.430 \longrightarrow 00:51:51.096$ And could we even use this to

 $00:51:51.096 \longrightarrow 00:51:52.824$ discover new treatments that might

NOTE Confidence: 0.8348008825

 $00:51:52.824 \longrightarrow 00:51:54.930$ be able to help our patient?

NOTE Confidence: 0.8348008825 00:51:54.930 --> 00:51:55.243 Yes, NOTE Confidence: 0.8348008825

 $00:51:55.243 \longrightarrow 00:51:57.434$ so I simulated a clinical trial that

NOTE Confidence: 0.8348008825

 $00:51:57.434 \longrightarrow 00:52:00.298$ aims to modify someone's trajectory of aging.

NOTE Confidence: 0.8348008825

00:52:00.298 --> 00:52:01.639 Measuring these epigenetic

NOTE Confidence: 0.8348008825

 $00:52:01.639 \longrightarrow 00:52:02.533$ clocks longitudinally.

NOTE Confidence: 0.8348008825

 $00:52:02.540 \longrightarrow 00:52:04.145$ Now the issue of reliability

NOTE Confidence: 0.8348008825

 $00:52:04.145 \longrightarrow 00:52:05.108$ is critical here,

NOTE Confidence: 0.8348008825

 $00:52:05.110 \longrightarrow 00:52:07.475$ because noise affects both baseline

NOTE Confidence: 0.8348008825

 $00{:}52{:}07.475 \dashrightarrow 00{:}52{:}09.367$ and follow up measurements.

NOTE Confidence: 0.8348008825

 $00:52:09.370 \longrightarrow 00:52:11.848$ It's cures our ability to detect the

NOTE Confidence: 0.8348008825

 $00:52:11.848 \longrightarrow 00:52:14.452$ effect of an intervention and power

NOTE Confidence: 0.8348008825

 $00:52:14.452 \longrightarrow 00:52:15.932$ analysis indicate that these new

NOTE Confidence: 0.8348008825

00:52:15.932 --> 00:52:17.890 PC clocks are far more sensitive,

NOTE Confidence: 0.8348008825

 $00{:}52{:}17.890 \dashrightarrow 00{:}52{:}20.122$ reducing the sample size needed to

 $00:52:20.122 \longrightarrow 00:52:22.650$ detect an effect by up to tenfold.

NOTE Confidence: 0.8348008825

 $00:52:22.650 \longrightarrow 00:52:24.258$ Now given how challenging clinical trials

NOTE Confidence: 0.8348008825

 $00:52:24.258 \longrightarrow 00:52:25.776$ are, this could save a lot of money.

NOTE Confidence: 0.8348008825

 $00:52:25.780 \longrightarrow 00:52:28.988$ And resources next slide.

NOTE Confidence: 0.8348008825

 $00:52:28.990 \longrightarrow 00:52:30.610$ So what can we actually do for our patient?

NOTE Confidence: 0.8348008825 00:52:30.610 --> 00:52:30.877 Well,

NOTE Confidence: 0.8348008825

 $00:52:30.877 \longrightarrow 00:52:32.746$ we can measure his biological age and

NOTE Confidence: 0.8348008825

00:52:32.746 --> 00:52:34.640 find that all those years of living,

NOTE Confidence: 0.8348008825

 $00:52:34.640 \longrightarrow 00:52:36.614$ mental illness and all the discrimination

NOTE Confidence: 0.8348008825

 $00:52:36.614 \longrightarrow 00:52:39.141$ is a social hardships that put him at a

NOTE Confidence: 0.8348008825

00:52:39.141 --> 00:52:40.870 high risk of morbidity and mortality.

NOTE Confidence: 0.8348008825

 $00:52:40.870 \longrightarrow 00:52:42.926$ And we can look to the many aging

NOTE Confidence: 0.8348008825

00:52:42.926 --> 00:52:44.029 treatments currently being investigated

NOTE Confidence: 0.8348008825

 $00:52:44.029 \longrightarrow 00:52:46.197$ and we may eventually be able to treat

NOTE Confidence: 0.8348008825

 $00:52:46.247 \longrightarrow 00:52:47.767$ this problem at least partially.

00:52:47.770 --> 00:52:48.095 Importantly,

NOTE Confidence: 0.8348008825

 $00{:}52{:}48.095 \dashrightarrow 00{:}52{:}50.370$ this would help prevent all the diseases

NOTE Confidence: 0.8348008825

 $00:52:50.370 \longrightarrow 00:52:52.709$ of aging or once cardiovascular disease,

NOTE Confidence: 0.8348008825

 $00:52:52.710 \longrightarrow 00:52:53.492$ cancer, dementia,

NOTE Confidence: 0.834800882500:52:53.492 --> 00:52:53.883 etc. NOTE Confidence: 0.8348008825

 $00:52:53.883 \longrightarrow 00:52:56.620$ And this will be made possible by

NOTE Confidence: 0.8348008825

 $00:52:56.694 \longrightarrow 00:52:58.999$ aging biomarkers that are highly.

NOTE Confidence: 0.8348008825

00:52:59.000 --> 00:53:01.712 Reliable so I will wrap up a wrap

NOTE Confidence: 0.8348008825

 $00:53:01.712 \longrightarrow 00:53:03.370$ up there next slide.

NOTE Confidence: 0.8348008825

 $00{:}53{:}03.370 \dashrightarrow 00{:}53{:}05.232$ And I like to think that less on

NOTE Confidence: 0.8348008825

 $00{:}53{:}05.232 \dashrightarrow 00{:}53{:}07.205$ family that live in lab is she

NOTE Confidence: 0.8348008825

00:53:07.205 --> 00:53:08.595 doctor been all my collaborators

NOTE Confidence: 0.8348008825

 $00:53:08.595 \longrightarrow 00:53:10.538$ and every body else in time trade.

NOTE Confidence: 0.9079392

00:53:14.620 --> 00:53:16.675 Great thank you. Albert was very

NOTE Confidence: 0.9079392

 $00:53:16.675 \longrightarrow 00:53:17.602$ clearly presented presentation

NOTE Confidence: 0.9079392

00:53:17.602 --> 00:53:20.420 of very important work.

 $00:53:20.420 \longrightarrow 00:53:22.028$ We're not going to have time

NOTE Confidence: 0.872676488181818

 $00:53:22.028 \longrightarrow 00:53:23.480$ for questions at this point.

NOTE Confidence: 0.872676488181818

 $00{:}53{:}23.480 \to 00{:}53{:}27.040$ We're going to move on to our our

NOTE Confidence: 0.872676488181818

 $00:53:27.040 \longrightarrow 00:53:29.336$ second honorable mentioning here.

NOTE Confidence: 0.872676488181818

 $00:53:29.336 \longrightarrow 00:53:31.466$ The 2021 Last Minute awards.

NOTE Confidence: 0.872676488181818

 $00:53:31.470 \longrightarrow 00:53:33.423$ Peter Na and I want to invite

NOTE Confidence: 0.872676488181818

 $00:53:33.423 \longrightarrow 00:53:34.986$ Rob Pietrzak to come and

NOTE Confidence: 0.872676488181818

 $00:53:34.986 \longrightarrow 00:53:36.626$ introduce Peter in his work.

NOTE Confidence: 0.831199593333333

 $00:53:38.120 \longrightarrow 00:53:39.260$ Thank you Chris.

NOTE Confidence: 0.831199593333333

00:53:39.260 --> 00:53:40.400 Good morning everyone.

NOTE Confidence: 0.831199593333333

 $00{:}53{:}40.400 \dashrightarrow 00{:}53{:}42.152$ Thank you to the Lawson family

NOTE Confidence: 0.831199593333333

 $00{:}53{:}42.152 \dashrightarrow 00{:}53{:}43.616$ and congratulations to all of

NOTE Confidence: 0.831199593333333

00:53:43.616 --> 00:53:44.880 the last minute award ease.

NOTE Confidence: 0.831199593333333

00:53:44.880 --> 00:53:46.704 Well it's my pleasure this morning

NOTE Confidence: 0.831199593333333

00:53:46.704 --> 00:53:48.284 to introduce Doctor Peter Na,

00:53:48.290 --> 00:53:50.180 recipient of an honorable mention

NOTE Confidence: 0.831199593333333

00:53:50.180 --> 00:53:52.240 for the 2021 lesson in the world.

NOTE Confidence: 0.831199593333333

00:53:52.240 --> 00:53:54.132 I first met Peter just five months ago

NOTE Confidence: 0.831199593333333

00:53:54.132 --> 00:53:56.324 when he reached out to me to inquire

NOTE Confidence: 0.831199593333333

 $00:53:56.324 \longrightarrow 00:53:57.748$ about potential research opportunities

NOTE Confidence: 0.831199593333333

00:53:57.748 --> 00:53:59.971 using data from the National Health

NOTE Confidence: 0.831199593333333

 $00:53:59.971 \longrightarrow 00:54:01.816$ and Resilience and Veterans Study.

NOTE Confidence: 0.831199593333333

 $00:54:01.820 \longrightarrow 00:54:03.760$ This is a nationally representative

NOTE Confidence: 0.831199593333333

00:54:03.760 --> 00:54:05.312 prospective cohort study of

NOTE Confidence: 0.831199593333333

00:54:05.312 --> 00:54:06.888 veterans that Steve Southwick,

NOTE Confidence: 0.831199593333333

 $00{:}54{:}06.890 \dashrightarrow 00{:}54{:}07.336$ John Crystal,

NOTE Confidence: 0.831199593333333

00:54:07.336 --> 00:54:08.451 and I have been conducting

NOTE Confidence: 0.831199593333333

 $00:54:08.451 \longrightarrow 00:54:10.106$ for the past 10 years.

NOTE Confidence: 0.831199593333333

 $00:54:10.106 \longrightarrow 00:54:12.072$ I was immediately impressed by

NOTE Confidence: 0.831199593333333

00:54:12.072 --> 00:54:13.385 Peter's academic background,

NOTE Confidence: 0.831199593333333

 $00:54:13.385 \longrightarrow 00:54:15.124$ which includes an undergraduate

00:54:15.124 --> 00:54:17.064 degree from Seoul National University

NOTE Confidence: 0.831199593333333

00:54:17.064 --> 00:54:19.138 where he graduated Summa Cloudy.

NOTE Confidence: 0.831199593333333

 $00:54:19.140 \longrightarrow 00:54:21.168$ An MD degree also from Seoul

NOTE Confidence: 0.831199593333333

 $00:54:21.168 \longrightarrow 00:54:23.071$ National and MPH from Harvard

NOTE Confidence: 0.831199593333333

 $00:54:23.071 \longrightarrow 00:54:25.441$ internship training at the Male

NOTE Confidence: 0.831199593333333

00:54:25.441 --> 00:54:27.680 Clinic Psychiatry Residency at NYU,

NOTE Confidence: 0.831199593333333

 $00:54:27.680 \longrightarrow 00:54:29.648$ and most recently in addiction psychiatry.

NOTE Confidence: 0.831199593333333

 $00:54:29.650 \longrightarrow 00:54:31.330$ Quite so much chip at Yale.

NOTE Confidence: 0.831199593333333

 $00:54:31.330 \longrightarrow 00:54:32.860$ In recognition of his work,

NOTE Confidence: 0.831199593333333

 $00:54:32.860 \longrightarrow 00:54:35.146$ Peter has already received three awards,

NOTE Confidence: 0.831199593333333

00:54:35.150 --> 00:54:37.628 including a Samsung Fellowship from Appa,

NOTE Confidence: 0.831199593333333

00:54:37.630 --> 00:54:39.810 the NMH Outstanding Resident Award,

NOTE Confidence: 0.831199593333333

00:54:39.810 --> 00:54:40.412 Honorable mention.

NOTE Confidence: 0.831199593333333

00:54:40.412 --> 00:54:42.218 And the John Runner Award from

NOTE Confidence: 0.831199593333333

00:54:42.218 --> 00:54:43.898 the American Academy of Addiction,

 $00:54:43.900 \longrightarrow 00:54:44.490$ Psychiatry.

NOTE Confidence: 0.831199593333333

 $00:54:44.490 \longrightarrow 00:54:48.030$ In addition to his academic accomplishments,

NOTE Confidence: 0.831199593333333

 $00:54:48.030 \longrightarrow 00:54:50.106$ Peter has served as a senior

NOTE Confidence: 0.831199593333333

 $00:54:50.106 \longrightarrow 00:54:51.490$ noncommissioned officer of the

NOTE Confidence: 0.831199593333333

 $00:54:51.550 \longrightarrow 00:54:53.804$ G3 Liaison Office as part of the

NOTE Confidence: 0.831199593333333

00:54:53.804 --> 00:54:55.538 Korean augmentation of the US Army,

NOTE Confidence: 0.831199593333333

 $00:54:55.540 \longrightarrow 00:54:58.636$ otherwise known as Catoosa in Camp Red Cloud.

NOTE Confidence: 0.831199593333333

00:54:58.640 --> 00:55:00.220 In recognition of his service,

NOTE Confidence: 0.831199593333333

00:55:00.220 --> 00:55:02.135 he received 2US Army achievement

NOTE Confidence: 0.831199593333333

 $00:55:02.135 \longrightarrow 00:55:04.050$ medals and a certificate of

NOTE Confidence: 0.831199593333333

 $00:55:04.114 \longrightarrow 00:55:06.539$ Achievement for excellence in service.

NOTE Confidence: 0.831199593333333

 $00:55:06.540 \longrightarrow 00:55:07.735$ Peter came to Yale,

NOTE Confidence: 0.831199593333333

 $00:55:07.735 \longrightarrow 00:55:09.472$ already quite accomplished with three.

NOTE Confidence: 0.831199593333333

00:55:09.472 --> 00:55:11.650 First off in manuscripts and eight

NOTE Confidence: 0.831199593333333

 $00:55:11.714 \longrightarrow 00:55:13.504$ first offered meeting abstract since

NOTE Confidence: 0.831199593333333

 $00:55:13.504 \longrightarrow 00:55:16.078$ the start of his fellowship in July 2020.

 $00:55:16.080 \longrightarrow 00:55:17.274$ Peter has accelerated

NOTE Confidence: 0.831199593333333

 $00:55:17.274 \longrightarrow 00:55:18.468$ his productivity further.

NOTE Confidence: 0.831199593333333

 $00:55:18.470 \longrightarrow 00:55:19.814$ First authoring 9 manuscripts

NOTE Confidence: 0.831199593333333

 $00:55:19.814 \longrightarrow 00:55:21.494$ that are currently in press,

NOTE Confidence: 0.831199593333333

 $00:55:21.500 \longrightarrow 00:55:23.080$ except that are under review,

NOTE Confidence: 0.831199593333333

 $00:55:23.080 \longrightarrow 00:55:25.066$ including five in which I've been

NOTE Confidence: 0.831199593333333

 $00:55:25.066 \longrightarrow 00:55:26.820$ fortunate to services primary mentor.

NOTE Confidence: 0.831199593333333

00:55:26.820 --> 00:55:28.368 So run on average right now

NOTE Confidence: 0.831199593333333

 $00{:}55{:}28.368 {\:\raisebox{--}{\text{--}}}{\:\raisebox{--}{\text{--}}}{\:\raisebox{--}{\text{--}}} 00{:}55{:}30.121$ with paper per month will see if

NOTE Confidence: 0.831199593333333

 $00:55:30.121 \longrightarrow 00:55:31.447$ we can keep up with that.

NOTE Confidence: 0.831199593333333

 $00:55:31.450 \longrightarrow 00:55:33.154$ Peters exemplary productivity speaks

NOTE Confidence: 0.831199593333333

 $00:55:33.154 \longrightarrow 00:55:35.146$ to his deep rooted interest and

NOTE Confidence: 0.831199593333333

 $00{:}55{:}35.146 \dashrightarrow 00{:}55{:}37.096$ commitment to a career as a clinician,

NOTE Confidence: 0.831199593333333 00:55:37.100 --> 00:55:37.408 scholar, NOTE Confidence: 0.831199593333333

 $00:55:37.408 \longrightarrow 00:55:39.564$ and psychiatry as well as to his

00:55:39.564 --> 00:55:41.100 dedication to make meaningful

NOTE Confidence: 0.831199593333333

00:55:41.100 --> 00:55:43.260 scientific contributions to our field,

NOTE Confidence: 0.831199593333333

 $00:55:43.260 \longrightarrow 00:55:45.276$ particularly in veteran mental

NOTE Confidence: 0.831199593333333

 $00:55:45.276 \longrightarrow 00:55:46.788$ health this July.

NOTE Confidence: 0.831199593333333

 $00:55:46.790 \longrightarrow 00:55:48.500$ Peter will join the VA Connecticut

NOTE Confidence: 0.831199593333333

 $00:55:48.500 \longrightarrow 00:55:50.003$ staff and is currently already

NOTE Confidence: 0.831199593333333

 $00:55:50.003 \longrightarrow 00:55:51.553$ preparing an application for a

NOTE Confidence: 0.831199593333333

00:55:51.553 --> 00:55:53.367 be a career development award to

NOTE Confidence: 0.831199593333333

 $00{:}55{:}53.367 \dashrightarrow 00{:}55{:}54.667$ expand his research to consider

NOTE Confidence: 0.831199593333333

00:55:54.667 --> 00:55:56.731 the role of genetic factors in

NOTE Confidence: 0.831199593333333

 $00{:}55{:}56.731 \dashrightarrow 00{:}55{:}58.866$ suicide and substance use disorders.

NOTE Confidence: 0.831199593333333

 $00:55:58.870 \longrightarrow 00:56:00.560$ He's already developed a detailed

NOTE Confidence: 0.831199593333333

 $00:56:00.560 \longrightarrow 00:56:02.250$ research and primary mentor ship

NOTE Confidence: 0.831199593333333

 $00:56:02.303 \longrightarrow 00:56:03.989$ plan with Doctor Joel Gelernter and

NOTE Confidence: 0.831199593333333

 $00:56:03.989 \longrightarrow 00:56:06.124$ me that will enable him to develop

NOTE Confidence: 0.831199593333333

 $00:56:06.124 \longrightarrow 00:56:08.026$ new skills and expertise in genetic

 $00:56:08.026 \longrightarrow 00:56:09.940$ psychiatric epidemiology with the

NOTE Confidence: 0.831199593333333

00:56:09.940 --> 00:56:12.060 ultimate goal of identifying

NOTE Confidence: 0.831199593333333

00:56:12.060 --> 00:56:13.120 modifiable psychosocial,

NOTE Confidence: 0.831199593333333

00:56:13.120 --> 00:56:15.125 moderators or polygenic risk for

NOTE Confidence: 0.831199593333333

 $00:56:15.125 \longrightarrow 00:56:17.130$ suicide and substance use disorders.

NOTE Confidence: 0.831199593333333

00:56:17.130 --> 00:56:18.504 I must note that throughout my

NOTE Confidence: 0.831199593333333

00:56:18.504 --> 00:56:19.710 experience of working with Peter,

NOTE Confidence: 0.831199593333333

 $00:56:19.710 \longrightarrow 00:56:21.368$ I've been impressed by his scientific

NOTE Confidence: 0.831199593333333

00:56:21.370 --> 00:56:23.400 writing and critical thinking skills,

NOTE Confidence: 0.831199593333333

 $00:56:23.400 \longrightarrow 00:56:25.698$ as well as his remarkable ability

NOTE Confidence: 0.831199593333333

 $00{:}56{:}25.698 {\:{\mbox{--}}\!>}\ 00{:}56{:}27.230$ to translate very complex

NOTE Confidence: 0.831199593333333

 $00:56:27.298 \longrightarrow 00:56:29.344$ epidemiological findings into

NOTE Confidence: 0.831199593333333

 $00{:}56{:}29.344 \dashrightarrow 00{:}56{:}31.390$ actionable clinical implications.

NOTE Confidence: 0.831199593333333

 $00:56:31.390 \longrightarrow 00:56:33.304$ I'd also like to highlight that

NOTE Confidence: 0.831199593333333

 $00:56:33.304 \longrightarrow 00:56:34.580$ Peters research and veteran

 $00:56:34.643 \longrightarrow 00:56:36.806$ mental health and suicide is inspired by

NOTE Confidence: 0.809649877333333

 $00:56:36.806 \longrightarrow 00:56:38.796$ his own military experience and losses

NOTE Confidence: 0.809649877333333

00:56:38.796 --> 00:56:41.148 that he personally endured on that note,

NOTE Confidence: 0.809649877333333

00:56:41.150 --> 00:56:43.187 as we head into Memorial Day weekend,

NOTE Confidence: 0.809649877333333

00:56:43.190 --> 00:56:44.855 I'd like to share a quote from the author,

NOTE Confidence: 0.809649877333333

00:56:44.860 --> 00:56:48.059 Richelle Goodrich, who said on Memorial Day.

NOTE Confidence: 0.809649877333333

 $00:56:48.060 \longrightarrow 00:56:50.844$ Take time to remember those who have fallen,

NOTE Confidence: 0.809649877333333

 $00.56.50.850 \longrightarrow 00.56.52.370$ but on every day after,

NOTE Confidence: 0.809649877333333

 $00:56:52.370 \longrightarrow 00:56:54.953$ do more put the freedoms that they

NOTE Confidence: 0.809649877333333

 $00:56:54.953 \longrightarrow 00:56:58.078$ died for to greater and nobler uses.

NOTE Confidence: 0.809649877333333

 $00{:}56{:}58.080 \dashrightarrow 00{:}57{:}00.103$ Today Peter will present results of a

NOTE Confidence: 0.809649877333333

 $00:57:00.103 \longrightarrow 00:57:02.135$ recent study on factors associated with

NOTE Confidence: 0.809649877333333

 $00:57:02.135 \longrightarrow 00:57:04.355$ suicidal thinking during the pandemic and

NOTE Confidence: 0.809649877333333

 $00:57:04.355 \longrightarrow 00:57:06.860$ US veterans with pre-existing conditions.

NOTE Confidence: 0.809649877333333

00:57:06.860 --> 00:57:09.056 To speak to Peter's amazing efficiency,

NOTE Confidence: 0.809649877333333

 $00:57:09.060 \longrightarrow 00:57:11.335$ he wrote this paper in one week.

 $00:57:11.340 \longrightarrow 00:57:14.290$ Peter, please. Thank

NOTE Confidence: 0.777818684285714

 $00{:}57{:}14.300 \dashrightarrow 00{:}57{:}16.310$ you Doctor Pietrzak for your

NOTE Confidence: 0.777818684285714

 $00:57:16.310 \longrightarrow 00:57:17.473$ kind introduction. Today.

NOTE Confidence: 0.777818684285714

00:57:17.473 --> 00:57:19.944 I'll be presenting our research on factors

NOTE Confidence: 0.777818684285714

 $00:57:19.944 \longrightarrow 00:57:21.976$ associated with suicidal ideations during

NOTE Confidence: 0.777818684285714

00:57:21.976 --> 00:57:24.514 the COVID-19 pandemic and veterans with

NOTE Confidence: 0.777818684285714

 $00:57:24.514 \longrightarrow 00:57:26.250$ pre-existing psychiatric conditions.

NOTE Confidence: 0.777818684285714

 $00{:}57{:}26.250 \dashrightarrow 00{:}57{:}29.010$ I do not have any disclosures to report.

NOTE Confidence: 0.777818684285714

 $00:57:29.010 \dashrightarrow 00:57:31.770$ This paper was published in the Journal of

NOTE Confidence: 0.777818684285714

 $00:57:31.770 \longrightarrow 00:57:33.509$ Psychiatric Research earlier this year.

NOTE Confidence: 0.777818684285714 00:57:33.510 --> 00:57:35.750 Next slide. NOTE Confidence: 0.777818684285714

 $00:57:35.750 \longrightarrow 00:57:37.520$ As we're all aware mental health

NOTE Confidence: 0.777818684285714

 $00{:}57{:}37.520 {\:{\circ}{\circ}{\circ}}>00{:}57{:}39.300$ burden during the pandemic is on

NOTE Confidence: 0.777818684285714

 $00:57:39.300 \longrightarrow 00:57:40.920$ the rise with reports of increased

NOTE Confidence: 0.777818684285714

 $00:57:40.920 \longrightarrow 00:57:42.070$ prevalence of depression,

 $00:57:42.070 \longrightarrow 00:57:43.722$ anxiety and alcohol consumption

NOTE Confidence: 0.777818684285714

 $00:57:43.722 \longrightarrow 00:57:45.374$ of the general public.

NOTE Confidence: 0.777818684285714

 $00:57:45.380 \longrightarrow 00:57:47.240$ There were also concerns about

NOTE Confidence: 0.777818684285714

 $00:57:47.240 \longrightarrow 00:57:48.728$ possible increase in suicidal

NOTE Confidence: 0.777818684285714

 $00:57:48.728 \longrightarrow 00:57:51.199$ behavior based on the fact that during

NOTE Confidence: 0.777818684285714

00:57:51.199 --> 00:57:52.559 previous pandemics and outbreaks,

NOTE Confidence: 0.777818684285714

 $00:57:52.560 \longrightarrow 00:57:54.632$ suicide rate increased historically.

NOTE Confidence: 0.777818684285714

 $00:57:54.632 \longrightarrow 00:57:55.668$ For example,

NOTE Confidence: 0.777818684285714

00:57:55.670 --> 00:57:58.085 during the SARS outbreak in Hong Kong,

NOTE Confidence: 0.777818684285714

 $00:57:58.090 \longrightarrow 00:58:00.005$ the most significant increase were

NOTE Confidence: 0.777818684285714

 $00{:}58{:}00.005 \dashrightarrow 00{:}58{:}04.130$ found among older adults, next light.

NOTE Confidence: 0.777818684285714

00:58:04.130 --> 00:58:05.225 Possible vulnerable groups

NOTE Confidence: 0.777818684285714

 $00:58:05.225 \longrightarrow 00:58:06.320$ during the pandemic.

NOTE Confidence: 0.777818684285714

 $00:58:06.320 \longrightarrow 00:58:08.536$ Identify where older adults,

NOTE Confidence: 0.777818684285714

 $00:58:08.536 \longrightarrow 00:58:11.860$ possibly due to more physical comorbidities.

NOTE Confidence: 0.777818684285714

 $00:58:11.860 \longrightarrow 00:58:13.612$ Also, who experienced greater

 $00{:}58{:}13.612 \dashrightarrow 00{:}58{:}15.364$ social isolation and loneliness.

NOTE Confidence: 0.777818684285714

00:58:15.370 --> 00:58:15.766 Also,

NOTE Confidence: 0.777818684285714

 $00{:}58{:}15.766 \dashrightarrow 00{:}58{:}17.746$ individuals with mental disorders retreat

NOTE Confidence: 0.777818684285714

 $00:58:17.746 \longrightarrow 00:58:20.858$ who may be uniquely vulnerable to increase

NOTE Confidence: 0.777818684285714

 $00:58:20.858 \longrightarrow 00:58:23.298$ psychological distress during the pandemic.

NOTE Confidence: 0.777818684285714

 $00.58:23.300 \longrightarrow 00.58:23.640$ Also,

NOTE Confidence: 0.777818684285714

 $00:58:23.640 \longrightarrow 00:58:25.340$ military veterans were well known.

NOTE Confidence: 0.777818684285714

 $00{:}58{:}25.340 \to 00{:}58{:}27.447$ High risk group for suicide as well

NOTE Confidence: 0.777818684285714

 $00:58:27.447 \longrightarrow 00:58:29.163$ as COVID-19 survivors who showed

NOTE Confidence: 0.777818684285714

 $00:58:29.163 \longrightarrow 00:58:30.675$ higher prevalence of depression,

NOTE Confidence: 0.777818684285714

 $00{:}58{:}30.680 \dashrightarrow 00{:}58{:}34.586$ anxiety and PTSD compared to non survivors.

NOTE Confidence: 0.777818684285714

 $00:58:34.590 \longrightarrow 00:58:37.194$ Next slide so to meet this

NOTE Confidence: 0.777818684285714

 $00{:}58{:}37.194 \dashrightarrow 00{:}58{:}38.930$ urgent public health concern,

NOTE Confidence: 0.777818684285714

 $00:58:38.930 \longrightarrow 00:58:41.730$ we analyze the 2019 to 20 National

NOTE Confidence: 0.777818684285714

 $00:58:41.730 \longrightarrow 00:58:44.014$ Health and resilience in various study

 $00:58:44.014 \dashrightarrow 00:58:47.010$ and HRV S among the total 4000 samples.

NOTE Confidence: 0.777818684285714

00:58:47.010 --> 00:58:49.740 We analyze subsample of 6061 veterans

NOTE Confidence: 0.777818684285714

 $00:58:49.740 \longrightarrow 00:58:52.614$ who screen positive for pre pandemic

NOTE Confidence: 0.777818684285714

00:58:52.614 --> 00:58:55.524 major psychiatric disorders such as MD,

NOTE Confidence: 0.777818684285714

 $00:58:55.530 \longrightarrow 00:58:59.370$ JD, PTSD, and or SUT.

NOTE Confidence: 0.777818684285714

00:58:59.370 --> 00:59:02.106 Baseline survey or wave one survey as well.

NOTE Confidence: 0.777818684285714

 $00:59:02.110 \dashrightarrow 00:59:04.462$ Call Pre Pandemic survey was completed

NOTE Confidence: 0.777818684285714

 $00:59:04.462 \longrightarrow 00:59:06.827$ prior to the first known identified

NOTE Confidence: 0.777818684285714

 $00:59:06.827 \longrightarrow 00:59:09.739$ COVID-19 case in the US and then the

NOTE Confidence: 0.777818684285714

00:59:09.813 --> 00:59:12.173 follow up survey wave two or refer to

NOTE Confidence: 0.777818684285714

 $00{:}59{:}12.173 \dashrightarrow 00{:}59{:}14.426$ a pre pandemic survey was conducted

NOTE Confidence: 0.777818684285714

 $00:59:14.426 \longrightarrow 00:59:16.496$ nine months into the pandemic.

NOTE Confidence: 0.777818684285714

 $00{:}59{:}16.500 \dashrightarrow 00{:}59{:}17.160$ Next line.

NOTE Confidence: 0.828337100714286

 $00{:}59{:}19.230 \dashrightarrow 00{:}59{:}20.830$ Suicidal ideations was measured,

NOTE Confidence: 0.828337100714286

 $00:59:20.830 \longrightarrow 00:59:23.966$ but the two items adapted from page tonight

NOTE Confidence: 0.828337100714286

 $00{:}59{:}23.966 \dashrightarrow 00{:}59{:}26.654$ at 9:00 and purpose in life was assessed

00:59:26.723 --> 00:59:29.387 using the purpose in life test short form.

NOTE Confidence: 0.828337100714286

 $00:59:29.390 \dashrightarrow 00:59:31.980$ Here's a sample item here and also

NOTE Confidence: 0.828337100714286

 $00:59:31.980 \longrightarrow 00:59:34.213$ be gathered kovin related variables

NOTE Confidence: 0.828337100714286

00:59:34.213 --> 00:59:36.843 including COVID-19 infection status that

NOTE Confidence: 0.828337100714286

 $00:59:36.843 \longrightarrow 00:59:41.810$ was reported by Self Report next line.

NOTE Confidence: 0.828337100714286

 $00:59:41.810 \longrightarrow 00:59:44.180$ We ran multivariable logistic regression

NOTE Confidence: 0.828337100714286

 $00:59:44.180 \longrightarrow 00:59:47.011$ analysis as well as interaction analysis

NOTE Confidence: 0.828337100714286

 $00{:}59{:}47.011 \dashrightarrow 00{:}59{:}49.776$ of COVID-19 infection by age and also

NOTE Confidence: 0.828337100714286

 $00{:}59{:}49.776 \dashrightarrow 00{:}59{:}51.640$ by protective psychosocial factors

NOTE Confidence: 0.828337100714286

 $00:59:51.640 \dashrightarrow 00:59:56.130$ based on prior literature next line.

NOTE Confidence: 0.828337100714286

 $00:59:56.130 \longrightarrow 00:59:58.776$ The results mean age was 55.2

NOTE Confidence: 0.828337100714286

00:59:58.776 --> 01:00:01.120 predominantly male and white,

NOTE Confidence: 0.828337100714286

 $01:00:01.120 \dashrightarrow 01:00:04.690$ 40% were combat veterans and apparel.

NOTE Confidence: 0.828337100714286

 $01:00:04.690 \longrightarrow 01:00:05.986$ Pandemic assessment.

NOTE Confidence: 0.828337100714286

 $01:00:05.986 \longrightarrow 01:00:09.874$ Almost 20% screen positive for suicidal

 $01:00:09.874 \longrightarrow 01:00:13.086$ ideations and among them 58.9% reported

NOTE Confidence: 0.828337100714286

01:00:13.086 --> 01:00:15.924 both pre and Peri pandemic aside,

NOTE Confidence: 0.828337100714286

 $01:00:15.930 \longrightarrow 01:00:19.365$ an 8.9% developed essay during

NOTE Confidence: 0.828337100714286

 $01:00:19.365 \longrightarrow 01:00:22.113$ the pandemic next slide.

NOTE Confidence: 0.828337100714286

 $01:00:22.120 \longrightarrow 01:00:23.484$ This is the multivariable

NOTE Confidence: 0.828337100714286

 $01:00:23.484 \longrightarrow 01:00:24.848$ regression model we found.

NOTE Confidence: 0.828337100714286

 $01:00:24.850 \longrightarrow 01:00:26.480$ So as you can see,

NOTE Confidence: 0.828337100714286

 $01:00:26.480 \longrightarrow 01:00:28.810$ higher household income and also

NOTE Confidence: 0.828337100714286

 $01{:}00{:}28.810 \dashrightarrow 01{:}00{:}31.140$ greater scores and purpose in

NOTE Confidence: 0.828337100714286

 $01:00:31.222 \longrightarrow 01:00:33.627$ life scale were associated with.

NOTE Confidence: 0.828337100714286

 $01{:}00{:}33.630 \dashrightarrow 01{:}00{:}35.630$ Lower risk of suicidal ideations

NOTE Confidence: 0.828337100714286

01:00:35.630 --> 01:00:36.830 during the pandemic,

NOTE Confidence: 0.828337100714286

 $01:00:36.830 \longrightarrow 01:00:38.918$ whereas Eyecatch greater psychiatric

NOTE Confidence: 0.828337100714286

 $01:00:38.918 \longrightarrow 01:00:42.050$ symptoms severity as well as previous

NOTE Confidence: 0.828337100714286

 $01:00:42.123 \longrightarrow 01:00:45.087$ suicidal behaviors and cover 19 infection

NOTE Confidence: 0.828337100714286

 $01:00:45.087 \longrightarrow 01:00:47.908$ were associated with greater risk of SI.

 $01:00:47.910 \longrightarrow 01:00:50.560$ Next slide.

NOTE Confidence: 0.828337100714286

 $01:00:50.560 \longrightarrow 01:00:52.785$ Interaction analysis showed that among

NOTE Confidence: 0.828337100714286

 $01:00:52.785 \longrightarrow 01:00:55.690$ those who are infected with COVID-19,

NOTE Confidence: 0.828337100714286

 $01:00:55.690 \longrightarrow 01:00:57.760$ those age 45 or older were

NOTE Confidence: 0.828337100714286

 $01:00:57.760 \longrightarrow 01:00:59.569$ more likely to endorse as I,

NOTE Confidence: 0.828337100714286

 $01{:}00{:}59.570 --> 01{:}01{:}02.160$ as you can see during in the 45 to 59

NOTE Confidence: 0.828337100714286

 $01:01:02.230 \longrightarrow 01:01:05.160$ year old bracket, it's close to 60%.

NOTE Confidence: 0.828337100714286

 $01{:}01{:}05.160 \dashrightarrow 01{:}01{:}07.335$ One possible mechanism to this

NOTE Confidence: 0.828337100714286

01:01:07.335 --> 01:01:09.877 finding is that older adults tend

NOTE Confidence: 0.828337100714286

01:01:09.877 --> 01:01:12.590 to have more severe illness courses,

NOTE Confidence: 0.828337100714286

01:01:12.590 --> 01:01:14.912 or also when they're infected they

NOTE Confidence: 0.828337100714286

 $01:01:14.912 \longrightarrow 01:01:17.411$ may suffer more anticipo Tori anxiety

NOTE Confidence: 0.828337100714286

 $01{:}01{:}17.411 \dashrightarrow 01{:}01{:}20.063$ because of the possible higher mortality.

NOTE Confidence: 0.828337100714286 01:01:20.070 --> 01:01:22.600 Next line. NOTE Confidence: 0.828337100714286

01:01:22.600 --> 01:01:24.445 Another interaction analysis found that

01:01:24.445 --> 01:01:26.819 among those were infected with covin 19,

NOTE Confidence: 0.828337100714286

 $01:01:26.820 \longrightarrow 01:01:29.142$ those in the lowest quartile of

NOTE Confidence: 0.828337100714286

 $01:01:29.142 \longrightarrow 01:01:31.702$ purpose in life score almost 80%,

NOTE Confidence: 0.828337100714286

 $01:01:31.702 \longrightarrow 01:01:33.220$ indoors suicidal ideations,

NOTE Confidence: 0.828337100714286

 $01:01:33.220 \longrightarrow 01:01:36.808$ and during the pandemic next line.

NOTE Confidence: 0.828337100714286

01:01:36.810 --> 01:01:39.160 Policy clinical implications of our

NOTE Confidence: 0.828337100714286

 $01:01:39.160 \longrightarrow 01:01:42.003$ findings that veterans age 45 or

NOTE Confidence: 0.828337100714286

 $01:01:42.003 \longrightarrow 01:01:44.098$ older with COVID-19 infection and

NOTE Confidence: 0.828337100714286

01:01:44.098 --> 01:01:46.342 also has pre existing psychiatric

NOTE Confidence: 0.828337100714286

01:01:46.342 --> 01:01:48.797 disorders may require more close

NOTE Confidence: 0.828337100714286

 $01{:}01{:}48.797 \dashrightarrow 01{:}01{:}51.508$ monitoring as an policy measures to

NOTE Confidence: 0.828337100714286

 $01:01:51.508 \longrightarrow 01:01:52.984$ mitigate financial stress interventions

NOTE Confidence: 0.828337100714286

 $01:01:52.984 \longrightarrow 01:01:54.770$ to enhance purpose in life.

NOTE Confidence: 0.828337100714286

01:01:54.770 --> 01:01:56.084 Size chess acceptance,

NOTE Confidence: 0.828337100714286

 $01:01:56.084 \longrightarrow 01:01:58.274$ commitment therapy logotherapy as well

NOTE Confidence: 0.828337100714286

 $01:01:58.274 \longrightarrow 01:02:01.044$ as chaplain care which is known to

 $01:02:01.044 \longrightarrow 01:02:02.874$ enhance religiosity or in spirituality

NOTE Confidence: 0.828337100714286

 $01:02:02.937 \longrightarrow 01:02:04.647$ which is closely associated with

NOTE Confidence: 0.828337100714286

01:02:04.647 --> 01:02:07.278 purpose in life may help mitigate suicide.

NOTE Confidence: 0.828337100714286

01:02:07.278 --> 01:02:09.573 Risk behavior risk in veterans

NOTE Confidence: 0.828337100714286

01:02:09.573 --> 01:02:10.950 during the pandemic.

NOTE Confidence: 0.828337100714286 01:02:10.950 --> 01:02:12.970 Nick lied. NOTE Confidence: 0.828337100714286

01:02:12.970 --> 01:02:14.870 Future directions it's Doctor Pietrzak

NOTE Confidence: 0.828337100714286

 $01:02:14.870 \longrightarrow 01:02:17.230$ measured with Doctor Pietrzak angoul counter.

NOTE Confidence: 0.828337100714286

 $01:02:17.230 \longrightarrow 01:02:18.460$ I'll be applying for the VA,

NOTE Confidence: 0.828337100714286

 $01:02:18.460 \longrightarrow 01:02:22.372$ CDA this fall with plans with proposal to

NOTE Confidence: 0.828337100714286

 $01:02:22.372 \longrightarrow 01:02:25.119$ identify modifiable psychosocial factors.

NOTE Confidence: 0.828337100714286

 $01:02:25.120 \longrightarrow 01:02:27.196$ Then we interact with.

NOTE Confidence: 0.828337100714286

 $01{:}02{:}27.196 \dashrightarrow 01{:}02{:}29.791$ G was derived polygenic risk

NOTE Confidence: 0.828337100714286

 $01:02:29.791 \longrightarrow 01:02:32.553$ scores of suicidality as Westworld

NOTE Confidence: 0.828337100714286

 $01:02:32.553 \longrightarrow 01:02:34.719$ ssed and as a starter.

 $01:02:34.719 \longrightarrow 01:02:37.260$ We just submitted a paper to the

NOTE Confidence: 0.828337100714286

01:02:37.342 --> 01:02:39.868 journal looking at a seven year.

NOTE Confidence: 0.828337100714286

 $01:02:39.870 \longrightarrow 01:02:42.712$ Prospective cord of an HRV S looking

NOTE Confidence: 0.828337100714286

 $01:02:42.712 \longrightarrow 01:02:45.880$ at PRS by psychosocial factors.

NOTE Confidence: 0.828337100714286 01:02:45.880 --> 01:02:48.110 Next line. NOTE Confidence: 0.828337100714286

 $01:02:48.110 \longrightarrow 01:02:49.600$ These are the findings we

NOTE Confidence: 0.828337100714286

01:02:49.600 --> 01:02:51.420 found and as you can see,

NOTE Confidence: 0.828337100714286

 $01:02:51.420 \longrightarrow 01:02:54.040$ those with higher suicidality

NOTE Confidence: 0.828337100714286

 $01{:}02{:}54.040 \dashrightarrow 01{:}02{:}57.315$ collision at risk for suicidality.

NOTE Confidence: 0.828337100714286

 $01:02:57.320 \longrightarrow 01:03:00.265$ And also endorsed lower dispositional

NOTE Confidence: 0.828337100714286

 $01{:}03{:}00.265 --> 01{:}03{:}03.747$ optimism and lower social support were

NOTE Confidence: 0.828337100714286

01:03:03.747 --> 01:03:06.435 more likely to endorse Chronicus I

NOTE Confidence: 0.828337100714286

 $01:03:06.435 \longrightarrow 01:03:09.442$ or develop new onset SI respectively

NOTE Confidence: 0.828337100714286

 $01{:}03{:}09.442 \dashrightarrow 01{:}03{:}12.592$ during the seven year study period.

NOTE Confidence: 0.771111305

 $01:03:12.600 \longrightarrow 01:03:17.390$ I'll put this papers under review next line.

NOTE Confidence: 0.771111305

 $01:03:17.390 \longrightarrow 01:03:18.972$ We would like to thank the veterans

01:03:18.972 --> 01:03:19.940 participating in our study,

NOTE Confidence: 0.771111305

 $01{:}03{:}19.940 \to 01{:}03{:}22.055$ especially with Memorial Day coming

NOTE Confidence: 0.771111305

 $01:03:22.055 \longrightarrow 01:03:24.605$ around and also our collaborators as

NOTE Confidence: 0.771111305

 $01:03:24.605 \longrightarrow 01:03:27.181$ well as the crew addictions that country

NOTE Confidence: 0.771111305

 $01:03:27.181 \longrightarrow 01:03:28.810$ fellowship crew, including doctors,

NOTE Confidence: 0.771111305

 $01{:}03{:}28.810 \dashrightarrow 01{:}03{:}31.480$ meaning that raucous thank you everyone.

NOTE Confidence: 0.8094771

01:03:35.390 --> 01:03:38.230 Wonderful thank you one week, really.

NOTE Confidence: 0.7182042

 $01:03:39.510 \longrightarrow 01:03:42.910$ Wow, wait. We have time crunch.

NOTE Confidence: 0.860790427142857

 $01{:}03{:}45.090 \dashrightarrow 01{:}03{:}47.134$ Important to get this out so we're

NOTE Confidence: 0.841051909375

01:03:47.560 --> 01:03:48.619 very timely. Alright,

NOTE Confidence: 0.841051909375

01:03:48.619 --> 01:03:51.611 so we'll move on to our last speaker

NOTE Confidence: 0.841051909375

 $01:03:51.611 \longrightarrow 01:03:53.846$ in our last honorable mention,

NOTE Confidence: 0.841051909375

 $01:03:53.850 \longrightarrow 01:03:56.270$ and this is Ryan O'dell.

NOTE Confidence: 0.841051909375

01:03:56.270 --> 01:03:57.710 He'll be introduced by his mentor,

NOTE Confidence: 0.841051909375

01:03:57.710 --> 01:03:58.739 Chris Van ****.

 $01:04:01.060 \longrightarrow 01:04:03.908$ Great thank you Chris and I want to

NOTE Confidence: 0.875938383529412

 $01{:}04{:}03.908 \dashrightarrow 01{:}04{:}06.656$ congratulate all of the awards and I'm

NOTE Confidence: 0.875938383529412

01:04:06.656 --> 01:04:09.393 especially honored to introduce Ryan O'dell,

NOTE Confidence: 0.875938383529412

 $01:04:09.393 \longrightarrow 01:04:13.817$ whom we felt very fortunate to have as

NOTE Confidence: 0.875938383529412

 $01:04:13.817 \longrightarrow 01:04:17.046$ a member of our research team with the

NOTE Confidence: 0.875938383529412

01:04:17.046 --> 01:04:19.471 Alzheimer's Research Unit and Alzheimer's

NOTE Confidence: 0.875938383529412

 $01:04:19.471 \longrightarrow 01:04:23.230$ Research Center for the past three years.

NOTE Confidence: 0.875938383529412

01:04:23.230 --> 01:04:24.570 Starting with this case,

NOTE Confidence: 0.875938383529412

 $01:04:24.570 \longrightarrow 01:04:27.405$ rotation and then continuing throughout this

NOTE Confidence: 0.875938383529412

 $01:04:27.405 \longrightarrow 01:04:32.000$ Presidency through the N R&R TP program

Ryan.

NOTE Confidence: 0.875938383529412

01:04:32.000 --> 01:04:33.056 As you will see,

NOTE Confidence: 0.875938383529412

01:04:33.056 --> 01:04:36.488 has focused his research in Nuro pet imaging,

NOTE Confidence: 0.875938383529412

 $01:04:36.488 \longrightarrow 01:04:41.009$ in which he is Co mentored by Adam Mecca,

NOTE Confidence: 0.875938383529412

 $01:04:41.010 \longrightarrow 01:04:43.580$ an in which he's proven to be a very,

NOTE Confidence: 0.875938383529412

 $01:04:43.580 \longrightarrow 01:04:48.278$ very quick study of complex neuroimaging

 $01:04:48.280 \longrightarrow 01:04:50.050$ methodology's in statistics,

NOTE Confidence: 0.875938383529412

 $01:04:50.050 \longrightarrow 01:04:52.242$ and I think his research abilities will speak

NOTE Confidence: 0.875938383529412

 $01:04:52.242 \longrightarrow 01:04:54.126$ for themselves through the paper that Hill.

NOTE Confidence: 0.875938383529412

 $01:04:54.126 \longrightarrow 01:04:57.138$ Be presenting to you.

NOTE Confidence: 0.875938383529412

 $01:04:57.140 \longrightarrow 01:04:59.954$ But Ryan, I wanted to really emphasize

NOTE Confidence: 0.875938383529412

01:04:59.954 --> 01:05:03.260 as a person of extraordinary ability,

NOTE Confidence: 0.875938383529412

 $01:05:03.260 \longrightarrow 01:05:05.672$ curiosity, and dedication,

NOTE Confidence: 0.875938383529412

 $01{:}05{:}05{:}05{:}05{:}05{:}10.496$ but also compassion as a clinician.

NOTE Confidence: 0.875938383529412

01:05:10.500 --> 01:05:14.868 And also you know rare generosity as he

NOTE Confidence: 0.875938383529412

 $01:05:14.868 \longrightarrow 01:05:18.151$ regularly shares his his knowledge and

NOTE Confidence: 0.875938383529412

 $01{:}05{:}18.151 \dashrightarrow 01{:}05{:}22.840$ experience with our students with our.

NOTE Confidence: 0.875938383529412

 $01:05:22.840 \longrightarrow 01:05:24.384$ Staff and with collaborators.

NOTE Confidence: 0.875938383529412

 $01:05:24.384 \longrightarrow 01:05:26.700$ And I think he certainly has

NOTE Confidence: 0.875938383529412

 $01:05:26.773 \longrightarrow 01:05:28.039$ a brilliant future.

NOTE Confidence: 0.875938383529412

01:05:28.040 --> 01:05:28.190 As NOTE Confidence: 0.90567448

01:05:28.200 --> 01:05:30.744 a physician, scientist and

 $01:05:30.744 \longrightarrow 01:05:32.788$ teacher, but maybe at least as

NOTE Confidence: 0.900792508

 $01:05:32.800 \longrightarrow 01:05:35.060$ importantly as a new father.

NOTE Confidence: 0.900792508

 $01:05:35.060 \longrightarrow 01:05:37.395$ So multiple. Congratulations

NOTE Confidence: 0.900792508

 $01:05:37.395 \longrightarrow 01:05:41.085$ to Ryan and also to Milda.

NOTE Confidence: 0.900792508

 $01:05:41.090 \longrightarrow 01:05:44.340$ So take it away, Ryan. Thank

NOTE Confidence: 0.874919461428571

01:05:44.350 --> 01:05:47.087 you Chris for that very kind introduction.

NOTE Confidence: 0.874919461428571

01:05:47.090 --> 01:05:48.962 And before I begin I also want to

NOTE Confidence: 0.874919461428571

 $01{:}05{:}48.962 \dashrightarrow 01{:}05{:}51.090$ thank the Lussman family as well as

NOTE Confidence: 0.874919461428571

 $01:05:51.090 \longrightarrow 01:05:52.695$ the award selection committee for

NOTE Confidence: 0.874919461428571

 $01:05:52.751 \longrightarrow 01:05:54.707$ this opportunity to present my work.

NOTE Confidence: 0.874919461428571

01:05:54.710 --> 01:05:57.094 My recent work using a novel pet imaging

NOTE Confidence: 0.874919461428571

 $01:05:57.094 \longrightarrow 01:05:59.197$ tracer to characterize the relationship

NOTE Confidence: 0.874919461428571

 $01{:}05{:}59.197 \dashrightarrow 01{:}06{:}01.607$ between amyloid accumulation and synaptic

NOTE Confidence: 0.874919461428571

 $01{:}06{:}01.607 \dashrightarrow 01{:}06{:}04.188$ health in early Alzheimer's disease.

NOTE Confidence: 0.874919461428571

01:06:04.190 --> 01:06:07.725 Next slide, I have no personal disclosures,

 $01:06:07.730 \longrightarrow 01:06:08.888$ so next slide.

NOTE Confidence: 0.874919461428571

 $01:06:08.888 \longrightarrow 01:06:11.590$ And so just to dive right in.

NOTE Confidence: 0.874919461428571

 $01:06:11.590 \longrightarrow 01:06:13.738$ So synaptic loss has been demonstrated

NOTE Confidence: 0.874919461428571

 $01:06:13.738 \longrightarrow 01:06:15.788$ both as an early pathological

NOTE Confidence: 0.874919461428571

 $01:06:15.788 \longrightarrow 01:06:17.976$ event in Alzheimer's disease,

NOTE Confidence: 0.874919461428571

01:06:17.980 --> 01:06:20.932 but also a significant major structural

NOTE Confidence: 0.874919461428571

 $01:06:20.932 \longrightarrow 01:06:22.900$ correlate with cognitive impairment.

NOTE Confidence: 0.874919461428571

 $01:06:22.900 \longrightarrow 01:06:25.378$ An as synaptic loss in Alzheimer's

NOTE Confidence: 0.874919461428571

 $01:06:25.378 \longrightarrow 01:06:27.561$ disease has been investigated primarily

NOTE Confidence: 0.874919461428571

 $01:06:27.561 \longrightarrow 01:06:30.327$ via postmortem and brain biopsy studies.

NOTE Confidence: 0.874919461428571

 $01:06:30.330 \longrightarrow 01:06:32.268$ The ability to measure synaptic density

NOTE Confidence: 0.874919461428571

 $01:06:32.268 \longrightarrow 01:06:35.170$ in vivo would not only allow for a

NOTE Confidence: 0.874919461428571

 $01:06:35.170 \longrightarrow 01:06:37.100$ more complete understanding of synaptic

NOTE Confidence: 0.874919461428571

01:06:37.100 --> 01:06:38.910 alterations in early disease stages,

NOTE Confidence: 0.874919461428571

 $01:06:38.910 \longrightarrow 01:06:40.828$ but would also be a great utility.

NOTE Confidence: 0.874919461428571

 $01:06:40.830 \longrightarrow 01:06:43.357$ For tracking a deep regression and also

01:06:43.357 --> 01:06:45.202 monitoring the efficacy of potential

NOTE Confidence: 0.874919461428571

 $01:06:45.202 \longrightarrow 01:06:47.308$ therapies in clinical trials and so,

NOTE Confidence: 0.874919461428571

01:06:47.310 --> 01:06:49.434 one suitable target is the synaptic

NOTE Confidence: 0.874919461428571

01:06:49.434 --> 01:06:51.393 vesicle glycoprotein 2 which is circled

NOTE Confidence: 0.874919461428571

 $01:06:51.393 \longrightarrow 01:06:53.730$ in blue in the bottom left of the slide.

NOTE Confidence: 0.874919461428571

 $01:06:53.730 \longrightarrow 01:06:56.194$ This is an essential component of synaptic

NOTE Confidence: 0.874919461428571

 $01:06:56.194 \longrightarrow 01:06:58.319$ vesicles is located in the presynaptic

NOTE Confidence: 0.874919461428571

 $01:06:58.319 \longrightarrow 01:07:00.329$ terminals and one of its isoforms.

NOTE Confidence: 0.874919461428571

01:07:00.330 --> 01:07:01.930 SV 2A is ubiquitously expressed

NOTE Confidence: 0.874919461428571

 $01:07:01.930 \longrightarrow 01:07:03.909$ in almost all of the synapses

NOTE Confidence: 0.874919461428571

 $01:07:03.909 \longrightarrow 01:07:06.065$ in the CNS and could be useful.

NOTE Confidence: 0.874919461428571

 $01{:}07{:}06.070 \dashrightarrow 01{:}07{:}07.438$ Useful biomarker for synaptic

NOTE Confidence: 0.874919461428571

 $01:07:07.438 \longrightarrow 01:07:09.490$ density and so to that end,

NOTE Confidence: 0.874919461428571

 $01:07:09.490 \longrightarrow 01:07:10.967$ such a tracer known as you see,

NOTE Confidence: 0.874919461428571

 $01:07:10.970 \longrightarrow 01:07:12.836$ BJ's shown in the bottom right.

01:07:12.840 --> 01:07:14.886 Has been developed for quantitative SV

NOTE Confidence: 0.874919461428571

01:07:14.886 --> 01:07:17.528 2A pet imaging at the Yale Pet Center.

NOTE Confidence: 0.874919461428571

 $01:07:17.530 \longrightarrow 01:07:18.664$ Next slide please.

NOTE Confidence: 0.874919461428571

01:07:18.664 --> 01:07:20.932 So in our previous work with

NOTE Confidence: 0.874919461428571

01:07:20.932 --> 01:07:22.429 UCB JPEG image Ng,

NOTE Confidence: 0.874919461428571

 $01:07:22.430 \longrightarrow 01:07:24.178$ we've demonstrated widespread reductions

NOTE Confidence: 0.874919461428571

 $01:07:24.178 \longrightarrow 01:07:26.363$ in synaptic density in Alzheimer's

NOTE Confidence: 0.874919461428571

 $01:07:26.363 \longrightarrow 01:07:28.330$ disease in the medial temporal lobe,

NOTE Confidence: 0.874919461428571

 $01{:}07{:}28.330 \dashrightarrow 01{:}07{:}29.482$ and also neocortical regions,

NOTE Confidence: 0.874919461428571

 $01:07:29.482 \longrightarrow 01:07:32.311$ and on the left this is a slide from

NOTE Confidence: 0.874919461428571

 $01{:}07{:}32.311 \dashrightarrow 01{:}07{:}34.051$ a recent publication that displays

NOTE Confidence: 0.874919461428571

 $01:07:34.051 \longrightarrow 01:07:35.761$ average coronial images of synaptic

NOTE Confidence: 0.874919461428571

01:07:35.761 --> 01:07:37.361 density for 19 cognitively normal

NOTE Confidence: 0.874919461428571

 $01:07:37.361 \longrightarrow 01:07:39.630$ on the left and 34 Alzheimer's

NOTE Confidence: 0.874919461428571

01:07:39.630 --> 01:07:41.655 disease participants on the right,

NOTE Confidence: 0.874919461428571

 $01:07:41.660 \longrightarrow 01:07:43.884$ and you can see visibly reduced you CBJ

 $01:07:43.884 \longrightarrow 01:07:45.897$ binding in the medial temporal lobe,

NOTE Confidence: 0.874919461428571

 $01:07:45.900 \longrightarrow 01:07:48.940$ which is the bottom row of corona sections.

NOTE Confidence: 0.874919461428571

 $01:07:48.940 \longrightarrow 01:07:50.704$ But you can also see there's a

NOTE Confidence: 0.874919461428571

01:07:50.704 --> 01:07:52.183 reduction in synaptic density throughout

NOTE Confidence: 0.874919461428571

 $01:07:52.183 \longrightarrow 01:07:53.848$ the NEO cortex and subcortex.

NOTE Confidence: 0.874919461428571

01:07:53.850 --> 01:07:55.800 Which we have quantified below,

NOTE Confidence: 0.874919461428571

01:07:55.800 --> 01:07:58.670 and although this study did seek out,

NOTE Confidence: 0.874919461428571 01:07:58.670 --> 01:07:59.432 you know,

NOTE Confidence: 0.874919461428571

 $01:07:59.432 \longrightarrow 01:08:01.337$ to fully characterize the extent

NOTE Confidence: 0.874919461428571

 $01:08:01.337 \dashrightarrow 01:08:03.411$ of synaptic alterations in early AD

NOTE Confidence: 0.874919461428571

01:08:03.411 --> 01:08:05.145 using SV2 Apetit did leave unclear

NOTE Confidence: 0.874919461428571

 $01:08:05.145 \longrightarrow 01:08:07.225$ the relationship of these synaptic

NOTE Confidence: 0.874919461428571

 $01:08:07.225 \longrightarrow 01:08:08.957$ alterations with more traditional

NOTE Confidence: 0.874919461428571

01:08:08.957 --> 01:08:10.766 markers of 80 pathology,

NOTE Confidence: 0.874919461428571

01:08:10.766 --> 01:08:13.574 specifically amyloid or a beta deposition,

 $01:08:13.580 \longrightarrow 01:08:15.080$ and so therefore in the present

NOTE Confidence: 0.874919461428571

01:08:15.080 --> 01:08:16.841 study we set out to characterize

NOTE Confidence: 0.874919461428571

 $01:08:16.841 \longrightarrow 01:08:18.611$ the relationship between a measure

NOTE Confidence: 0.874919461428571

 $01:08:18.611 \longrightarrow 01:08:20.556$ of global amyloid deposition and SV

NOTE Confidence: 0.874919461428571

01:08:20.556 --> 01:08:22.488 two way binding in early Ady across

NOTE Confidence: 0.874919461428571

01:08:22.490 --> 01:08:24.248 a broad range of cortical regions.

NOTE Confidence: 0.874919461428571 01:08:24.250 --> 01:08:24.932 Next slide.

NOTE Confidence: 0.874919461428571

01:08:24.932 --> 01:08:28.350 And so, in the era of amyloid PET imaging,

NOTE Confidence: 0.874919461428571

01:08:28.350 --> 01:08:30.606 longitudinal studies have generally

NOTE Confidence: 0.874919461428571

 $01:08:30.606 \longrightarrow 01:08:32.298$ demonstrated that continued.

NOTE Confidence: 0.874919461428571

 $01:08:32.300 \longrightarrow 01:08:34.056$ There's continued amyloid accumulation

NOTE Confidence: 0.874919461428571

01:08:34.056 --> 01:08:36.251 throughout the prodromal or mild

NOTE Confidence: 0.874919461428571

 $01:08:36.251 \longrightarrow 01:08:38.088$ cognitive impairment stages of

NOTE Confidence: 0.874919461428571

01:08:38.088 --> 01:08:38.978 Alzheimer's disease,

NOTE Confidence: 0.874919461428571

 $01:08:38.980 \longrightarrow 01:08:41.507$ with minimal change by the time of

NOTE Confidence: 0.874919461428571

 $01:08:41.507 \longrightarrow 01:08:44.108$ conversion to a dementia next slide.

 $01:08:44.110 \longrightarrow 01:08:45.730$ And additionally limited postmortem

NOTE Confidence: 0.874919461428571

 $01:08:45.730 \longrightarrow 01:08:48.160$ work in these prodromal or mild

NOTE Confidence: 0.825297698571428

 $01{:}08{:}48.221 \dashrightarrow 01{:}08{:}50.001$ Adie stages has demonstrated the

NOTE Confidence: 0.825297698571428

 $01:08:50.001 \longrightarrow 01:08:52.572$ hippocampus to be the site of the

NOTE Confidence: 0.825297698571428

 $01:08:52.572 \longrightarrow 01:08:54.744$ earliest and most profound synaptic loss.

NOTE Confidence: 0.825297698571428

01:08:54.750 --> 01:08:56.970 Next slide and so therefore,

NOTE Confidence: 0.825297698571428

 $01:08:56.970 \longrightarrow 01:08:59.810$ in the prodromal stage of a D when

NOTE Confidence: 0.825297698571428

 $01:08:59.810 \longrightarrow 01:09:02.230$ amyloid plaques are still accumulating,

NOTE Confidence: 0.825297698571428

 $01:09:02.230 \longrightarrow 01:09:04.309$ we might expect them to be associated

NOTE Confidence: 0.825297698571428

01:09:04.309 --> 01:09:06.159 with industries of disease severity,

NOTE Confidence: 0.825297698571428

 $01{:}09{:}06.160 \dashrightarrow 01{:}09{:}07.716$ which includes synaptic loss,

NOTE Confidence: 0.825297698571428

 $01:09:07.716 \longrightarrow 01:09:09.661$ particularly in those brain regions

NOTE Confidence: 0.825297698571428

 $01{:}09{:}09.661 \dashrightarrow 01{:}09{:}11.890$ that show marked early synaptic loss,

NOTE Confidence: 0.825297698571428

 $01:09:11.890 \longrightarrow 01:09:14.836$ such as the hippocampus. Next slide.

NOTE Confidence: 0.825297698571428

01:09:14.840 --> 01:09:17.036 Oh yes, there's the primary hypothesis,

 $01:09:17.040 \longrightarrow 01:09:21.920$ Yep, so in in this study we have

NOTE Confidence: 0.825297698571428

 $01:09:21.920 \longrightarrow 01:09:23.555$ recruited participants between the age

NOTE Confidence: 0.825297698571428

 $01:09:23.555 \longrightarrow 01:09:26.776$ of 55 and 85 years old that either had

NOTE Confidence: 0.825297698571428

 $01:09:26.776 \longrightarrow 01:09:28.686$ normal cognition or Alzheimer's disease.

NOTE Confidence: 0.825297698571428

01:09:28.690 --> 01:09:30.414 The cognitively normal participants,

NOTE Confidence: 0.825297698571428

01:09:30.414 --> 01:09:32.569 Ramel Lloyd negative NAD participants

NOTE Confidence: 0.825297698571428

01:09:32.569 --> 01:09:34.485 either had mild dementia or

NOTE Confidence: 0.825297698571428

01:09:34.485 --> 01:09:35.556 mild cognitive impairment,

NOTE Confidence: 0.825297698571428

 $01:09:35.560 \longrightarrow 01:09:37.310$ and all were employed positive.

NOTE Confidence: 0.825297698571428

01:09:37.310 --> 01:09:39.326 We perform pipet for brain amyloid.

NOTE Confidence: 0.825297698571428

01:09:39.330 --> 01:09:41.810 You see BJ to measure the synaptic density

NOTE Confidence: 0.825297698571428

 $01:09:41.810 \longrightarrow 01:09:44.744$ and we did MRI for volumetric segmentation

NOTE Confidence: 0.825297698571428

 $01:09:44.744 \longrightarrow 01:09:47.320$ and ROI determination using free surfer.

NOTE Confidence: 0.825297698571428

 $01:09:47.320 \longrightarrow 01:09:49.024$ Parameters and the model parameters that

NOTE Confidence: 0.825297698571428

01:09:49.024 --> 01:09:51.258 I'm going to be reporting are distribution

NOTE Confidence: 0.825297698571428

 $01:09:51.258 \longrightarrow 01:09:53.701$ volume ratios that use a whole cerebellum.

 $01:09:53.710 \longrightarrow 01:09:55.430$ Reference region for both tracers.

NOTE Confidence: 0.825297698571428 01:09:55.430 --> 01:09:56.276 Next slide. NOTE Confidence: 0.825297698571428

 $01:09:56.276 \longrightarrow 01:09:58.814$ So this is some demographic information.

NOTE Confidence: 0.825297698571428

01:09:58.820 --> 01:10:01.130 The sample consisted of 19 cognitively

NOTE Confidence: 0.825297698571428

 $01:10:01.130 \longrightarrow 01:10:03.715$ normal 14 amnestic mild cognitive impairment

NOTE Confidence: 0.825297698571428

 $01:10:03.715 \longrightarrow 01:10:06.155$ and 24 mild dementia participants.

NOTE Confidence: 0.825297698571428

 $01:10:06.160 \longrightarrow 01:10:08.304$ It was well balanced for age and sex

NOTE Confidence: 0.825297698571428

 $01:10:08.304 \longrightarrow 01:10:09.526$ and demonstrated slightly decreased

NOTE Confidence: 0.825297698571428

01:10:09.526 --> 01:10:11.410 years of education in the dementia

NOTE Confidence: 0.825297698571428

 $01:10:11.410 \longrightarrow 01:10:13.357$ group as compared to the CN Group.

NOTE Confidence: 0.825297698571428

01:10:13.360 --> 01:10:14.836 I have that highlighted in red,

NOTE Confidence: 0.825297698571428

 $01:10:14.840 \longrightarrow 01:10:17.122$ but overall we do see expected group

NOTE Confidence: 0.825297698571428

 $01{:}10{:}17.122 \dashrightarrow 01{:}10{:}19.511$ differences in measures of disease stage as

NOTE Confidence: 0.825297698571428

 $01:10:19.511 \longrightarrow 01:10:21.533$ indicated by the clinical dementia rating.

NOTE Confidence: 0.825297698571428

 $01:10:21.540 \longrightarrow 01:10:22.776$ Some boxes score.

01:10:22.776 --> 01:10:25.660 Global cognition is shown with the MSE,

NOTE Confidence: 0.825297698571428

 $01{:}10{:}25.660 {\:{\mbox{--}}\!>}\ 01{:}10{:}28.068$ an episodic memory as shown is an average

NOTE Confidence: 0.825297698571428

01:10:28.068 --> 01:10:31.102 of the logical memory 2IN revolt delay.

NOTE Confidence: 0.825297698571428

 $01:10:31.102 \longrightarrow 01:10:34.107$ Onoro psychological tests next slide.

NOTE Confidence: 0.825297698571428

 $01:10:34.110 \longrightarrow 01:10:36.441$ So then looking at our primary analysis

NOTE Confidence: 0.825297698571428

 $01:10:36.441 \longrightarrow 01:10:38.312$ of the association between global

NOTE Confidence: 0.825297698571428

01:10:38.312 --> 01:10:40.730 amyloid deposition and hippocampal SV 2A,

NOTE Confidence: 0.825297698571428

 $01:10:40.730 \longrightarrow 01:10:43.502$ we see a marginally significant inverse

NOTE Confidence: 0.825297698571428

 $01{:}10{:}43.502 \dashrightarrow 01{:}10{:}45.694$ correlation in participants with MCI

NOTE Confidence: 0.825297698571428

 $01:10:45.694 \longrightarrow 01:10:47.918$ as shown by the green dots in line,

NOTE Confidence: 0.825297698571428

01:10:47.920 --> 01:10:50.384 but not in dementia shown in red,

NOTE Confidence: 0.825297698571428

 $01:10:50.390 \longrightarrow 01:10:52.590$ and this significant correlation did

NOTE Confidence: 0.825297698571428

01:10:52.590 --> 01:10:54.350 survive partial volume correction,

NOTE Confidence: 0.825297698571428

01:10:54.350 --> 01:10:56.975 although I'm not going to be discussing

NOTE Confidence: 0.825297698571428

 $01:10:56.975 \longrightarrow 01:10:58.621$ that technique and methodology

NOTE Confidence: 0.825297698571428

01:10:58.621 --> 01:10:59.980 here next slide.

 $01:10:59.980 \longrightarrow 01:11:01.820$ And we can also see this difference in

NOTE Confidence: 0.825297698571428

 $01:11:01.820 \longrightarrow 01:11:02.967$ the correlation coefficients between

NOTE Confidence: 0.825297698571428

01:11:02.967 --> 01:11:04.577 the true groups was significant,

NOTE Confidence: 0.825297698571428

 $01:11:04.580 \longrightarrow 01:11:06.946$ as assessed by the Fisher Z transform

NOTE Confidence: 0.825297698571428

 $01:11:06.946 \longrightarrow 01:11:09.450$ with a one tailed P value next.

NOTE Confidence: 0.825297698571428

01:11:09.450 --> 01:11:10.590 And so finally,

NOTE Confidence: 0.825297698571428

01:11:10.590 --> 01:11:11.730 surrounding our exploratory

NOTE Confidence: 0.825297698571428

 $01:11:11.730 \longrightarrow 01:11:13.763$ analysis of the association between

NOTE Confidence: 0.825297698571428

01:11:13.763 --> 01:11:15.128 global amyloid deposition,

NOTE Confidence: 0.825297698571428

 $01{:}11{:}15.130 \dashrightarrow 01{:}11{:}17.328$ an regional S V2 and the remaining

NOTE Confidence: 0.825297698571428

01:11:17.328 --> 01:11:18.270 medial temporal structures,

NOTE Confidence: 0.825297698571428

01:11:18.270 --> 01:11:20.102 amygdala, and to rhino,

NOTE Confidence: 0.825297698571428

 $01{:}11{:}20.102 \dashrightarrow 01{:}11{:}21.476$ in parahippocampal cortices,

NOTE Confidence: 0.825297698571428

 $01:11:21.480 \longrightarrow 01:11:23.755$ as well as cortical composite are wise.

NOTE Confidence: 0.825297698571428

 $01:11:23.760 \longrightarrow 01:11:25.480$ We do observe many negative

 $01:11:25.480 \longrightarrow 01:11:26.512$ but nonsignificant correlations

NOTE Confidence: 0.825297698571428

01:11:26.512 --> 01:11:27.930 in both participants,

NOTE Confidence: 0.825297698571428

 $01:11:27.930 \longrightarrow 01:11:29.970$ with MCI and mild dementia.

NOTE Confidence: 0.825297698571428

 $01:11:29.970 \longrightarrow 01:11:32.682$ We do have other do see a significant

NOTE Confidence: 0.825297698571428

 $01:11:32.682 \longrightarrow 01:11:34.119$ inverse correlation between global

NOTE Confidence: 0.825297698571428

01:11:34.119 --> 01:11:35.844 amyloid and lateral parietal SV

NOTE Confidence: 0.825297698571428

01:11:35.844 --> 01:11:38.260 2A and mild dementia participants,

NOTE Confidence: 0.825297698571428

 $01:11:38.260 \longrightarrow 01:11:39.403$ but this significant.

NOTE Confidence: 0.825297698571428

01:11:39.403 --> 01:11:40.927 Correlation did not survive

NOTE Confidence: 0.825297698571428

 $01:11:40.927 \longrightarrow 01:11:42.070$ partial volume correction.

NOTE Confidence: 0.825297698571428 01:11:42.070 --> 01:11:42.708 Next slide, NOTE Confidence: 0.825297698571428

 $01:11:42.708 \longrightarrow 01:11:45.260$ and so in conclusion we this is the

NOTE Confidence: 0.825297698571428

01:11:45.333 --> 01:11:47.948 first in vivo study investigating

NOTE Confidence: 0.825297698571428

01:11:47.948 --> 01:11:50.040 the relationship between amyloid

NOTE Confidence: 0.825297698571428

 $01:11:50.040 \longrightarrow 01:11:52.266$ deposition and synaptic alterations

NOTE Confidence: 0.825297698571428

 $01:11:52.266 \longrightarrow 01:11:53.895$ in Alzheimer's disease.

 $01:11:53.900 \longrightarrow 01:11:55.215$ We feel our findings lend

NOTE Confidence: 0.825297698571428

01:11:55.215 --> 01:11:56.530 in vivo support to this

NOTE Confidence: 0.813743420526316

 $01:11:56.590 \longrightarrow 01:11:58.320$ hypothesis that in the earlier

NOTE Confidence: 0.813743420526316

01:11:58.320 --> 01:11:59.704 stages of clinical disease,

NOTE Confidence: 0.813743420526316

01:11:59.710 --> 01:12:01.540 amyloid deposition may still be

NOTE Confidence: 0.813743420526316

 $01:12:01.540 \longrightarrow 01:12:03.370$ accumulating across the broad range

NOTE Confidence: 0.813743420526316

01:12:03.430 --> 01:12:05.596 of cortical regions having yet to

NOTE Confidence: 0.813743420526316

 $01{:}12{:}05.596 \dashrightarrow 01{:}12{:}07.040$ reach this hypothesized plateau,

NOTE Confidence: 0.813743420526316

 $01:12:07.040 \longrightarrow 01:12:09.007$ and we also feel these results are

NOTE Confidence: 0.813743420526316

 $01:12:09.007 \longrightarrow 01:12:10.583$ consistent with prior evidence that

NOTE Confidence: 0.813743420526316

 $01{:}12{:}10.583 \dashrightarrow 01{:}12{:}12.617$ amyloid plaques are not well correlated.

NOTE Confidence: 0.813743420526316

01:12:12.620 --> 01:12:14.456 With the indices of disease severity,

NOTE Confidence: 0.813743420526316

 $01:12:14.460 \longrightarrow 01:12:16.110$ at least in the dementia stage,

NOTE Confidence: 0.813743420526316

01:12:16.110 --> 01:12:17.262 and of course,

NOTE Confidence: 0.813743420526316

 $01:12:17.262 \longrightarrow 01:12:18.798$ to better characterize this

01:12:18.798 --> 01:12:19.950 relationship moving forward,

NOTE Confidence: 0.813743420526316

01:12:19.950 --> 01:12:21.346 we're recruiting or continuing

NOTE Confidence: 0.813743420526316

01:12:21.346 --> 01:12:23.440 to recruit a larger cohort of

NOTE Confidence: 0.813743420526316

 $01:12:23.503 \longrightarrow 01:12:25.945$ participants with MCI and mild dementia

NOTE Confidence: 0.813743420526316

01:12:25.945 --> 01:12:27.573 to be followed longitudinally,

NOTE Confidence: 0.813743420526316

 $01:12:27.580 \longrightarrow 01:12:30.154$ as well as investigating a separate

NOTE Confidence: 0.813743420526316

 $01:12:30.154 \longrightarrow 01:12:31.870$ cohort with preclinical Alzheimer's

NOTE Confidence: 0.813743420526316

 $01{:}12{:}31.933 \dashrightarrow 01{:}12{:}34.123$ disease for longitudinal multi tracer

NOTE Confidence: 0.813743420526316

01:12:34.123 --> 01:12:37.580 PET imaging studies. Next slide.

NOTE Confidence: 0.813743420526316

 $01:12:37.580 \longrightarrow 01:12:38.917$ So that's all I have for today.

NOTE Confidence: 0.813743420526316

01:12:38.920 --> 01:12:40.612 Thank you again for allowing me

NOTE Confidence: 0.813743420526316

01:12:40.612 --> 01:12:42.118 this opportunity to tell everyone

NOTE Confidence: 0.813743420526316

01:12:42.118 --> 01:12:43.406 about our ongoing work.

NOTE Confidence: 0.813743420526316

 $01:12:43.410 \longrightarrow 01:12:45.276$ I really, really can't give enough

NOTE Confidence: 0.813743420526316

01:12:45.276 --> 01:12:46.520 thanks to my faculty,

NOTE Confidence: 0.813743420526316

01:12:46.520 --> 01:12:48.448 mentors, Doctor Vandyken, Dr.

01:12:48.448 --> 01:12:48.930 Mecca,

NOTE Confidence: 0.813743420526316

 $01:12:48.930 \longrightarrow 01:12:50.771$ as well as all of the research

NOTE Confidence: 0.813743420526316

01:12:50.771 --> 01:12:52.356 faculty and staff that have

NOTE Confidence: 0.813743420526316

01:12:52.356 --> 01:12:54.196 contributed contributed to this work,

NOTE Confidence: 0.813743420526316

 $01:12:54.200 \longrightarrow 01:12:55.748$ many of whom are listed here.

NOTE Confidence: 0.813743420526316

01:12:55.750 --> 01:12:56.455 And, of course,

NOTE Confidence: 0.813743420526316

 $01:12:56.455 \longrightarrow 01:12:57.865$ we can't give enough thanks to

NOTE Confidence: 0.813743420526316

 $01{:}12{:}57.865 \dashrightarrow 01{:}12{:}59.134$ the research participants who

NOTE Confidence: 0.813743420526316

 $01{:}12{:}59.134 \dashrightarrow 01{:}13{:}00.779$ generously do nated their time and

NOTE Confidence: 0.813743420526316

01:13:00.779 --> 01:13:02.330 efforts to make these studies

NOTE Confidence: 0.914235496666667

01:13:02.340 --> 01:13:03.888 possible. Thank you.

NOTE Confidence: 0.865959557916666

 $01:13:09.770 \longrightarrow 01:13:12.398$ Thank you Ryan for a great talk and I

NOTE Confidence: 0.865959557916666

 $01{:}13{:}12.398 \dashrightarrow 01{:}13{:}15.074$ think we've really seen an extraordinary

NOTE Confidence: 0.865959557916666

01:13:15.074 --> 01:13:17.384 breadth of wonderful science here

NOTE Confidence: 0.865959557916666

01:13:17.459 --> 01:13:19.499 across many different domains.

 $01:13:19.500 \longrightarrow 01:13:21.600$ It just speaks to the the wonderful

NOTE Confidence: 0.865959557916666

01:13:21.600 --> 01:13:23.763 things that are going on among the

NOTE Confidence: 0.865959557916666

 $01:13:23.763 \longrightarrow 01:13:25.563$ trainees in our department and I

NOTE Confidence: 0.865959557916666

 $01:13:25.625 \longrightarrow 01:13:27.510$ congratulate all of the winners.

NOTE Confidence: 0.865959557916666

01:13:27.510 --> 01:13:28.246 Since we've had 11:30,

NOTE Confidence: 0.865959557916666

01:13:28.246 --> 01:13:29.838 I think we're not going to have time

NOTE Confidence: 0.865959557916666

01:13:29.838 --> 01:13:31.188 for more questions and discussion now,

NOTE Confidence: 0.865959557916666

 $01:13:31.190 \longrightarrow 01:13:33.334$ but I invite you if people have questions

NOTE Confidence: 0.865959557916666

 $01:13:33.334 \longrightarrow 01:13:35.030$ for individual winners and presenters,

NOTE Confidence: 0.865959557916666

 $01:13:35.030 \longrightarrow 01:13:37.936$ please follow up by email with them.

NOTE Confidence: 0.865959557916666

 $01{:}13{:}37.936 \dashrightarrow 01{:}13{:}40.654$ Thank you everyone for being here.

NOTE Confidence: 0.865959557916666

01:13:40.660 --> 01:13:42.599 Thanks again to the last man family

NOTE Confidence: 0.865959557916666

 $01:13:42.599 \longrightarrow 01:13:44.958$ and the last and Family Foundation

NOTE Confidence: 0.865959557916666

01:13:44.958 --> 01:13:46.910 for supporting this wonderful

NOTE Confidence: 0.865959557916666

 $01:13:46.910 \longrightarrow 01:13:47.886$ departmental transition.

NOTE Confidence: 0.865959557916666

01:13:47.890 --> 01:13:48.996 We'll see you all again next year.