WEBVTT

NOTE duration:"01:21:36" NOTE recognizability:0.896

NOTE language:en-us

NOTE Confidence: 0.882581222

 $00:00:00.000 \longrightarrow 00:00:01.620$ But we're still going to have

NOTE Confidence: 0.882581222

 $00:00:01.620 \longrightarrow 00:00:02.700$ people joining for a while,

NOTE Confidence: 0.882581222

 $00:00:02.700 \dashrightarrow 00:00:05.400$ but I I would like to make sure that I leave.

NOTE Confidence: 0.882581222

 $00:00:05.400 \longrightarrow 00:00:07.212$ There's not much time as possible

NOTE Confidence: 0.882581222

00:00:07.212 --> 00:00:09.000 for Doctor Janik to to speak,

NOTE Confidence: 0.882581222

 $00{:}00{:}09.000 \dashrightarrow 00{:}00{:}11.796$ so I will begin our introduction.

NOTE Confidence: 0.882581222

00:00:11.800 --> 00:00:14.957 Those of you who heard The Chieftains,

NOTE Confidence: 0.882581222

 $00:00:14.960 \longrightarrow 00:00:18.986$ this music was in honor of

NOTE Confidence: 0.882581222

00:00:18.986 --> 00:00:20.650 Doctor Flynn's Irish heritage,

NOTE Confidence: 0.882581222

 $00:00:20.650 \dashrightarrow 00:00:22.630$ although I do think The Chieftains

NOTE Confidence: 0.882581222

 $00{:}00{:}22.630 \dashrightarrow 00{:}00{:}25.260$ might be Scottish, as Jane mentioned,

NOTE Confidence: 0.882581222

 $00:00:25.260 \longrightarrow 00:00:30.574$ but the IT was still lively and Celtic, and.

NOTE Confidence: 0.882581222

 $00:00:30.574 \longrightarrow 00:00:35.886$ As as you'll hear Doctor John Patrick Flynn,

00:00:35.890 --> 00:00:39.670 who for whom this lecture is name is named,

NOTE Confidence: 0.882581222

 $00:00:39.670 \dashrightarrow 00:00:42.683$ was a member of the L faculty from 1954

NOTE Confidence: 0.882581222

00:00:42.683 --> 00:00:45.539 until his retirement in July 1979,

NOTE Confidence: 0.882581222

00:00:45.539 --> 00:00:47.873 and he was really an extraordinary

NOTE Confidence: 0.882581222

 $00:00:47.873 \longrightarrow 00:00:50.808$ person who had quite a remarkable life.

NOTE Confidence: 0.882581222

 $00:00:50.810 \longrightarrow 00:00:52.168$ And so I'm going to take a

NOTE Confidence: 0.882581222

 $00:00:52.168 \longrightarrow 00:00:53.650$ little time to tell you about it.

NOTE Confidence: 0.882581222

 $00:00:53.650 \longrightarrow 00:00:55.502$ We have his daughter,

NOTE Confidence: 0.882581222

 $00{:}00{:}55.502 \dashrightarrow 00{:}00{:}57.530$ Sarah Flynn, with us here today.

NOTE Confidence: 0.882581222

00:00:57.530 --> 00:00:59.360 Thank you for coming, Sarah, and.

NOTE Confidence: 0.882581222

 $00{:}00{:}59.360 \dashrightarrow 00{:}01{:}01.915$ She was the one who helped me

NOTE Confidence: 0.882581222

 $00{:}01{:}01.915 \dashrightarrow 00{:}01{:}03.982$ gather the information that I'm

NOTE Confidence: 0.882581222

 $00:01:03.982 \longrightarrow 00:01:06.940$ going to be sharing with you today.

NOTE Confidence: 0.882581222

00:01:06.940 --> 00:01:08.120 So first of all,

NOTE Confidence: 0.882581222

 $00:01:08.120 \longrightarrow 00:01:09.300$ I I knew Dr.

NOTE Confidence: 0.882581222

 $00:01:09.300 \longrightarrow 00:01:11.575$ Finn Flynn's work because of his focus

 $00:01:11.575 \longrightarrow 00:01:13.765$ on the neural basis of aggressive

NOTE Confidence: 0.882581222

 $00{:}01{:}13.765 \dashrightarrow 00{:}01{:}15.675$ behavior and he's recognized as

NOTE Confidence: 0.882581222

 $00{:}01{:}15.675 \dashrightarrow 00{:}01{:}18.202$ a pioneer in neuroscience and in

NOTE Confidence: 0.882581222

 $00:01:18.202 \longrightarrow 00:01:20.327$ general for his contribution to

NOTE Confidence: 0.882581222

 $00:01:20.327 \longrightarrow 00:01:22.760$ understanding the function of the

NOTE Confidence: 0.882581222

 $00:01:22.760 \longrightarrow 00:01:24.760$ hippocampus in the hypothalamus.

NOTE Confidence: 0.882581222

00:01:24.760 --> 00:01:28.030 He also served from 1968 to 1978

NOTE Confidence: 0.882581222

 $00:01:28.030 \longrightarrow 00:01:30.070$ as director of the Abraham Ribicoff

NOTE Confidence: 0.882581222

 $00:01:30.070 \longrightarrow 00:01:32.010$ Research facilities at the Connecticut

NOTE Confidence: 0.882581222

 $00{:}01{:}32.010 \dashrightarrow 00{:}01{:}34.170$ Mental Connecticut Mental Health Center,

NOTE Confidence: 0.882581222

 $00:01:34.170 \longrightarrow 00:01:36.922$ which is where most of our basic or

NOTE Confidence: 0.882581222

 $00:01:36.922 \longrightarrow 00:01:39.775$ a large chunk of our basic science

NOTE Confidence: 0.882581222

 $00:01:39.775 \longrightarrow 00:01:42.820$ labs remain right in in proximity

NOTE Confidence: 0.882581222

 $00{:}01{:}42.820 \dashrightarrow 00{:}01{:}46.245$ to our clinical research facilities,

NOTE Confidence: 0.882581222

 $00:01:46.250 \longrightarrow 00:01:48.980$ which was something that was essential

 $00:01:48.980 \longrightarrow 00:01:50.800$ for establishing the translational

NOTE Confidence: 0.882581222

 $00{:}01{:}50.869 \dashrightarrow 00{:}01{:}53.049$ and collaborative nature of the

NOTE Confidence: 0.882581222

 $00:01:53.049 \longrightarrow 00:01:54.417$ department and since 1982.

NOTE Confidence: 0.882581222

00:01:54.417 --> 00:01:56.580 We've had a lecture held in his

NOTE Confidence: 0.882581222

 $00:01:56.647 \longrightarrow 00:01:58.882$ honor recognizing his quote pivotal

NOTE Confidence: 0.882581222

 $00:01:58.882 \longrightarrow 00:02:01.117$ role in establishing the central

NOTE Confidence: 0.882581222

 $00:02:01.185 \longrightarrow 00:02:03.077$ importance of basic neuroscience

NOTE Confidence: 0.882581222

 $00:02:03.077 \longrightarrow 00:02:05.442$ research as their frontier for

NOTE Confidence: 0.882581222

 $00{:}02{:}05.442 \dashrightarrow 00{:}02{:}06.596$ clinical psychiatric studies.

NOTE Confidence: 0.882581222

 $00:02:06.596 \longrightarrow 00:02:08.624$ And that's a tradition that we

NOTE Confidence: 0.882581222

 $00{:}02{:}08.624 \dashrightarrow 00{:}02{:}10.269$ honor today with our speaker,

NOTE Confidence: 0.882581222

 $00:02:10.270 \longrightarrow 00:02:11.533$ doctor Patricia Janik.

NOTE Confidence: 0.882581222

 $00:02:11.533 \longrightarrow 00:02:15.030$ So here's now why we had Celtic music.

NOTE Confidence: 0.882581222

 $00{:}02{:}15.030 \dashrightarrow 00{:}02{:}17.432$ Doctor Flynn was born in Superior, WI.

NOTE Confidence: 0.882581222

00:02:17.432 --> 00:02:19.966 The of an Irish immigrant mother and

NOTE Confidence: 0.882581222

 $00{:}02{:}19.966 \dashrightarrow 00{:}02{:}22.281$ a first generation Irish American

 $00:02:22.281 \longrightarrow 00:02:25.311$ father who worked as a railroad

NOTE Confidence: 0.882581222

 $00:02:25.311 \longrightarrow 00:02:27.740$ switchman and he studied for the priesthood.

NOTE Confidence: 0.882581222

 $00{:}02{:}27.740 \dashrightarrow 00{:}02{:}30.156$ He was ordained in Rome and then he

NOTE Confidence: 0.882581222

 $00:02:30.156 \longrightarrow 00:02:32.510$ returned to the United States in 1938

NOTE Confidence: 0.882581222

 $00:02:32.510 \longrightarrow 00:02:34.910$ to Loyola University and when his

NOTE Confidence: 0.882581222

 $00:02:34.910 \longrightarrow 00:02:37.012$ superiors there decided that they

NOTE Confidence: 0.882581222

00:02:37.012 --> 00:02:39.087 needed someone to teach psychology,

NOTE Confidence: 0.882581222

 $00:02:39.090 \longrightarrow 00:02:41.260$ he volunteered to study it.

NOTE Confidence: 0.882581222

 $00:02:41.260 \longrightarrow 00:02:42.680$ And then he went out.

NOTE Confidence: 0.882581222

 $00:02:42.680 \longrightarrow 00:02:44.871$ To find the best teacher so he

NOTE Confidence: 0.882581222

 $00:02:44.871 \longrightarrow 00:02:46.670$ could actually teach his students.

NOTE Confidence: 0.882581222

 $00:02:46.670 \longrightarrow 00:02:48.734$ This led him to Columbia University

NOTE Confidence: 0.882581222

 $00{:}02{:}48.734 \dashrightarrow 00{:}02{:}50.569$ and there he studied psychology

NOTE Confidence: 0.882581222

 $00:02:50.569 \longrightarrow 00:02:52.939$ and he remained in the priesthood.

NOTE Confidence: 0.882581222

 $00:02:52.940 \longrightarrow 00:02:55.155$ But throughout this time he

 $00:02:55.155 \longrightarrow 00:02:56.927$ was examining his conscience.

NOTE Confidence: 0.882581222

 $00{:}02{:}56.930 \dashrightarrow 00{:}02{:}59.390$ And he ultimately resigned from the

NOTE Confidence: 0.882581222

 $00:02:59.390 \longrightarrow 00:03:02.236$ priesthood and left the church in 1944,

NOTE Confidence: 0.882581222

 $00:03:02.236 \longrightarrow 00:03:04.766$ having received his PhD in

NOTE Confidence: 0.882581222

00:03:04.766 --> 00:03:06.772 experimental psychology in 1943.

NOTE Confidence: 0.882581222

00:03:06.772 --> 00:03:11.110 In 1944, Dr Flynn went to work at Harvard,

NOTE Confidence: 0.882581222

 $00:03:11.110 \longrightarrow 00:03:13.366$ where he did war work related

NOTE Confidence: 0.882581222

00:03:13.366 --> 00:03:14.870 to aviation and audition,

NOTE Confidence: 0.882581222

 $00:03:14.870 \longrightarrow 00:03:17.732$ and in late 1945 he married

NOTE Confidence: 0.882581222

00:03:17.732 --> 00:03:19.640 a holder Isma Garvey,

NOTE Confidence: 0.882581222

 $00{:}03{:}19.640 \dashrightarrow 00{:}03{:}22.195$ and she was someone who I met

NOTE Confidence: 0.882581222

 $00:03:22.195 \longrightarrow 00:03:23.290$ when I first

NOTE Confidence: 0.872659905

 $00:03:23.379 \longrightarrow 00:03:24.767$ came to Yale,

NOTE Confidence: 0.872659905

 $00:03:24.770 \longrightarrow 00:03:26.996$ and she would come with Sarah.

NOTE Confidence: 0.872659905

 $00:03:27.000 \longrightarrow 00:03:29.916$ To listen to the lecture and she was also

NOTE Confidence: 0.872659905

 $00:03:29.916 \longrightarrow 00:03:33.039$ an incredible and remarkable individual.

 $00:03:33.040 \longrightarrow 00:03:36.143$ She was a psychologist and in 1946

NOTE Confidence: 0.872659905

 $00{:}03{:}36.143 \dashrightarrow 00{:}03{:}38.887$ and she was here actually as a member

NOTE Confidence: 0.872659905

00:03:38.887 --> 00:03:41.520 of the Department of Psychiatry,

NOTE Confidence: 0.872659905

 $00:03:41.520 \longrightarrow 00:03:44.642$ first appointed in 1962 as a research

NOTE Confidence: 0.872659905

 $00:03:44.642 \longrightarrow 00:03:47.200$ assistant and then serving on the

NOTE Confidence: 0.872659905

 $00{:}03{:}47.200 \dashrightarrow 00{:}03{:}49.310$ planning project for the Connecticut

NOTE Confidence: 0.872659905

00:03:49.386 --> 00:03:51.706 Mental Health Center and finally.

NOTE Confidence: 0.872659905

00:03:51.710 --> 00:03:54.830 Working closely with Doctor Boris Astrachan,

NOTE Confidence: 0.872659905

 $00:03:54.830 \longrightarrow 00:03:58.190$ who was instrumental in founding the

NOTE Confidence: 0.872659905

 $00:03:58.190 \longrightarrow 00:03:59.965$ the Connecticut Mental Health Center,

NOTE Confidence: 0.872659905

 $00:03:59.970 \longrightarrow 00:04:02.497$ she she retired as a valued member

NOTE Confidence: 0.872659905

 $00:04:02.497 \longrightarrow 00:04:04.759$ of the medical school faculty.

NOTE Confidence: 0.872659905

 $00:04:04.760 \longrightarrow 00:04:08.304$ So in 1946,

NOTE Confidence: 0.872659905

 $00{:}04{:}08.304 \dashrightarrow 00{:}04{:}09.924$ Doctor Flynn was appointed head

NOTE Confidence: 0.872659905

 $00:04:09.924 \longrightarrow 00:04:11.676$ of the psychology and Statistics

00:04:11.676 --> 00:04:13.656 Division at the Naval Medical

NOTE Confidence: 0.872659905

 $00{:}04{:}13.656 \dashrightarrow 00{:}04{:}15.240$ Research Institute in Bethesda,

NOTE Confidence: 0.872659905

 $00:04:15.240 \longrightarrow 00:04:17.988$ and there he began his work

NOTE Confidence: 0.872659905

00:04:17.988 --> 00:04:19.362 in physiological psychology.

NOTE Confidence: 0.872659905

 $00:04:19.370 \longrightarrow 00:04:22.065$ And here's where the story gets even

NOTE Confidence: 0.872659905

00:04:22.065 --> 00:04:23.763 more interesting despite excellent

NOTE Confidence: 0.872659905

 $00:04:23.763 \longrightarrow 00:04:25.659$ performance reviews and general

NOTE Confidence: 0.872659905

 $00{:}04{:}25.659 \dashrightarrow 00{:}04{:}28.029$ acclaim by his fellow scientists.

NOTE Confidence: 0.872659905

 $00:04:28.030 \longrightarrow 00:04:30.812$ Doctor Flynn was fired in 1953,

NOTE Confidence: 0.872659905

 $00:04:30.812 \longrightarrow 00:04:33.136$ and he was deemed a risk to

NOTE Confidence: 0.872659905

 $00{:}04{:}33.136 \dashrightarrow 00{:}04{:}34.780$ national security for his quote.

NOTE Confidence: 0.872659905

 $00:04:34.780 \longrightarrow 00:04:36.880$ Close and continuing association

NOTE Confidence: 0.872659905

00:04:36.880 --> 00:04:39.505 End Quote with his wife,

NOTE Confidence: 0.872659905

 $00:04:39.510 \longrightarrow 00:04:41.934$ whose name had been named before the house.

NOTE Confidence: 0.872659905

 $00:04:41.940 \longrightarrow 00:04:44.394$ A committee on UN American activities

NOTE Confidence: 0.872659905

 $00{:}04{:}44{.}394 \dashrightarrow 00{:}04{:}46{.}670$ during the McCarthy ERA era.

 $00{:}04{:}46.670 \dashrightarrow 00{:}04{:}50.510$ Because an error is a correct

NOTE Confidence: 0.872659905

 $00{:}04{:}50.510 \dashrightarrow 00{:}04{:}54.158$ Freudian slip because of of whole,

NOTE Confidence: 0.872659905

00:04:54.160 --> 00:04:57.040 this political activities in the 1930s

NOTE Confidence: 0.872659905

00:04:57.040 --> 00:04:59.836 and early 40s and doctor Flynn was

NOTE Confidence: 0.872659905

 $00:04:59.836 \longrightarrow 00:05:02.540$ offered the chance to keep his job if

NOTE Confidence: 0.872659905

 $00:05:02.540 \longrightarrow 00:05:05.970$ he divorced and he of course declined.

NOTE Confidence: 0.872659905

 $00:05:05.970 \longrightarrow 00:05:07.888$ Over the next six months or so,

NOTE Confidence: 0.872659905

 $00:05:07.890 \longrightarrow 00:05:10.158$ he received offers of employment from

NOTE Confidence: 0.872659905

 $00{:}05{:}10.158 \dashrightarrow 00{:}05{:}12.452$ colleagues across the country at 13

NOTE Confidence: 0.872659905

 $00:05:12.452 \longrightarrow 00:05:14.550$ universities, and he told his daughter,

NOTE Confidence: 0.872659905

00:05:14.550 --> 00:05:15.434 Sarah Flynn,

NOTE Confidence: 0.872659905

 $00{:}05{:}15.434 \dashrightarrow 00{:}05{:}18.086$ that each time his name reached

NOTE Confidence: 0.872659905

 $00:05:18.086 \longrightarrow 00:05:19.416$ the provost's office,

NOTE Confidence: 0.872659905

 $00{:}05{:}19.416 \dashrightarrow 00{:}05{:}21.631$ the colleague was informed that

NOTE Confidence: 0.872659905

00:05:21.631 --> 00:05:23.789 the university could not hire Dr.

00:05:23.790 --> 00:05:27.210 Flynn, and in September 1954,

NOTE Confidence: 0.872659905

 $00:05:27.210 \longrightarrow 00:05:29.526$ Yale hired him to work with

NOTE Confidence: 0.872659905

00:05:29.526 --> 00:05:30.684 Doctor Paul McLean,

NOTE Confidence: 0.872659905

 $00:05:30.690 \longrightarrow 00:05:32.718$ who then held a joint appointment

NOTE Confidence: 0.872659905

 $00:05:32.718 \longrightarrow 00:05:34.434$ in Physiology and Psychiatry and

NOTE Confidence: 0.872659905

 $00:05:34.434 \longrightarrow 00:05:36.258$ who was studying the limbic system.

NOTE Confidence: 0.872659905

 $00{:}05{:}36.260 \dashrightarrow 00{:}05{:}40.201$ So yell was able to benefit from

NOTE Confidence: 0.872659905

 $00{:}05{:}40.201 \dashrightarrow 00{:}05{:}42.612$ his neuroscience area addition

NOTE Confidence: 0.872659905

 $00{:}05{:}42.612 \dashrightarrow 00{:}05{:}46.278$ in the face of strong headwinds.

NOTE Confidence: 0.872659905

 $00:05:46.280 \longrightarrow 00:05:51.800$ Come upon a learning of so.

NOTE Confidence: 0.872659905

 $00:05:51.800 \longrightarrow 00:05:53.684$ Doctor Flynn then became a member

NOTE Confidence: 0.872659905

 $00:05:53.684 \longrightarrow 00:05:55.922$ of this of the department and

NOTE Confidence: 0.872659905

 $00{:}05{:}55.922 \dashrightarrow 00{:}06{:}00.202$ worked until he retired in 79 on the

NOTE Confidence: 0.872659905

 $00{:}06{:}00.202 \dashrightarrow 00{:}06{:}02.500$ physiological basis of aggression

NOTE Confidence: 0.872659905

 $00:06:02.500 \longrightarrow 00:06:06.868$ and he really made a an incredible

NOTE Confidence: 0.872659905

 $00:06:06.868 \longrightarrow 00:06:09.420$ mark on the department,

 $00:06:09.420 \longrightarrow 00:06:11.028$ and upon learning of his death,

NOTE Confidence: 0.872659905

00:06:11.030 --> 00:06:11.560 Fritz Redlich,

NOTE Confidence: 0.872659905

00:06:11.560 --> 00:06:13.415 whose chair of the Department of Psychiatry,

NOTE Confidence: 0.872659905

 $00:06:13.420 \longrightarrow 00:06:15.400$ wrote to holder Flynn at John

NOTE Confidence: 0.872659905

 $00:06:15.400 \longrightarrow 00:06:17.778$ was all I ever wanted to be.

NOTE Confidence: 0.872659905

00:06:17.780 --> 00:06:19.690 A fine scientist and teacher,

NOTE Confidence: 0.872659905

 $00:06:19.690 \longrightarrow 00:06:21.062$ and most of all,

NOTE Confidence: 0.872659905

00:06:21.062 --> 00:06:22.434 an extraordinary human being.

NOTE Confidence: 0.872659905

00:06:22.440 --> 00:06:24.080 I've always admired his courage

NOTE Confidence: 0.872659905

00:06:24.080 --> 00:06:25.844 and integrity, two virtues,

NOTE Confidence: 0.872659905

 $00:06:25.844 \longrightarrow 00:06:28.604$ high value above anything else,

NOTE Confidence: 0.872659905

 $00{:}06{:}28.610 \dashrightarrow 00{:}06{:}29.926$ and similar sentiments were

NOTE Confidence: 0.872659905

 $00{:}06{:}29.926 \dashrightarrow 00{:}06{:}31.571$ expressed by other colleagues both

NOTE Confidence: 0.872659905

 $00:06:31.571 \longrightarrow 00:06:33.280$ at Yale and around the world.

NOTE Confidence: 0.872659905

 $00:06:33.280 \longrightarrow 00:06:36.367$ And the last thing I want to say before

00:06:36.367 --> 00:06:39.716 I I move on is also about Sarah Flint.

NOTE Confidence: 0.872659905

 $00{:}06{:}39.716 \dashrightarrow 00{:}06{:}42.206$ Generate generosity to the department.

NOTE Confidence: 0.872659905

00:06:42.210 --> 00:06:44.818 So in April 2005,

NOTE Confidence: 0.872659905

 $00:06:44.818 \longrightarrow 00:06:46.774$ Sarah donated Dr.

NOTE Confidence: 0.872659905

00:06:46.780 --> 00:06:48.464 Flynn's most prized possession,

NOTE Confidence: 0.872659905

 $00:06:48.464 \longrightarrow 00:06:50.990$ the three volume set of romantica

NOTE Confidence: 0.872659905

00:06:51.054 --> 00:06:52.218 halls fixed Judah,

NOTE Confidence: 0.872659905

 $00:06:52.220 \longrightarrow 00:06:54.278$ their system and nervioso de Lumbre.

NOTE Confidence: 0.872659905

 $00{:}06{:}54.280 --> 00{:}06{:}55.915$ Elizabeth brought us.

NOTE Confidence: 0.872659905

 $00:06:55.915 \longrightarrow 00:06:58.892$ In the original Spanish to the

NOTE Confidence: 0.872659905

 $00:06:58.892 \longrightarrow 00:07:00.476$ Yale Medical Historical Library

NOTE Confidence: 0.872659905

 $00:07:00.476 \longrightarrow 00:07:02.060$ and inside the first

NOTE Confidence: 0.888918560555556

 $00{:}07{:}02.133 \dashrightarrow 00{:}07{:}03.978$ volume is an inscription written

NOTE Confidence: 0.888918560555556

 $00:07:03.978 \longrightarrow 00:07:07.794$ by Cahal in 1910, which reads in

NOTE Confidence: 0.888918560555556

 $00:07:07.794 \longrightarrow 00:07:09.874$ translation because of the brain.

NOTE Confidence: 0.888918560555556

 $00:07:09.874 \longrightarrow 00:07:11.980$ Man is the king of Creation,

 $00:07:11.980 \longrightarrow 00:07:13.842$ and to clarify the structure of the

NOTE Confidence: 0.888918560555556

 $00{:}07{:}13.842 \dashrightarrow 00{:}07{:}15.782$ brain is to understand why that figure

NOTE Confidence: 0.888918560555556

 $00:07:15.782 \longrightarrow 00:07:18.248$ is at the head of the animal Kingdom

NOTE Confidence: 0.888918560555556

 $00:07:18.248 \longrightarrow 00:07:20.188$ and how civilization was created.

NOTE Confidence: 0.888918560555556

00:07:20.190 --> 00:07:21.575 A sign of human superiority

NOTE Confidence: 0.888918560555556

 $00:07:21.575 \longrightarrow 00:07:23.450$ to the rest of the beings.

NOTE Confidence: 0.888918560555556

 $00:07:23.450 \longrightarrow 00:07:25.040$ This may not actually translate

NOTE Confidence: 0.888918560555556

 $00:07:25.040 \longrightarrow 00:07:26.630$ so well to the current.

NOTE Confidence: 0.888918560555556

 $00:07:26.630 \dashrightarrow 00:07:28.686$ Sarah, my original idea was that I would

NOTE Confidence: 0.88891856055556

 $00{:}07{:}28.686 \dashrightarrow 00{:}07{:}30.782$ not be able to take a sure step in

NOTE Confidence: 0.88891856055556

00:07:30.782 --> 00:07:32.582 the study of Physiology and pathology

NOTE Confidence: 0.888918560555556

 $00:07:32.582 \longrightarrow 00:07:34.748$ of the nervous system without knowing

NOTE Confidence: 0.888918560555556

 $00{:}07{:}34.748 \dashrightarrow 00{:}07{:}36.666$ the cerebral machine with precision,

NOTE Confidence: 0.888918560555556

 $00:07:36.666 \longrightarrow 00:07:39.399$ and that the mysteries of the science

NOTE Confidence: 0.888918560555556

 $00:07:39.399 \longrightarrow 00:07:41.723$ of the spirit will only be clarified

00:07:41.723 --> 00:07:43.991 when all the unknowns relative to

NOTE Confidence: 0.888918560555556

 $00:07:43.991 \longrightarrow 00:07:46.289$ the chemistry of the fine structures

NOTE Confidence: 0.888918560555556

 $00:07:46.290 \longrightarrow 00:07:48.565$ of the nerve cell are cleared up.

NOTE Confidence: 0.888918560555556

00:07:48.570 --> 00:07:49.690 And luckily, of course,

NOTE Confidence: 0.888918560555556

 $00:07:49.690 \longrightarrow 00:07:51.090$ we're completely done with that,

NOTE Confidence: 0.88891856055556

00:07:51.090 --> 00:07:54.261 and Doctor Janik will will give the

NOTE Confidence: 0.888918560555556

 $00:07:54.261 \longrightarrow 00:07:57.669$ the the heading to that to that quote.

NOTE Confidence: 0.888918560555556

00:07:57.670 --> 00:07:59.850 And just to finish up,

NOTE Confidence: 0.888918560555556

00:07:59.850 --> 00:08:01.998 Doctor Flynn always took delight in

NOTE Confidence: 0.888918560555556

 $00:08:01.998 \longrightarrow 00:08:04.128$ telling the story that he acquired

NOTE Confidence: 0.888918560555556

 $00:08:04.128 \longrightarrow 00:08:06.648$ these books for \$10 at a used bookstore

NOTE Confidence: 0.888918560555556

 $00{:}08{:}06.713 \dashrightarrow 00{:}08{:}11.070$ in New York sometime during the 1940s.

NOTE Confidence: 0.88891856055556

 $00{:}08{:}11.070 \dashrightarrow 00{:}08{:}16.006$ So I I will finish there and I I hope

NOTE Confidence: 0.888918560555556

 $00:08:16.006 \longrightarrow 00:08:20.340$ that you will now join me in welcoming.

NOTE Confidence: 0.888918560555556

 $00:08:20.340 \longrightarrow 00:08:21.570$ Sorry I will get through.

NOTE Confidence: 0.888918560555556

 $00:08:21.570 \longrightarrow 00:08:26.338$ These are 2021 lecturer in the

00:08:26.338 --> 00:08:27.946 Flynn Memorial Lecture series,

NOTE Confidence: 0.888918560555556

 $00{:}08{:}27.950 \dashrightarrow 00{:}08{:}32.766$ Doctor Patricia Janik and so Dr
 Janik

NOTE Confidence: 0.888918560555556

 $00{:}08{:}32.770 \dashrightarrow 00{:}08{:}34.820$ is the A Bloomberg distinguished

NOTE Confidence: 0.888918560555556

00:08:34.820 --> 00:08:36.870 professor at Johns Hopkins University

NOTE Confidence: 0.888918560555556

 $00:08:36.936 \longrightarrow 00:08:39.046$ with appointments in the Department

NOTE Confidence: 0.888918560555556

 $00:08:39.046 \longrightarrow 00:08:41.156$ of Psychological and Brain Sciences.

NOTE Confidence: 0.888918560555556

00:08:41.160 --> 00:08:43.170 And the Krieger School of Arts

NOTE Confidence: 0.888918560555556

00:08:43.170 --> 00:08:44.989 and Sciences and the Department

NOTE Confidence: 0.888918560555556

 $00{:}08{:}44.989 \dashrightarrow 00{:}08{:}47.467$ of Neuroscience in the School of

NOTE Confidence: 0.88891856055556

 $00{:}08{:}47.467 \dashrightarrow 00{:}08{:}49.700$ Medicine and Doctor Janik Studies.

NOTE Confidence: 0.888918560555556

00:08:49.700 --> 00:08:51.680 Neural processes of reward learning.

NOTE Confidence: 0.888918560555556

 $00:08:51.680 \longrightarrow 00:08:53.744$ And you'll hear a lot about that today.

NOTE Confidence: 0.888918560555556

 $00:08:53.750 \longrightarrow 00:08:55.830$ She's especially interested in learning

NOTE Confidence: 0.888918560555556

 $00:08:55.830 \longrightarrow 00:08:57.078$ mechanisms underlying addiction,

NOTE Confidence: 0.888918560555556

 $00:08:57.080 \longrightarrow 00:09:00.754$ which is an area where this department

 $00:09:00.754 \longrightarrow 00:09:03.664$ certainly has extremely strong interest.

NOTE Confidence: 0.888918560555556

 $00{:}09{:}03.670 \dashrightarrow 00{:}09{:}04.670$ She earned her pH.

NOTE Confidence: 0.888918560555556

00:09:04.670 --> 00:09:04.920 D.

NOTE Confidence: 0.888918560555556

00:09:04.920 --> 00:09:06.190 From the University of California,

NOTE Confidence: 0.888918560555556 00:09:06.190 --> 00:09:06.550 Berkeley, NOTE Confidence: 0.888918560555556

110 1 E Connachee. 0.00031030033330

 $00:09:06.550 \longrightarrow 00:09:08.710$ and then she conducted postdoctoral research

NOTE Confidence: 0.888918560555556

 $00:09:08.710 \longrightarrow 00:09:11.416$ at Wake Forest and at the National Institute.

NOTE Confidence: 0.888918560555556

 $00:09:11.420 \longrightarrow 00:09:15.462$ And drug abuse and come in from 1999 to 2014,

NOTE Confidence: 0.88891856055556

 $00{:}09{:}15.462 \dashrightarrow 00{:}09{:}17.520$ which is the period when I first

NOTE Confidence: 0.888918560555556

 $00:09:17.591 \longrightarrow 00:09:18.679$ came to know her.

NOTE Confidence: 0.888918560555556

 $00:09:18.680 \longrightarrow 00:09:20.510$ She was faculty at the University

NOTE Confidence: 0.888918560555556

00:09:20.510 --> 00:09:22.360 of California at San Francisco,

NOTE Confidence: 0.888918560555556

00:09:22.360 --> 00:09:24.719 where she was the Howard J Weinberger,

NOTE Confidence: 0.888918560555556

00:09:24.720 --> 00:09:27.395 MD endowed Chair and addiction

NOTE Confidence: 0.888918560555556

 $00:09:27.395 \longrightarrow 00:09:28.806$ research at UCSF.

NOTE Confidence: 0.888918560555556

 $00:09:28.806 \longrightarrow 00:09:30.996$ She's a pioneer in the

00:09:30.996 --> 00:09:33.100 identification of neural circuits,

NOTE Confidence: 0.888918560555556

 $00{:}09{:}33.100 \dashrightarrow 00{:}09{:}35.180$ underlying alcohol and drug seeking,

NOTE Confidence: 0.888918560555556

 $00:09:35.180 \longrightarrow 00:09:37.682$ and her work is really spanned

NOTE Confidence: 0.888918560555556

00:09:37.682 --> 00:09:39.967 levels of investigation from the

NOTE Confidence: 0.888918560555556

 $00:09:39.967 \longrightarrow 00:09:42.119$ molecular and synaptic plasticity.

NOTE Confidence: 0.888918560555556

 $00:09:42.120 \longrightarrow 00:09:43.807$ All the way to in vivo mechanisms

NOTE Confidence: 0.888918560555556

00:09:43.807 --> 00:09:45.456 that are relevant to complex models

NOTE Confidence: 0.888918560555556

00:09:45.456 --> 00:09:46.906 that are relevant to addiction,

NOTE Confidence: 0.888918560555556

 $00:09:46.910 \longrightarrow 00:09:49.400$ and this includes alcohol and

NOTE Confidence: 0.88891856055556

 $00:09:49.400 \longrightarrow 00:09:50.894$ drug seeking relapse,

NOTE Confidence: 0.888918560555556

00:09:50.900 --> 00:09:51.848 habit learning,

NOTE Confidence: 0.888918560555556

00:09:51.848 --> 00:09:52.796 extinction learning,

NOTE Confidence: 0.88891856055556

 $00{:}09{:}52.796 \dashrightarrow 00{:}09{:}54.218$ and she's used.

NOTE Confidence: 0.888918560555556

 $00:09:54.220 \longrightarrow 00:09:55.759$ Everything from electrophysiological

NOTE Confidence: 0.888918560555556

 $00:09:55.759 \longrightarrow 00:09:57.298$ approaches to neuronal

 $00:09:57.298 \longrightarrow 00:09:58.837$ imaging and optogenetics,

NOTE Confidence: 0.888918560555556

 $00{:}09{:}58.840 \dashrightarrow 00{:}10{:}00.716$ which I'm sure you'll hear about today,

NOTE Confidence: 0.888918560555556

 $00:10:00.720 \longrightarrow 00:10:02.388$ and if you're looking for a

NOTE Confidence: 0.888918560555556

 $00:10:02.388 \longrightarrow 00:10:03.953$ resource to understand the neural

NOTE Confidence: 0.888918560555556

00:10:03.953 --> 00:10:05.557 circuitry relevant to addiction,

NOTE Confidence: 0.888918560555556

 $00:10:05.560 \longrightarrow 00:10:07.898$ you need to read her 2021 review

NOTE Confidence: 0.888918560555556

00:10:07.898 --> 00:10:09.414 on consolidating the circuit

NOTE Confidence: 0.888918560555556

 $00:10:09.414 \longrightarrow 00:10:11.384$ model for addiction that she

NOTE Confidence: 0.888918560555556

 $00:10:11.384 \longrightarrow 00:10:12.968$ wrote with Christian luescher.

NOTE Confidence: 0.888918560555556

 $00:10:12.970 \longrightarrow 00:10:15.388$ And that appears in the annual

NOTE Confidence: 0.888918560555556

00:10:15.388 --> 00:10:16.597 review of neuroscience.

NOTE Confidence: 0.879787118571429

 $00:10:16.600 \longrightarrow 00:10:18.472$ And I first met Doctor Genich through her

NOTE Confidence: 0.879787118571429

 $00{:}10{:}18.472 \dashrightarrow 00{:}10{:}20.347$ roles at the Society for Neuroscience.

NOTE Confidence: 0.879787118571429

 $00:10:20.350 \longrightarrow 00:10:22.156$ She's she's done a lot for the

NOTE Confidence: 0.879787118571429

 $00:10:22.156 \longrightarrow 00:10:23.997$ society she served as reviewing editor

NOTE Confidence: 0.879787118571429

00:10:23.997 --> 00:10:25.677 at the Journal of Neuroscience.

00:10:25.680 --> 00:10:27.654 She's been chair of the program committee,

NOTE Confidence: 0.879787118571429

00:10:27.660 --> 00:10:30.242 probably for way too long, given kovid,

NOTE Confidence: 0.879787118571429

 $00:10:30.242 \longrightarrow 00:10:34.099$ and she's incoming secretary of the society.

NOTE Confidence: 0.879787118571429

 $00:10:34.100 \longrightarrow 00:10:36.753$ She's also served on the Program Committee

NOTE Confidence: 0.879787118571429

00:10:36.753 --> 00:10:39.439 for the Research Society and Alcoholism,

NOTE Confidence: 0.879787118571429

 $00:10:39.440 \longrightarrow 00:10:41.729$ and She's been Co Chair and Chair

NOTE Confidence: 0.879787118571429

 $00:10:41.729 \longrightarrow 00:10:43.609$ of the Catecholamines and the

NOTE Confidence: 0.879787118571429

00:10:43.609 --> 00:10:44.860 Alcohol Gordon Conferences,

NOTE Confidence: 0.879787118571429

 $00:10:44.860 \longrightarrow 00:10:46.516$ so you can see that the

NOTE Confidence: 0.879787118571429

00:10:46.516 --> 00:10:47.620 influence of her work.

NOTE Confidence: 0.879787118571429

 $00:10:47.620 \longrightarrow 00:10:50.875$ In the field is extremely broad and

NOTE Confidence: 0.879787118571429

00:10:50.875 --> 00:10:52.915 I just want to close by saying that

NOTE Confidence: 0.879787118571429

 $00:10:52.915 \dashrightarrow 00:10:54.957$ Doctor Janik is much more than her CV.

NOTE Confidence: 0.879787118571429

 $00:10:54.960 \longrightarrow 00:10:57.088$ She's been a mentor to leaders in the

NOTE Confidence: 0.879787118571429

00:10:57.088 --> 00:10:59.129 field who study the neurobiology of

00:10:59.129 --> 00:11:00.939 addiction and other behaviors that

NOTE Confidence: 0.879787118571429

 $00{:}11{:}00.939 \dashrightarrow 00{:}11{:}03.178$ are relevant to psychiatric illness.

NOTE Confidence: 0.879787118571429

 $00:11:03.180 \longrightarrow 00:11:05.596$ When my own lab was trying to figure

NOTE Confidence: 0.879787118571429

 $00:11:05.596 \longrightarrow 00:11:07.529$ out issues related to experimental

NOTE Confidence: 0.879787118571429

00:11:07.529 --> 00:11:10.055 design for in vivo calcium imaging,

NOTE Confidence: 0.879787118571429

 $00{:}11{:}10.060 \dashrightarrow 00{:}11{:}12.006$ Katie told me that the most important

NOTE Confidence: 0.879787118571429

 $00:11:12.006 \longrightarrow 00:11:14.204$ thing she ever learned in her scientific

NOTE Confidence: 0.879787118571429

00:11:14.204 --> 00:11:15.839 career was from Doctor Janik.

NOTE Confidence: 0.879787118571429

 $00:11:15.840 \longrightarrow 00:11:17.660$ And that is how to start with.

NOTE Confidence: 0.879787118571429

 $00:11:17.660 \longrightarrow 00:11:19.410$ Robust experimental design that gives

NOTE Confidence: 0.879787118571429

 $00{:}11{:}19.410 \dashrightarrow 00{:}11{:}21.590$ you the adequate power to that's

NOTE Confidence: 0.879787118571429

00:11:21.590 --> 00:11:23.828 essential to get robust Physiology data,

NOTE Confidence: 0.879787118571429

 $00:11:23.830 \longrightarrow 00:11:25.240$ and we've certainly taken this

NOTE Confidence: 0.879787118571429

 $00:11:25.240 \longrightarrow 00:11:26.368$ to heart as well.

NOTE Confidence: 0.879787118571429

 $00:11:26.370 \longrightarrow 00:11:28.380$ So as you can tell,

NOTE Confidence: 0.879787118571429

 $00:11:28.380 \longrightarrow 00:11:31.004$ Dr Janik is a role model for many

 $00:11:31.004 \longrightarrow 00:11:34.346$ and she is the ideal person to give

NOTE Confidence: 0.879787118571429

 $00{:}11{:}34.346 \to 00{:}11{:}36.960$ the Flynn Memorial lecture this year.

NOTE Confidence: 0.879787118571429

 $00:11:36.960 \longrightarrow 00:11:39.040$ So please let's welcome Dr.

NOTE Confidence: 0.879787118571429

00:11:39.040 --> 00:11:41.673 Janik and I will stop sharing and

NOTE Confidence: 0.879787118571429

00:11:41.673 --> 00:11:44.330 allow her to share her data with you.

NOTE Confidence: 0.970280178333333

 $00:11:46.830 \longrightarrow 00:11:48.666$ Thank you so much for that.

NOTE Confidence: 0.970280178333333

 $00:11:48.670 \longrightarrow 00:11:52.030$ It was such an an amazing introduction

NOTE Confidence: 0.970280178333333

 $00{:}11{:}52.030 \dashrightarrow 00{:}11{:}55.738$ and I really enjoyed learning about.

NOTE Confidence: 0.970280178333333

00:11:55.740 --> 00:11:57.770 Now, Doctor Flynn and can

NOTE Confidence: 0.970280178333333

 $00:11:57.770 \longrightarrow 00:12:00.080$ you see my screen? Yes.

NOTE Confidence: 0.902419785

 $00:12:01.550 \longrightarrow 00:12:02.298$ I have to say

NOTE Confidence: 0.87350712375

 $00:12:02.310 \longrightarrow 00:12:04.340$ what it, what a deep honor it is to be

NOTE Confidence: 0.87350712375

 $00:12:04.403 \longrightarrow 00:12:06.545$ invited to give this particular lecture.

NOTE Confidence: 0.87350712375

00:12:06.550 --> 00:12:08.734 I it was such an interesting history

NOTE Confidence: 0.87350712375

 $00:12:08.734 \longrightarrow 00:12:11.169$ and I really hope that you find that

00:12:11.169 --> 00:12:13.533 the kind of work that I talk about

NOTE Confidence: 0.87350712375

 $00{:}12{:}13.533 \dashrightarrow 00{:}12{:}15.110$ today resonates with the kinds of

NOTE Confidence: 0.87350712375

 $00:12:15.110 \longrightarrow 00:12:16.590$ things that he was interested in,

NOTE Confidence: 0.87350712375

 $00:12:16.590 \longrightarrow 00:12:19.330$ so that you can see that it's a good fit.

NOTE Confidence: 0.87350712375

 $00{:}12{:}19.330 \dashrightarrow 00{:}12{:}21.358$ And I I want to thank you, Marina for

NOTE Confidence: 0.87350712375

 $00{:}12{:}21.358 \dashrightarrow 00{:}12{:}23.228$ the invitation and this opportunity.

NOTE Confidence: 0.87350712375

 $00:12:23.230 \longrightarrow 00:12:26.534$ And thank you for such an A.

NOTE Confidence: 0.87350712375

00:12:26.540 --> 00:12:28.880 Humbling introduction that that was really,

NOTE Confidence: 0.87350712375

 $00:12:28.880 \longrightarrow 00:12:30.973$ really so nice and I'll try my

NOTE Confidence: 0.87350712375

 $00:12:30.973 \longrightarrow 00:12:33.301$ best to live up to everything that

NOTE Confidence: 0.87350712375

 $00:12:33.301 \longrightarrow 00:12:35.341$ has been said in this talk.

NOTE Confidence: 0.87350712375

00:12:35.350 --> 00:12:38.497 So, uh, welcome to everybody and I'm

NOTE Confidence: 0.87350712375

00:12:38.497 --> 00:12:41.193 sorry I'm not meeting you today in person,

NOTE Confidence: 0.87350712375

00:12:41.200 --> 00:12:43.153 but I'm so happy to talk to you even

NOTE Confidence: 0.87350712375

00:12:43.153 --> 00:12:44.963 though we're over zoom and if any

NOTE Confidence: 0.87350712375

 $00{:}12{:}44.963 \dashrightarrow 00{:}12{:}46.500$ questions come up during the talk.

 $00:12:46.500 \longrightarrow 00:12:49.218$ I'm sure people will help me to try to

NOTE Confidence: 0.87350712375

 $00:12:49.218 \longrightarrow 00:12:51.408$ answer those since I don't think I'll

NOTE Confidence: 0.87350712375

00:12:51.408 --> 00:12:53.967 be able to see the chat very well.

NOTE Confidence: 0.87350712375

00:12:53.970 --> 00:12:56.994 OK, so I'd like to tell you about

NOTE Confidence: 0.87350712375

00:12:56.994 --> 00:12:58.290 our experiments today.

NOTE Confidence: 0.87350712375

00:12:58.290 --> 00:13:00.162 Looking at reward processing

NOTE Confidence: 0.87350712375

 $00:13:00.162 \longrightarrow 00:13:02.034$ in the nervous system,

NOTE Confidence: 0.87350712375

 $00:13:02.040 \longrightarrow 00:13:04.108$ specifically focusing on the

NOTE Confidence: 0.87350712375

00:13:04.108 --> 00:13:06.693 area called the ventral pallidum,

NOTE Confidence: 0.87350712375

 $00{:}13{:}06.700 \dashrightarrow 00{:}13{:}08.476$ and I want to first tell you a

NOTE Confidence: 0.87350712375

 $00:13:08.476 \longrightarrow 00:13:10.069$ little bit about the motivation

NOTE Confidence: 0.87350712375

 $00:13:10.070 \longrightarrow 00:13:13.014$ for us in our lab in looking at

NOTE Confidence: 0.87350712375

 $00{:}13{:}13.014 \dashrightarrow 00{:}13{:}15.584$ reward seeking behavior models and

NOTE Confidence: 0.87350712375

 $00:13:15.584 \longrightarrow 00:13:18.270$ the underlying neural circuitry and

NOTE Confidence: 0.87350712375

 $00:13:18.270 \longrightarrow 00:13:20.370$ what we are interested in broadly,

 $00:13:20.370 \longrightarrow 00:13:22.530$ is what the processes are.

NOTE Confidence: 0.87350712375

 $00:13:22.530 \longrightarrow 00:13:25.020$ The determined reward seeking behavior.

NOTE Confidence: 0.87350712375

 $00:13:25.020 \longrightarrow 00:13:27.337$ Whether that reward is a food reward,

NOTE Confidence: 0.87350712375

 $00:13:27.340 \longrightarrow 00:13:29.330$ something that our nervous system

NOTE Confidence: 0.87350712375

00:13:29.330 --> 00:13:32.109 evolved to help us discover and ingest,

NOTE Confidence: 0.87350712375

 $00:13:32.110 \longrightarrow 00:13:33.598$ or whether it's a drug reward,

NOTE Confidence: 0.87350712375

00:13:33.600 --> 00:13:35.220 something that it's very important

NOTE Confidence: 0.87350712375

 $00:13:35.220 \longrightarrow 00:13:37.325$ for us to understand as we think

NOTE Confidence: 0.87350712375

 $00:13:37.325 \longrightarrow 00:13:39.349$ about how we can help individuals

NOTE Confidence: 0.87350712375

 $00:13:39.349 \longrightarrow 00:13:41.377$ with substance use disorders,

NOTE Confidence: 0.87350712375

 $00{:}13{:}41.380 \dashrightarrow 00{:}13{:}43.480$ and we conceive of these processes

NOTE Confidence: 0.87350712375

 $00:13:43.480 \longrightarrow 00:13:45.660$ through the lens of psychology.

NOTE Confidence: 0.87350712375

 $00:13:45.660 \longrightarrow 00:13:48.576$ And there really are three interrelated

NOTE Confidence: 0.87350712375

 $00:13:48.576 \longrightarrow 00:13:50.034$ psychological processes that

NOTE Confidence: 0.87350712375

00:13:50.034 --> 00:13:52.018 determine at any one moment whether

NOTE Confidence: 0.87350712375

00:13:52.018 --> 00:13:54.329 an agent will seek a given reward.

 $00:13:54.330 \longrightarrow 00:13:55.380$ And so first you have the.

NOTE Confidence: 0.87350712375

 $00:13:55.380 \longrightarrow 00:13:56.496$ Real time decision.

NOTE Confidence: 0.87350712375

00:13:56.496 --> 00:13:58.728 Will you decide to reach out

NOTE Confidence: 0.87350712375

00:13:58.728 --> 00:14:01.150 your arm and grab that hamburger?

NOTE Confidence: 0.87350712375

 $00:14:01.150 \longrightarrow 00:14:03.110$ Will the attic decide to call up

NOTE Confidence: 0.87350712375

 $00:14:03.110 \longrightarrow 00:14:05.255$ the dealer to try to get that next

NOTE Confidence: 0.87350712375

 $00:14:05.255 \longrightarrow 00:14:07.466$ fix so you have your real time

NOTE Confidence: 0.87350712375

 $00:14:07.466 \longrightarrow 00:14:09.266$ decision that's impacted critically

NOTE Confidence: 0.87350712375

 $00{:}14{:}09.266 \dashrightarrow 00{:}14{:}11.515$ by your current motivational state?

NOTE Confidence: 0.87350712375

00:14:11.515 --> 00:14:12.850 Whether you're hungry,

NOTE Confidence: 0.87350712375

 $00:14:12.850 \longrightarrow 00:14:14.185$ whether you're thirsty,

NOTE Confidence: 0.87350712375

 $00:14:14.190 \longrightarrow 00:14:16.010$ whether you are a person with an

NOTE Confidence: 0.87350712375

 $00{:}14{:}16.010 \dashrightarrow 00{:}14{:}17.949$ abuse disorder, whose craving drug,

NOTE Confidence: 0.87350712375

 $00:14:17.949 \longrightarrow 00:14:20.787$ your decision is is necessarily filtered

NOTE Confidence: 0.87350712375

 $00:14:20.787 \longrightarrow 00:14:22.998$ through your motivational state.

 $00:14:23.000 \longrightarrow 00:14:25.562$ And both of these critically depend

NOTE Confidence: 0.87350712375

 $00{:}14{:}25.562 \dashrightarrow 00{:}14{:}28.030$ on past experience or learning.

NOTE Confidence: 0.87350712375

 $00:14:28.030 \longrightarrow 00:14:30.508$ So your past evaluation of the

NOTE Confidence: 0.87350712375

 $00:14:30.508 \longrightarrow 00:14:32.812$ subjective effects of the rewards

NOTE Confidence: 0.87350712375

00:14:32.812 --> 00:14:35.290 that you've experienced and you're

NOTE Confidence: 0.87350712375

 $00{:}14{:}35.290 \dashrightarrow 00{:}14{:}37.190$ learning about the conditions under

NOTE Confidence: 0.87350712375

 $00:14:37.190 \longrightarrow 00:14:39.365$ which you obtain those rewards is

NOTE Confidence: 0.87350712375

 $00:14:39.365 \longrightarrow 00:14:41.297$ critical for you in the future.

NOTE Confidence: 0.87350712375

00:14:41.300 --> 00:14:42.930 When you're making that decision

NOTE Confidence: 0.87350712375

 $00:14:42.930 \longrightarrow 00:14:45.448$ to get that reward so you know the

NOTE Confidence: 0.87350712375

 $00{:}14{:}45.448 \dashrightarrow 00{:}14{:}47.098$ actions to take or not take,

NOTE Confidence: 0.87350712375

00:14:47.100 --> 00:14:48.966 and you understand the meaning of

NOTE Confidence: 0.87350712375

 $00:14:48.966 \longrightarrow 00:14:50.990$ the stimuli in the environment.

NOTE Confidence: 0.87350712375

 $00{:}14{:}50.990 \dashrightarrow 00{:}14{:}53.118$ So we have these three interacting processes.

NOTE Confidence: 0.87350712375

 $00:14:53.120 \longrightarrow 00:14:54.944$ And we're interested in the neural

NOTE Confidence: 0.87350712375

00:14:54.944 --> 00:14:56.160 circuits that underlie them,

 $00{:}14{:}56.160 \dashrightarrow 00{:}14{:}58.800$ so we can understand decision

NOTE Confidence: 0.87350712375

 $00:14:58.800 \longrightarrow 00:15:00.912$ making both in normal

NOTE Confidence: 0.92424688466667

 $00:15:00.920 \longrightarrow 00:15:03.026$ conditions like feeding and also our

NOTE Confidence: 0.924246884666667

 $00:15:03.026 \longrightarrow 00:15:05.808$ end goal is to better understand this

NOTE Confidence: 0.924246884666667

 $00:15:05.808 \longrightarrow 00:15:08.846$ circuit so we can help explain decisions

NOTE Confidence: 0.924246884666667

00:15:08.920 --> 00:15:11.720 made by people who have substance use

NOTE Confidence: 0.924246884666667

 $00:15:11.720 \longrightarrow 00:15:13.848$ disorders or alcohol use disorders,

NOTE Confidence: 0.924246884666667

 $00{:}15{:}13.848 \dashrightarrow 00{:}15{:}16.168$ because these same processes of

NOTE Confidence: 0.924246884666667

 $00{:}15{:}16.168 {\:\raisebox{--}{\text{--}}}{\:\raisebox{--}{\text{--}}}{\:\raisebox{--}{\text{--}}} 00{:}15{:}18.789$ course are occurring when one makes

NOTE Confidence: 0.924246884666667

 $00:15:18.789 \longrightarrow 00:15:21.099$ the decision to continue taking it.

NOTE Confidence: 0.924246884666667

00:15:21.100 --> 00:15:23.347 Drink for example, or to take another.

NOTE Confidence: 0.924246884666667

 $00:15:23.350 \longrightarrow 00:15:25.810$ Hit of that drug.

NOTE Confidence: 0.924246884666667

00:15:25.810 --> 00:15:29.149 So through many, many decades of work

NOTE Confidence: 0.924246884666667

 $00:15:29.149 \longrightarrow 00:15:32.740$ in a nonhuman animals and in humans.

NOTE Confidence: 0.924246884666667

 $00:15:32.740 \longrightarrow 00:15:35.570$ We've discovered as a field.

 $00:15:35.570 \longrightarrow 00:15:37.600$ A group of interconnected circuits

NOTE Confidence: 0.924246884666667

 $00{:}15{:}37.600 \dashrightarrow 00{:}15{:}40.084$ that are called the canonical reward

NOTE Confidence: 0.92424688466667

 $00:15:40.084 \longrightarrow 00:15:42.039$ seeking circuit and of course.

NOTE Confidence: 0.924246884666667

00:15:42.040 --> 00:15:45.071 This circuit as many of you know

NOTE Confidence: 0.924246884666667

 $00:15:45.071 \longrightarrow 00:15:47.270$ overlaps extensively with the with the

NOTE Confidence: 0.924246884666667

 $00{:}15{:}47.270 \dashrightarrow 00{:}15{:}49.070$ limbic system so something that doctor,

NOTE Confidence: 0.924246884666667

 $00:15:49.070 \longrightarrow 00:15:51.254$ Flynn would have been very well acquainted

NOTE Confidence: 0.924246884666667

 $00:15:51.254 \longrightarrow 00:15:53.689$ with and a circuit with in which he

NOTE Confidence: 0.924246884666667

 $00{:}15{:}53.689 \dashrightarrow 00{:}15{:}55.946$ would have spent much of his time.

NOTE Confidence: 0.924246884666667

 $00:15:55.946 \longrightarrow 00:15:58.250$ In his research efforts.

NOTE Confidence: 0.924246884666667

 $00{:}15{:}58.250 \dashrightarrow 00{:}16{:}00.722$ I'm going to focus on a subset of

NOTE Confidence: 0.924246884666667

 $00:16:00.722 \longrightarrow 00:16:02.581$ regions within this circuit here

NOTE Confidence: 0.924246884666667

 $00{:}16{:}02.581 \dashrightarrow 00{:}16{:}04.541$ depicted in this cartoon schematic

NOTE Confidence: 0.924246884666667

 $00:16:04.541 \longrightarrow 00:16:06.812$ from the rodent brain where we

NOTE Confidence: 0.924246884666667

 $00:16:06.812 \longrightarrow 00:16:08.096$ see the nucleus incumbents.

NOTE Confidence: 0.924246884666667

 $00:16:08.100 \longrightarrow 00:16:11.124$ The most ventral aspect of this striatum,

 $00:16:11.130 \longrightarrow 00:16:13.110$ where we see its output,

NOTE Confidence: 0.924246884666667

 $00:16:13.110 \longrightarrow 00:16:14.670$ then one of its outputs,

NOTE Confidence: 0.924246884666667

00:16:14.670 --> 00:16:15.786 the ventral pallidum,

NOTE Confidence: 0.924246884666667

 $00:16:15.786 \longrightarrow 00:16:18.018$ which is analogous to globis pallidus

NOTE Confidence: 0.924246884666667

 $00:16:18.018 \longrightarrow 00:16:20.524$ in more dorsal striatal circuits and

NOTE Confidence: 0.924246884666667

 $00:16:20.524 \longrightarrow 00:16:22.594$ dopaminergic input to these regions

NOTE Confidence: 0.924246884666667

 $00:16:22.657 \longrightarrow 00:16:24.547$ from the VTA and other areas that

NOTE Confidence: 0.924246884666667

 $00{:}16{:}24.547 \dashrightarrow 00{:}16{:}26.540$ we know and love like the amygdala.

NOTE Confidence: 0.924246884666667

 $00:16:26.540 \longrightarrow 00:16:29.299$ And so my lab is very interested in.

NOTE Confidence: 0.924246884666667

 $00{:}16{:}29.300 \dashrightarrow 00{:}16{:}32.660$ How these reward circuits evaluate

NOTE Confidence: 0.924246884666667

00:16:32.660 --> 00:16:35.940 reward when it's being experienced?

NOTE Confidence: 0.92424688466667

 $00:16:35.940 \longrightarrow 00:16:38.382$ And then how that current evaluation

NOTE Confidence: 0.924246884666667

 $00:16:38.382 \longrightarrow 00:16:41.024$ can impact how the animals learn

NOTE Confidence: 0.924246884666667

 $00:16:41.024 \longrightarrow 00:16:44.019$ about what just happened so that it

NOTE Confidence: 0.92424688466667

00:16:44.019 --> 00:16:45.884 can impact their future behavior?

00:16:45.890 --> 00:16:48.778 So how are rewards processed in this circuit?

NOTE Confidence: 0.924246884666667

00:16:48.780 --> 00:16:50.803 So we're going to focus today specifically

NOTE Confidence: 0.92424688466667

 $00:16:50.803 \longrightarrow 00:16:53.225$ on trying to understand how the ventral

NOTE Confidence: 0.924246884666667

 $00:16:53.225 \longrightarrow 00:16:55.095$ pallidum contributes to that process,

NOTE Confidence: 0.924246884666667

 $00:16:55.100 \longrightarrow 00:16:57.392$ and it's much more interesting than

NOTE Confidence: 0.924246884666667

00:16:57.392 --> 00:17:00.030 perhaps we once thought few decades ago,

NOTE Confidence: 0.924246884666667

 $00:17:00.030 \longrightarrow 00:17:01.698$ the ventral pallidum historically

NOTE Confidence: 0.924246884666667

 $00:17:01.698 \longrightarrow 00:17:03.783$ has been considered somewhat of

NOTE Confidence: 0.924246884666667

 $00:17:03.783 \longrightarrow 00:17:06.006$ a way station or pass through.

NOTE Confidence: 0.924246884666667

00:17:06.010 --> 00:17:08.638 For information from this stried region,

NOTE Confidence: 0.924246884666667

 $00:17:08.640 \longrightarrow 00:17:09.780$ the accompagnes,

NOTE Confidence: 0.924246884666667

00:17:09.780 --> 00:17:11.490 but instead increasingly,

NOTE Confidence: 0.924246884666667

 $00:17:11.490 \longrightarrow 00:17:13.925$ we're understanding that really important

NOTE Confidence: 0.924246884666667

00:17:13.925 --> 00:17:15.873 integrative processing is happening

NOTE Confidence: 0.924246884666667

00:17:15.873 --> 00:17:18.651 at the level of the ventral pallidum

NOTE Confidence: 0.924246884666667

 $00:17:18.651 \longrightarrow 00:17:20.750$ that impacts reward seeking behavior.

 $00:17:20.750 \longrightarrow 00:17:23.174$ So I'm going to focus in on some

NOTE Confidence: 0.924246884666667

 $00{:}17{:}23.174 \dashrightarrow 00{:}17{:}24.847$ experiments conducted in the lab that

NOTE Confidence: 0.924246884666667

00:17:24.847 --> 00:17:26.744 I hope can can illuminate the function

NOTE Confidence: 0.924246884666667

 $00:17:26.744 \longrightarrow 00:17:28.670$ of the ventral pallidum for us,

NOTE Confidence: 0.924246884666667

 $00:17:28.670 \longrightarrow 00:17:31.757$ and when you wonder what a brain region does,

NOTE Confidence: 0.924246884666667

00:17:31.760 --> 00:17:33.713 of course one of the most traditional

NOTE Confidence: 0.924246884666667

00:17:33.713 --> 00:17:36.174 ways to look is to get rid of that brain.

NOTE Confidence: 0.924246884666667

 $00:17:36.180 \longrightarrow 00:17:37.828$ Region so decades ago,

NOTE Confidence: 0.924246884666667

00:17:37.828 --> 00:17:39.888 lesions of the ventral pallidum

NOTE Confidence: 0.924246884666667

 $00{:}17{:}39.888 \dashrightarrow 00{:}17{:}42.190$ were shown to decrease intake of

NOTE Confidence: 0.924246884666667

 $00:17:42.190 \longrightarrow 00:17:44.570$ drugs of abuse in animal models.

NOTE Confidence: 0.924246884666667

 $00:17:44.570 \longrightarrow 00:17:46.880$ So decrease opiate self administration,

NOTE Confidence: 0.924246884666667 00:17:46.880 --> 00:17:48.980 for example.

NOTE Confidence: 0.924246884666667

 $00{:}17{:}48.980 \dashrightarrow 00{:}17{:}51.383$ So that tells us this this area is very

NOTE Confidence: 0.924246884666667

00:17:51.383 --> 00:17:53.228 important for reward seeking behavior,

 $00:17:53.230 \longrightarrow 00:17:55.393$ but to figure out how it's very

NOTE Confidence: 0.92424688466667

 $00:17:55.393 \longrightarrow 00:17:57.615$ instructive to go in and use

NOTE Confidence: 0.924246884666667

 $00:17:57.615 \longrightarrow 00:17:59.665$ electrophysiology to record the neural

NOTE Confidence: 0.924246884666667

00:17:59.665 --> 00:18:01.785 activity during the behavior so you

NOTE Confidence: 0.924246884666667

 $00:18:01.785 \longrightarrow 00:18:03.860$ can see what the neurons care about,

NOTE Confidence: 0.924246884666667

 $00:18:03.860 \longrightarrow 00:18:05.840$ and so a number of labs have used this

NOTE Confidence: 0.924246884666667

 $00:18:05.892 \longrightarrow 00:18:07.844$ approach and I'd like to tell you about.

NOTE Confidence: 0.924246884666667

 $00:18:07.850 \longrightarrow 00:18:09.662$ Some of the data that has

NOTE Confidence: 0.924246884666667

 $00{:}18{:}09.662 \dashrightarrow 00{:}18{:}10.870$ already emerged that set

NOTE Confidence: 0.92190800375

00:18:10.939 --> 00:18:13.117 up our thinking for our experiments.

NOTE Confidence: 0.92190800375

 $00:18:13.120 \longrightarrow 00:18:17.584$ So these are our data obtained from in vivo

NOTE Confidence: 0.92190800375

 $00:18:17.584 \longrightarrow 00:18:19.638$ electrophysiological recordings in rats.

NOTE Confidence: 0.92190800375

 $00:18:19.640 \longrightarrow 00:18:20.964$ So electrodes are implanted

NOTE Confidence: 0.92190800375

 $00:18:20.964 \longrightarrow 00:18:22.288$ into the ventral pallidum,

NOTE Confidence: 0.92190800375

00:18:22.290 --> 00:18:23.966 and we're measuring extracellularly.

NOTE Confidence: 0.92190800375

 $00:18:23.966 \longrightarrow 00:18:26.480$ The spike activity of nearby neurons,

 $00:18:26.480 \longrightarrow 00:18:27.593$ and what Jocelyn,

NOTE Confidence: 0.92190800375

00:18:27.593 --> 00:18:29.819 Richard working with Howard Fields found,

NOTE Confidence: 0.92190800375

 $00:18:29.820 \longrightarrow 00:18:33.096$ is that neurons in the ventral pallidum

NOTE Confidence: 0.92190800375

 $00:18:33.096 \longrightarrow 00:18:35.806$ care about cues that tell the animal

NOTE Confidence: 0.92190800375

00:18:35.806 --> 00:18:37.850 it has an opportunity to get reward,

NOTE Confidence: 0.92190800375

 $00:18:37.850 \longrightarrow 00:18:39.943$ and so here we see an average

NOTE Confidence: 0.92190800375

 $00:18:39.943 \longrightarrow 00:18:41.859$ increase in activity when the animal.

NOTE Confidence: 0.92190800375

 $00:18:41.860 \longrightarrow 00:18:43.016$ Here's a Q, but.

NOTE Confidence: 0.92190800375

 $00{:}18{:}43.016 \dashrightarrow 00{:}18{:}45.165$ The increase in activity is much bigger

NOTE Confidence: 0.92190800375

 $00{:}18{:}45.165 \dashrightarrow 00{:}18{:}46.905$ when the animals actually motivated

NOTE Confidence: 0.92190800375

 $00:18:46.905 \longrightarrow 00:18:49.429$ to press a lever to get the reward,

NOTE Confidence: 0.92190800375

 $00:18:49.430 \longrightarrow 00:18:51.630$ so the the larger signal is from trials

NOTE Confidence: 0.92190800375

 $00{:}18{:}51.630 \dashrightarrow 00{:}18{:}53.917$ when animals press the lever to get reward

NOTE Confidence: 0.92190800375

 $00:18:53.917 \longrightarrow 00:18:56.049$ and the smaller signals when they fail to.

NOTE Confidence: 0.92190800375

 $00:18:56.050 \longrightarrow 00:18:58.498$ So motivation already is coming into

 $00:18:58.498 \longrightarrow 00:19:01.446$ play and modulating the way the ventral

NOTE Confidence: 0.92190800375

 $00{:}19{:}01.446 {\:\dashrightarrow\:} 00{:}19{:}03.526$ pallidal neurons respond to cues.

NOTE Confidence: 0.92190800375

 $00:19:03.530 \longrightarrow 00:19:06.092$ The valence of a reward also modulates

NOTE Confidence: 0.92190800375

 $00:19:06.092 \longrightarrow 00:19:08.390$ responses in the ventral pallidum.

NOTE Confidence: 0.92190800375

 $00:19:08.390 \longrightarrow 00:19:09.750$ So, as you might predict,

NOTE Confidence: 0.92190800375

00:19:09.750 --> 00:19:12.254 given the data I told you about lesions,

NOTE Confidence: 0.92190800375

 $00:19:12.260 \longrightarrow 00:19:13.600$ the ventral pallidum cares

NOTE Confidence: 0.92190800375

 $00:19:13.600 \longrightarrow 00:19:15.610$ about the nature of the reward.

NOTE Confidence: 0.92190800375

 $00:19:15.610 \longrightarrow 00:19:17.440$ So in this example from Kendall

NOTE Confidence: 0.92190800375

00:19:17.440 --> 00:19:19.619 at all this this classic paper,

NOTE Confidence: 0.92190800375

 $00{:}19{:}19.620 \dashrightarrow 00{:}19{:}21.858$ we see examples from 1 neuron

NOTE Confidence: 0.92190800375

 $00:19:21.858 \longrightarrow 00:19:23.970$ and its response to two QS.

NOTE Confidence: 0.92190800375

00:19:23.970 --> 00:19:26.266 One is a Q that predicts something good,

NOTE Confidence: 0.92190800375

 $00:19:26.270 \longrightarrow 00:19:28.106$ sucrose solution and one is a

NOTE Confidence: 0.92190800375

 $00:19:28.106 \longrightarrow 00:19:29.679$ cue that predicts something the

NOTE Confidence: 0.92190800375

 $00:19:29.679 \longrightarrow 00:19:31.425$ animal doesn't like a salty taste.

 $00:19:31.430 \longrightarrow 00:19:33.278$ We see the neuron has a big

NOTE Confidence: 0.92190800375

 $00:19:33.278 \longrightarrow 00:19:34.539$ increase in firing to the.

NOTE Confidence: 0.92190800375 00:19:34.540 --> 00:19:34.949 Q. NOTE Confidence: 0.92190800375

00:19:34.949 --> 00:19:36.176 Predicting something good

NOTE Confidence: 0.92190800375

 $00:19:36.176 \longrightarrow 00:19:38.630$ and not to the salt Q.

NOTE Confidence: 0.92190800375

00:19:38.630 --> 00:19:40.712 But if we use pharmacology to

NOTE Confidence: 0.92190800375

 $00:19:40.712 \longrightarrow 00:19:42.500$ deplete the subjects of salt,

NOTE Confidence: 0.92190800375

 $00:19:42.500 \dashrightarrow 00:19:45.524$ so we'd make them desire salt and want salt.

NOTE Confidence: 0.92190800375

 $00{:}19{:}45.530 \dashrightarrow 00{:}19{:}47.750$ We see suddenly that this neuron

NOTE Confidence: 0.92190800375

 $00{:}19{:}47.750 \dashrightarrow 00{:}19{:}49.653$ now responds with an increase

NOTE Confidence: 0.92190800375

 $00:19:49.653 \longrightarrow 00:19:51.747$ in activity to the salt Q.

NOTE Confidence: 0.92190800375

 $00:19:51.750 \longrightarrow 00:19:54.174$ Reminiscent of the response to the sucrose Q.

NOTE Confidence: 0.92190800375

 $00{:}19{:}54.180 \dashrightarrow 00{:}19{:}57.438$ So this is a clue that the Q responses

NOTE Confidence: 0.92190800375

 $00:19:57.438 \longrightarrow 00:19:59.923$ responses in the VP are sensitive to

NOTE Confidence: 0.92190800375

00:19:59.923 --> 00:20:02.461 the valence of the reward the animal

 $00:20:02.461 \longrightarrow 00:20:05.274$ is going to get, so it cares about.

NOTE Confidence: 0.92190800375

00:20:05.274 --> 00:20:06.618 Motivational state it cares

NOTE Confidence: 0.92190800375

00:20:06.618 --> 00:20:08.677 about the valence of the reward.

NOTE Confidence: 0.92190800375

 $00:20:08.680 \longrightarrow 00:20:09.566$ And interestingly,

NOTE Confidence: 0.92190800375

 $00:20:09.566 \longrightarrow 00:20:12.224$ in this work from bullies Lab,

NOTE Confidence: 0.92190800375

 $00:20:12.230 \longrightarrow 00:20:14.315$ it also cares about what's

NOTE Confidence: 0.92190800375

00:20:14.315 --> 00:20:16.400 actually happening in real time.

NOTE Confidence: 0.92190800375

 $00:20:16.400 \longrightarrow 00:20:18.129$ So here we see an example of

NOTE Confidence: 0.92190800375

00:20:18.129 --> 00:20:20.118 data from a paper recording from

NOTE Confidence: 0.92190800375

00:20:20.118 --> 00:20:22.093 neurons in the ventral pallidum

NOTE Confidence: 0.92190800375

 $00:20:22.093 \longrightarrow 00:20:24.552$ in subjects that receive the queue

NOTE Confidence: 0.92190800375

 $00:20:24.552 \longrightarrow 00:20:26.156$ that predicts sucrose reward.

NOTE Confidence: 0.92190800375

 $00:20:26.160 \longrightarrow 00:20:27.816$ The neurons respond to the queue,

NOTE Confidence: 0.92190800375

 $00{:}20{:}27.820 \dashrightarrow 00{:}20{:}29.770$ and they respond to the reward.

NOTE Confidence: 0.92190800375

00:20:29.770 --> 00:20:32.322 But if on some trials you omit reward

NOTE Confidence: 0.92190800375

 $00{:}20{:}32.322 \dashrightarrow 00{:}20{:}35.045$ that you see that there is a decrease.

 $00:20:35.050 \longrightarrow 00:20:37.090$ In firing by these neurons,

NOTE Confidence: 0.92190800375

 $00:20:37.090 \longrightarrow 00:20:39.268$ this tells us that neurons have

NOTE Confidence: 0.92190800375

 $00:20:39.268 \longrightarrow 00:20:41.547$ an expectation of the reward that

NOTE Confidence: 0.92190800375

 $00:20:41.547 \longrightarrow 00:20:43.809$ they should receive after the Q.

NOTE Confidence: 0.92190800375

 $00:20:43.810 \longrightarrow 00:20:45.970$ This is reminiscent of a negative

NOTE Confidence: 0.92190800375

 $00:20:45.970 \longrightarrow 00:20:47.694$ reward prediction error signal that

NOTE Confidence: 0.92190800375

 $00:20:47.694 \longrightarrow 00:20:50.335$ many of you may be familiar with from

NOTE Confidence: 0.92190800375

 $00:20:50.335 \longrightarrow 00:20:52.300$ thinking about how dopamine neurons

NOTE Confidence: 0.92190800375

 $00:20:52.374 \longrightarrow 00:20:55.279$ respond when expected reward fails to arrive.

NOTE Confidence: 0.92190800375

 $00:20:55.280 \longrightarrow 00:20:57.650$ So together these give us important

NOTE Confidence: 0.92190800375

 $00:20:57.650 \longrightarrow 00:21:00.157$ clues that neurons in the ventral

NOTE Confidence: 0.92190800375

00:21:00.157 --> 00:21:01.865 pallidum respond to cues.

NOTE Confidence: 0.92190800375

 $00:21:01.870 \longrightarrow 00:21:03.030$ They respond to reward,

NOTE Confidence: 0.92190800375

 $00{:}21{:}03.030 \dashrightarrow 00{:}21{:}05.270$ and they do so in interesting manners.

NOTE Confidence: 0.92190800375

 $00:21:05.270 \longrightarrow 00:21:07.010$ They care about the motivational state.

 $00:21:07.010 \longrightarrow 00:21:09.250$ They care about the valence of the reward,

NOTE Confidence: 0.906665728571428

00:21:09.250 --> 00:21:11.308 how much the subject likes the reward,

NOTE Confidence: 0.906665728571428

 $00:21:11.310 \longrightarrow 00:21:13.800$ and they have some sort of

NOTE Confidence: 0.906665728571428

 $00:21:13.800 \longrightarrow 00:21:14.630$ expectation information.

NOTE Confidence: 0.906665728571428

 $00:21:14.630 \longrightarrow 00:21:15.682$ They care about what's

NOTE Confidence: 0.906665728571428

00:21:15.682 --> 00:21:16.734 happening in real time.

NOTE Confidence: 0.906665728571428

 $00:21:16.740 \longrightarrow 00:21:19.140$ If the reward arrives or not.

NOTE Confidence: 0.906665728571428

 $00:21:19.140 \longrightarrow 00:21:21.258$ So these and many other levely

NOTE Confidence: 0.906665728571428

 $00{:}21{:}21.258 \dashrightarrow 00{:}21{:}24.022$ studies set the stage for the kinds

NOTE Confidence: 0.906665728571428

00:21:24.022 --> 00:21:26.147 of questions that David Ottenheimer,

NOTE Confidence: 0.906665728571428

 $00{:}21{:}26.150 \mathrel{--}{>} 00{:}21{:}27.650$ a graduate student in the lab,

NOTE Confidence: 0.906665728571428

 $00:21:27.650 \longrightarrow 00:21:30.254$ wanted to ask when he wondered about

NOTE Confidence: 0.906665728571428

 $00:21:30.254 \longrightarrow 00:21:32.826$ the details of how the outcomes

NOTE Confidence: 0.906665728571428

 $00:21:32.826 \longrightarrow 00:21:35.592$ themselves are processed by the neurons.

NOTE Confidence: 0.906665728571428

 $00:21:35.600 \longrightarrow 00:21:36.434$ In ventral pallidum.

NOTE Confidence: 0.906665728571428

 $00:21:36.434 \longrightarrow 00:21:38.790$ So David was a graduate student in my lab.

 $00:21:38.790 \longrightarrow 00:21:40.920$ He's now a postdoc in the

NOTE Confidence: 0.906665728571428

 $00:21:40.920 \longrightarrow 00:21:42.340$ Steinmetz and Stuber Labs,

NOTE Confidence: 0.906665728571428

00:21:42.340 --> 00:21:44.601 and he was aided through all of

NOTE Confidence: 0.906665728571428

00:21:44.601 --> 00:21:46.674 this with by Doctor Joslin Richard,

NOTE Confidence: 0.906665728571428

 $00{:}21{:}46.674 \dashrightarrow 00{:}21{:}49.560$ when she was a senior scientist in the lab.

NOTE Confidence: 0.906665728571428

 $00:21:49.560 \longrightarrow 00:21:52.199$ She was our resident ventral pallidum expert.

NOTE Confidence: 0.906665728571428

 $00:21:52.200 \longrightarrow 00:21:54.330$ Doctor Richard now runs her own

NOTE Confidence: 0.906665728571428

 $00{:}21{:}54.330 \rightarrow 00{:}21{:}56.689$ lab at the University of Minnesota,

NOTE Confidence: 0.906665728571428

 $00:21:56.690 \longrightarrow 00:21:58.590$ so together they designed a

NOTE Confidence: 0.906665728571428

 $00{:}21{:}58.590 \dashrightarrow 00{:}22{:}01.000$ series of studies to allow them

NOTE Confidence: 0.906665728571428

 $00:22:01.000 \longrightarrow 00:22:03.586$ to understand better how ventral

NOTE Confidence: 0.906665728571428

 $00:22:03.586 \longrightarrow 00:22:06.118$ pallidum neurons encode natural.

NOTE Confidence: 0.906665728571428

 $00:22:06.120 \longrightarrow 00:22:09.403$ Word outcomes and so that David was

NOTE Confidence: 0.906665728571428

 $00:22:09.403 \longrightarrow 00:22:11.950$ interested in doing these studies

NOTE Confidence: 0.906665728571428

 $00:22:11.950 \longrightarrow 00:22:15.088$ in the setting of multiple rewards.

00:22:15.090 --> 00:22:17.185 Because eventually we'd like to

NOTE Confidence: 0.906665728571428

 $00{:}22{:}17.185 \dashrightarrow 00{:}22{:}19.280$ understand how agents make choices

NOTE Confidence: 0.906665728571428

00:22:19.346 --> 00:22:21.656 among rewards because we'd like to apply

NOTE Confidence: 0.906665728571428

 $00:22:21.656 \longrightarrow 00:22:24.300$ this in the future to drug addiction.

NOTE Confidence: 0.906665728571428

 $00:22:24.300 \longrightarrow 00:22:27.620$ How to agents choose drugs or other rewards.

NOTE Confidence: 0.906665728571428

 $00:22:27.620 \longrightarrow 00:22:31.070$ So David began with very simple.

NOTE Confidence: 0.906665728571428

 $00:22:31.070 \longrightarrow 00:22:32.745$ Experimental designs where he could

NOTE Confidence: 0.906665728571428

 $00:22:32.745 \longrightarrow 00:22:34.835$ look at the activity of ventral

NOTE Confidence: 0.906665728571428

 $00{:}22{:}34.835 \dashrightarrow 00{:}22{:}36.620$ pallidal neurons when rats were

NOTE Confidence: 0.906665728571428

 $00:22:36.620 \longrightarrow 00:22:38.510$ receiving more than one reward.

NOTE Confidence: 0.906665728571428

 $00{:}22{:}38.510 \dashrightarrow 00{:}22{:}41.366$ So in this very initial simple design,

NOTE Confidence: 0.906665728571428

 $00:22:41.370 \longrightarrow 00:22:42.842$ he implanted electrodes into

NOTE Confidence: 0.906665728571428

 $00:22:42.842 \longrightarrow 00:22:44.682$ the ventral pallidum of rats,

NOTE Confidence: 0.906665728571428

 $00:22:44.690 \longrightarrow 00:22:46.598$ recorded extracellular spike activity.

NOTE Confidence: 0.906665728571428

 $00:22:46.598 \longrightarrow 00:22:48.983$ These are waveforms of example

NOTE Confidence: 0.906665728571428

 $00:22:48.983 \longrightarrow 00:22:50.579$ neurons that he recorded,

 $00:22:50.580 \longrightarrow 00:22:51.830$ and here's again our cartoon

NOTE Confidence: 0.906665728571428

 $00:22:51.830 \longrightarrow 00:22:52.830$ of the ventral pallidum.

NOTE Confidence: 0.906665728571428

 $00:22:52.830 \longrightarrow 00:22:55.656$ So the electrode tips are residing in the VP,

NOTE Confidence: 0.906665728571428

 $00:22:55.660 \longrightarrow 00:22:58.915$ and he exposed subjects to

NOTE Confidence: 0.906665728571428

00:22:58.915 --> 00:23:01.519 two different rewards liquid.

NOTE Confidence: 0.906665728571428

 $00:23:01.520 \longrightarrow 00:23:03.104$ Sucrose and maltodextrin.

NOTE Confidence: 0.906665728571428

 $00:23:03.104 \longrightarrow 00:23:05.216$ These are both carbohydrates,

NOTE Confidence: 0.906665728571428

00:23:05.220 --> 00:23:06.058 calorically equivalent,

NOTE Confidence: 0.906665728571428

 $00:23:06.058 \longrightarrow 00:23:09.410$ and that we know rats like they will

NOTE Confidence: 0.906665728571428

 $00{:}23{:}09.481 \dashrightarrow 00{:}23{:}12.120$ drink then avidly and he exposed them

NOTE Confidence: 0.906665728571428

 $00:23:12.120 \longrightarrow 00:23:14.859$ to these rewards in recording sessions,

NOTE Confidence: 0.906665728571428

 $00:23:14.860 \longrightarrow 00:23:17.062$ and the rewards were delivered randomly

NOTE Confidence: 0.906665728571428

 $00{:}23{:}17.062 \dashrightarrow 00{:}23{:}19.245$ so the animal didn't know which

NOTE Confidence: 0.906665728571428

 $00:23:19.245 \longrightarrow 00:23:21.527$ reward was coming on any given trial.

NOTE Confidence: 0.90666572857142800:23:21.530 --> 00:23:21.975 Specifically,

 $00:23:21.975 \longrightarrow 00:23:25.535$ he played the queue so white noise Q,

NOTE Confidence: 0.906665728571428

 $00:23:25.540 \longrightarrow 00:23:26.814$ and when the animal heard the cue,

NOTE Confidence: 0.906665728571428

 $00:23:26.820 \longrightarrow 00:23:29.340$ the animal knew it could go to the port

NOTE Confidence: 0.906665728571428

 $00:23:29.340 \longrightarrow 00:23:31.660$ when it put its snout in the reward.

NOTE Confidence: 0.906665728571428

00:23:31.660 --> 00:23:33.064 Port Reward was delivered,

NOTE Confidence: 0.906665728571428

 $00:23:33.064 \longrightarrow 00:23:35.170$ and then there's an Inter trial

NOTE Confidence: 0.906665728571428

 $00:23:35.233 \longrightarrow 00:23:36.803$ interval and that occurs again

NOTE Confidence: 0.906665728571428

 $00:23:36.803 \longrightarrow 00:23:39.108$ and so the the reward cannot be

NOTE Confidence: 0.906665728571428

 $00{:}23{:}39.108 \dashrightarrow 00{:}23{:}40.440$ predicted by the queue.

NOTE Confidence: 0.906665728571428

 $00:23:40.440 \longrightarrow 00:23:42.687$ The animal has to actually wait till

NOTE Confidence: 0.906665728571428

 $00{:}23{:}42.687 {\:{\mbox{--}}\!>}\ 00{:}23{:}44.541$ the reward squirt it out before

NOTE Confidence: 0.906665728571428

 $00:23:44.541 \longrightarrow 00:23:46.459$ it knows what it's getting so we

NOTE Confidence: 0.906665728571428

 $00:23:46.521 \longrightarrow 00:23:48.276$ can compare the neural response

NOTE Confidence: 0.906665728571428

 $00:23:48.276 \longrightarrow 00:23:49.680$ to these two rewards.

NOTE Confidence: 0.906665728571428

 $00:23:49.680 \longrightarrow 00:23:51.048$ There's an interesting feature

NOTE Confidence: 0.906665728571428

00:23:51.048 --> 00:23:53.320 about these two rewards and that is,

 $00:23:53.320 \longrightarrow 00:23:56.176$ although rats love both of them,

NOTE Confidence: 0.906665728571428

 $00{:}23{:}56.180 \dashrightarrow 00{:}23{:}57.652$ if you give them a full bottle of

NOTE Confidence: 0.906665728571428

00:23:57.652 --> 00:23:59.360 one or the other on their homepage,

NOTE Confidence: 0.906665728571428

 $00:23:59.360 \longrightarrow 00:24:00.820$ they'll drink it all up.

NOTE Confidence: 0.906665728571428

 $00:24:00.820 \longrightarrow 00:24:02.446$ If you give them two bottles.

NOTE Confidence: 0.906665728571428

 $00:24:02.450 \longrightarrow 00:24:03.428$ One with sucrose,

NOTE Confidence: 0.906665728571428

00:24:03.428 --> 00:24:05.710 one with maltodextrin at the same time,

NOTE Confidence: 0.906665728571428

00:24:05.710 --> 00:24:08.150 most rats prefer the sucrose,

NOTE Confidence: 0.906665728571428

00:24:08.150 --> 00:24:09.692 and that's what I'm showing here

NOTE Confidence: 0.906665728571428

 $00{:}24{:}09.692 \dashrightarrow 00{:}24{:}11.785$ in this behavioral data figure.

NOTE Confidence: 0.906665728571428

00:24:11.785 --> 00:24:14.222 This shows the preference subjects

NOTE Confidence: 0.906665728571428

 $00:24:14.222 \longrightarrow 00:24:15.550$ have for sucrose over

NOTE Confidence: 0.906665728571428

 $00{:}24{:}15.550 \dashrightarrow 00{:}24{:}17.210$ maltod extrin when tested in the

NOTE Confidence: 0.8566711805

00:24:17.269 --> 00:24:18.609 home cage when they just

NOTE Confidence: 0.8566711805

 $00:24:18.609 \longrightarrow 00:24:19.949$ have big bottles of both,

 $00:24:19.950 \longrightarrow 00:24:21.940$ they'll drink more of the

NOTE Confidence: 0.8566711805

 $00:24:21.940 \longrightarrow 00:24:23.532$ sucrose than multidex turn.

NOTE Confidence: 0.8566711805

 $00:24:23.540 \longrightarrow 00:24:25.280$ However, in this behavioral session

NOTE Confidence: 0.8566711805

 $00:24:25.280 \longrightarrow 00:24:27.800$ where we're giving A Q and then

NOTE Confidence: 0.8566711805

 $00:24:27.800 \longrightarrow 00:24:29.600$ squirting out maltodextrin or sucrose

NOTE Confidence: 0.8566711805

 $00{:}24{:}29.600 \dashrightarrow 00{:}24{:}32.030$ and they have to drink it in order

NOTE Confidence: 0.8566711805

 $00:24:32.030 \longrightarrow 00:24:33.753$ to have the next trial happen,

NOTE Confidence: 0.8566711805

 $00:24:33.753 \longrightarrow 00:24:36.351$ we see that the licking behavior

NOTE Confidence: 0.8566711805

 $00{:}24{:}36.351 \dashrightarrow 00{:}24{:}38.684$ when they're consuming the different

NOTE Confidence: 0.8566711805

 $00:24:38.684 \longrightarrow 00:24:40.608$ rewards is almost identical.

NOTE Confidence: 0.8566711805

 $00:24:40.610 \longrightarrow 00:24:42.038$ So we're left with a nice,

NOTE Confidence: 0.8566711805

 $00:24:42.040 \longrightarrow 00:24:44.284$ very simple behavioral model where their

NOTE Confidence: 0.8566711805

 $00:24:44.284 \longrightarrow 00:24:47.090$ preference for the two rewards is different.

NOTE Confidence: 0.8566711805

 $00:24:47.090 \longrightarrow 00:24:49.176$ We know that based on these

NOTE Confidence: 0.8566711805

 $00:24:49.176 \longrightarrow 00:24:50.600$ long term drinking studies,

NOTE Confidence: 0.8566711805

00:24:50.600 --> 00:24:53.066 but their motor behavior during this

00:24:53.066 --> 00:24:55.810 particular very simple task is very similar,

NOTE Confidence: 0.8566711805

 $00{:}24{:}55.810 \dashrightarrow 00{:}24{:}57.434$ so that gives us a nice way to

NOTE Confidence: 0.8566711805

 $00:24:57.434 \longrightarrow 00:24:59.361$ see what the signal related to the

NOTE Confidence: 0.8566711805

00:24:59.361 --> 00:25:00.821 preference might be when we're

NOTE Confidence: 0.8566711805

 $00:25:00.880 \longrightarrow 00:25:02.668$ basically making sure that the motor

NOTE Confidence: 0.8566711805

 $00:25:02.668 \longrightarrow 00:25:04.249$ behavior is not that different,

NOTE Confidence: 0.8566711805

00:25:04.249 --> 00:25:06.194 because that could motor behavior

NOTE Confidence: 0.8566711805

 $00:25:06.194 \longrightarrow 00:25:08.125$ could be an explanation for

NOTE Confidence: 0.8566711805

 $00:25:08.125 \longrightarrow 00:25:09.695$ some differences that we see.

NOTE Confidence: 0.8566711805

 $00:25:09.700 \longrightarrow 00:25:11.144$ So in the face.

NOTE Confidence: 0.8566711805

 $00:25:11.144 \longrightarrow 00:25:12.949$ In this very simple behavior,

NOTE Confidence: 0.8566711805

 $00:25:12.950 \longrightarrow 00:25:14.966$ what David found when he recorded from

NOTE Confidence: 0.8566711805

00:25:14.966 --> 00:25:17.010 many neurons in the ventral pallidum,

NOTE Confidence: 0.8566711805

 $00:25:17.010 \longrightarrow 00:25:19.176$ many individual neurons is that there's

NOTE Confidence: 0.8566711805

 $00:25:19.176 \longrightarrow 00:25:22.124$ a big difference in the way the neuron

 $00:25:22.124 \longrightarrow 00:25:23.924$ signal which reward they received.

NOTE Confidence: 0.8566711805

00:25:23.930 --> 00:25:27.115 So here I'm showing the average activity

NOTE Confidence: 0.8566711805

 $00:25:27.115 \longrightarrow 00:25:30.180$ of 205 neurons that were sensitive to

NOTE Confidence: 0.8566711805

 $00:25:30.180 \longrightarrow 00:25:32.630$ reward based on statistical analysis.

NOTE Confidence: 0.8566711805

 $00:25:32.630 \longrightarrow 00:25:34.966$ If we divide the trials into those in

NOTE Confidence: 0.8566711805

 $00:25:34.966 \longrightarrow 00:25:37.789$ which the animal receives sucrose and orange,

NOTE Confidence: 0.8566711805

 $00:25:37.790 \longrightarrow 00:25:40.800$ or maltodextrin in this pink purple color.

NOTE Confidence: 0.8566711805

 $00:25:40.800 \longrightarrow 00:25:43.072$ We see an average very large increase in

NOTE Confidence: 0.8566711805

 $00{:}25{:}43.072 \dashrightarrow 00{:}25{:}45.010$ activity when the animals are drinking.

NOTE Confidence: 0.8566711805

 $00:25:45.010 \longrightarrow 00:25:47.730$ The sucrose 0 is the time that rewards

NOTE Confidence: 0.8566711805

 $00:25:47.730 \longrightarrow 00:25:49.660$ delivered the first few seconds,

NOTE Confidence: 0.8566711805

 $00:25:49.660 \longrightarrow 00:25:51.837$ first three to four seconds is when

NOTE Confidence: 0.8566711805

00:25:51.837 --> 00:25:53.140 they're actually lapping it up.

NOTE Confidence: 0.8566711805

 $00:25:53.140 \longrightarrow 00:25:56.084$ We see a much lower response by the

NOTE Confidence: 0.8566711805

 $00:25:56.084 \longrightarrow 00:25:58.708$ population when maltodextrin is received.

NOTE Confidence: 0.8566711805

 $00{:}25{:}58.710 \dashrightarrow 00{:}26{:}00.677$ These heat maps here show you the

 $00:26:00.677 \longrightarrow 00:26:02.358$ activity of the individual neurons

NOTE Confidence: 0.8566711805

 $00:26:02.358 \longrightarrow 00:26:04.288$ that make up these averages,

NOTE Confidence: 0.8566711805

 $00:26:04.290 \longrightarrow 00:26:06.626$ so again we have the same time course

NOTE Confidence: 0.8566711805

 $00:26:06.626 \longrightarrow 00:26:09.454$ and each row is the color coded map of

NOTE Confidence: 0.8566711805

 $00:26:09.454 \longrightarrow 00:26:11.558$ the spike intensity from that neuron

NOTE Confidence: 0.8566711805

 $00:26:11.558 \longrightarrow 00:26:14.134$ arranged by most intense to less intense.

NOTE Confidence: 0.8566711805

00:26:14.140 --> 00:26:16.228 And you can see here by I many,

NOTE Confidence: 0.8566711805

 $00:26:16.230 \longrightarrow 00:26:17.770$ many neurons are showing an

NOTE Confidence: 0.8566711805

00:26:17.770 --> 00:26:19.310 increase at this exact time,

NOTE Confidence: 0.8566711805

 $00:26:19.310 \longrightarrow 00:26:21.697$ and it's much less present and sometimes

NOTE Confidence: 0.8566711805

 $00:26:21.697 \longrightarrow 00:26:24.598$ even more of a decrease for maltodextrin.

NOTE Confidence: 0.8566711805

 $00:26:24.600 \longrightarrow 00:26:26.959$ So the populations in the ventral pallidum

NOTE Confidence: 0.8566711805

 $00:26:26.959 \longrightarrow 00:26:29.179$ encode these two rewards differently.

NOTE Confidence: 0.8566711805

 $00:26:29.180 \longrightarrow 00:26:31.460$ Although the drinking behavior similar the

NOTE Confidence: 0.8566711805

 $00:26:31.460 \longrightarrow 00:26:33.720$ preference these subjects have is different,

 $00:26:33.720 \longrightarrow 00:26:37.086$ and that may be what is we're seeing here.

NOTE Confidence: 0.8566711805

00:26:37.090 --> 00:26:37.530 Alternatively,

NOTE Confidence: 0.8566711805

 $00:26:37.530 \longrightarrow 00:26:39.290$ you might propose will.

NOTE Confidence: 0.8566711805

 $00:26:39.290 \longrightarrow 00:26:42.468$ Sucrose is a very important natural sugar.

NOTE Confidence: 0.8566711805

00:26:42.470 --> 00:26:45.116 Maybe neurons in the brain are set

NOTE Confidence: 0.8566711805

00:26:45.116 --> 00:26:48.060 up already to fire in a very specific

NOTE Confidence: 0.8566711805

 $00:26:48.060 \longrightarrow 00:26:50.820$ way to sucrose as as it taste it.

NOTE Confidence: 0.8566711805

 $00:26:50.820 \longrightarrow 00:26:53.081$ So it could be that these responses

NOTE Confidence: 0.8566711805

 $00{:}26{:}53.081 \dashrightarrow 00{:}26{:}55.227$ are fixed and that they really

NOTE Confidence: 0.8566711805

 $00:26:55.227 \longrightarrow 00:26:56.699$ depend on the rewards.

NOTE Confidence: 0.8566711805

 $00:26:56.700 \longrightarrow 00:26:59.260$ So David tried to think of a way to to

NOTE Confidence: 0.8566711805

 $00{:}26{:}59.334 \longrightarrow 00{:}27{:}02.012$ examine that so to do that he repeated

NOTE Confidence: 0.8566711805

 $00:27:02.012 \longrightarrow 00:27:04.778$ the same behavioral procedure but he

NOTE Confidence: 0.8566711805

00:27:04.778 --> 00:27:07.396 swapped water for sucrose. So now.

NOTE Confidence: 0.8566711805

 $00:27:07.396 \longrightarrow 00:27:10.004$ He's going to give the animals up.

NOTE Confidence: 0.8566711805

 $00:27:10.004 \longrightarrow 00:27:10.852$ Interleaved sessions,

 $00:27:10.852 \longrightarrow 00:27:13.820$ when they receive sucrose or water after

NOTE Confidence: 0.8566711805

 $00:27:13.886 \longrightarrow 00:27:16.646$ the queue and you could see their behavior.

NOTE Confidence: 0.850628229285714

00:27:16.650 --> 00:27:18.425 In fact, they're licking behavior

NOTE Confidence: 0.850628229285714

 $00:27:18.425 \longrightarrow 00:27:20.200$ is different from water because

NOTE Confidence: 0.850628229285714

 $00{:}27{:}20.256 \dashrightarrow 00{:}27{:}21.840$ they're not water restricted.

NOTE Confidence: 0.850628229285714

00:27:21.840 --> 00:27:24.784 So they don't really want water very much

NOTE Confidence: 0.850628229285714

 $00:27:24.790 \longrightarrow 00:27:28.612$ and what he saw neurally is a switch in

NOTE Confidence: 0.850628229285714

 $00{:}27{:}28.612 \dashrightarrow 00{:}27{:}31.052$ the way neurons encoded maltod extrin.

NOTE Confidence: 0.850628229285714

 $00{:}27{:}31.060 \to 00{:}27{:}33.268$ So remember the activity for Maltod extrin

NOTE Confidence: 0.850628229285714

 $00{:}27{:}33.268 \dashrightarrow 00{:}27{:}35.868$ was much lower than for sucrose when

NOTE Confidence: 0.850628229285714

 $00:27:35.868 \longrightarrow 00:27:38.360$ those were the two rewards being compared.

NOTE Confidence: 0.850628229285714

 $00:27:38.360 \longrightarrow 00:27:40.670$ But now when we are comparing maltodextrin

NOTE Confidence: 0.850628229285714

 $00{:}27{:}40.670 \longrightarrow 00{:}27{:}43.340$ and water as the animals taste each one,

NOTE Confidence: 0.850628229285714

 $00:27:43.340 \longrightarrow 00:27:45.545$ we see a relative increase in the

NOTE Confidence: 0.850628229285714

 $00:27:45.545 \longrightarrow 00:27:47.083$ response from maltodextrin and a

 $00:27:47.083 \longrightarrow 00:27:48.697$ decrease in the response for water.

NOTE Confidence: 0.850628229285714

 $00:27:48.700 \longrightarrow 00:27:50.368$ And you could see that very

NOTE Confidence: 0.850628229285714

 $00:27:50.368 \longrightarrow 00:27:51.820$ clearly in these heat maps.

NOTE Confidence: 0.850628229285714

00:27:51.820 --> 00:27:54.750 Here's a sucrose here's water,

NOTE Confidence: 0.850628229285714

 $00:27:54.750 \longrightarrow 00:27:57.165$ but most interesting focus on

NOTE Confidence: 0.850628229285714

00:27:57.165 --> 00:27:59.097 maltodextrin relatively low activity

NOTE Confidence: 0.850628229285714

00:27:59.097 --> 00:28:01.239 across the population now very.

NOTE Confidence: 0.850628229285714

 $00:28:01.240 \longrightarrow 00:28:04.300$ High activity across the population.

NOTE Confidence: 0.850628229285714

 $00{:}28{:}04.300 \dashrightarrow 00{:}28{:}06.456$ So this readout is not fixed based

NOTE Confidence: 0.850628229285714

 $00{:}28{:}06.456 \dashrightarrow 00{:}28{:}08.880$ on the the actual chemical nature of

NOTE Confidence: 0.850628229285714

 $00{:}28{:}08.880 \longrightarrow 00{:}28{:}11.708$ the taste it so instead it it seems

NOTE Confidence: 0.850628229285714

 $00:28:11.708 \longrightarrow 00:28:14.172$ as if perhaps it relates more to the

NOTE Confidence: 0.850628229285714

 $00:28:14.180 \longrightarrow 00:28:17.480$ animal's current preference for example.

NOTE Confidence: 0.850628229285714

 $00:28:17.480 \longrightarrow 00:28:19.699$ And we can see very exactly similar

NOTE Confidence: 0.850628229285714

00:28:19.699 --> 00:28:21.805 results if we run a behavioral

NOTE Confidence: 0.850628229285714

 $00{:}28{:}21.805 \dashrightarrow 00{:}28{:}24.450$ session with all three liquid's

 $00:28:24.450 \longrightarrow 00:28:27.050$ randomly presented after the Q,

NOTE Confidence: 0.850628229285714

 $00{:}28{:}27.050 \dashrightarrow 00{:}28{:}30.038$ 2 hour rats and we see a much higher

NOTE Confidence: 0.850628229285714

 $00:28:30.038 \longrightarrow 00:28:32.982$ average neural response to sucrose medium

NOTE Confidence: 0.850628229285714

 $00:28:32.982 \longrightarrow 00:28:36.350$ for maltodextrin and big decrease for water.

NOTE Confidence: 0.850628229285714

 $00:28:36.350 \longrightarrow 00:28:38.898$ So so there there's a ranking in

NOTE Confidence: 0.850628229285714

 $00:28:38.898 \longrightarrow 00:28:40.983$ the neural activity that fits what

NOTE Confidence: 0.850628229285714

 $00:28:40.983 \longrightarrow 00:28:43.880$ we might think of as the ranking of

NOTE Confidence: 0.850628229285714

 $00{:}28{:}43.880 \dashrightarrow 00{:}28{:}45.908$ the animal subjective preference.

NOTE Confidence: 0.850628229285714

 $00:28:45.910 \longrightarrow 00:28:47.548$ So that's one way we could.

NOTE Confidence: 0.850628229285714

 $00{:}28{:}47.550 \dashrightarrow 00{:}28{:}49.236$ We could wonder about what the

NOTE Confidence: 0.850628229285714

 $00:28:49.236 \longrightarrow 00:28:50.810$ signal means for the animal.

NOTE Confidence: 0.850628229285714

 $00:28:50.810 \longrightarrow 00:28:53.826$ Is this just a readout of the animal's

NOTE Confidence: 0.850628229285714

 $00{:}28{:}53.826 \rightarrow 00{:}28{:}55.380$ current subjective preference?

NOTE Confidence: 0.850628229285714

00:28:55.380 --> 00:28:57.390 Another idea that David had when

NOTE Confidence: 0.850628229285714

 $00:28:57.390 \longrightarrow 00:28:59.682$ looking at this is that this signal

 $00:28:59.682 \longrightarrow 00:29:01.747$ also could be a readout of a

NOTE Confidence: 0.850628229285714

 $00:29:01.820 \longrightarrow 00:29:03.660$ difference from the animals,

NOTE Confidence: 0.850628229285714

 $00:29:03.660 \longrightarrow 00:29:05.756$ expectation of reward value.

NOTE Confidence: 0.850628229285714

 $00:29:05.756 \longrightarrow 00:29:08.376$ So each time the animal,

NOTE Confidence: 0.850628229285714

 $00:29:08.380 \longrightarrow 00:29:10.000$ here's the queue and is going

NOTE Confidence: 0.850628229285714

 $00:29:10.000 \longrightarrow 00:29:11.820$ to the port to get reward,

NOTE Confidence: 0.850628229285714

00:29:11.820 --> 00:29:13.710 it doesn't know which rewards coming,

NOTE Confidence: 0.850628229285714

 $00:29:13.710 \longrightarrow 00:29:16.566$ so it would have the same

NOTE Confidence: 0.850628229285714

 $00:29:16.566 \longrightarrow 00:29:17.994$ average reward expectation.

NOTE Confidence: 0.850628229285714

 $00:29:18.000 \longrightarrow 00:29:20.992$ And then the animal might receive a reward

NOTE Confidence: 0.850628229285714

 $00{:}29{:}20.992 \dashrightarrow 00{:}29{:}23.688$ better than average worse than average,

NOTE Confidence: 0.850628229285714 00:29:23.690 --> 00:29:24.380 you know, NOTE Confidence: 0.850628229285714

 $00:29:24.380 \longrightarrow 00:29:26.105$ just slightly better than average.

NOTE Confidence: 0.850628229285714

 $00{:}29{:}26.110 \dashrightarrow 00{:}29{:}28.707$ So this might also map onto what

NOTE Confidence: 0.850628229285714

 $00:29:28.707 \longrightarrow 00:29:31.207$ you might predict you would see

NOTE Confidence: 0.850628229285714

 $00:29:31.207 \longrightarrow 00:29:32.947$ with an expectation signal.

 $00:29:32.950 \longrightarrow 00:29:34.973$ So we were very interested in trying

NOTE Confidence: 0.850628229285714

 $00:29:34.973 \longrightarrow 00:29:37.464$ to figure out how could we tell the

NOTE Confidence: 0.850628229285714

 $00:29:37.464 \longrightarrow 00:29:39.426$ difference between a signal that might

NOTE Confidence: 0.850628229285714

 $00:29:39.426 \longrightarrow 00:29:41.823$ tell us something about if the animals

NOTE Confidence: 0.850628229285714

 $00:29:41.823 \longrightarrow 00:29:45.274$ using it to to read out violations of

NOTE Confidence: 0.850628229285714

 $00:29:45.274 \longrightarrow 00:29:48.634$ expectations or alterations of expectation.

NOTE Confidence: 0.850628229285714

 $00:29:48.640 \longrightarrow 00:29:50.481$ Or is the animal just using the

NOTE Confidence: 0.850628229285714

 $00:29:50.481 \longrightarrow 00:29:51.500$ signal to read up?

NOTE Confidence: 0.850628229285714

 $00:29:51.500 \longrightarrow 00:29:51.734 \text{ Yes},$

NOTE Confidence: 0.850628229285714

 $00:29:51.734 \longrightarrow 00:29:53.138$ this is what I like best.

NOTE Confidence: 0.850628229285714

 $00:29:53.140 \longrightarrow 00:29:54.310$ This is what I like worse.

NOTE Confidence: 0.89675105

 $00:29:57.230 \longrightarrow 00:29:59.414$ And this is basically a repeat

NOTE Confidence: 0.89675105

00:29:59.414 --> 00:30:01.769 of what I have just said.

NOTE Confidence: 0.89675105

 $00{:}30{:}01.770 \dashrightarrow 00{:}30{:}04.234$ So the the way in which David decided

NOTE Confidence: 0.89675105

 $00:30:04.234 \longrightarrow 00:30:06.356$ to tackle this was to collaborate

 $00:30:06.356 \longrightarrow 00:30:08.516$ with the lab of Jeremiah Cohen,

NOTE Confidence: 0.89675105

00:30:08.520 --> 00:30:10.540 also at Johns Hopkins University,

NOTE Confidence: 0.89675105

 $00:30:10.540 \longrightarrow 00:30:13.636$ and his then MD PhD student Bill Albari.

NOTE Confidence: 0.89675105

00:30:13.640 --> 00:30:16.405 And so Jeremiah and Bella had been

NOTE Confidence: 0.89675105

00:30:16.405 --> 00:30:19.120 using quantitative models to try

NOTE Confidence: 0.89675105

 $00:30:19.120 \longrightarrow 00:30:21.640$ to explain the activity of neurons

NOTE Confidence: 0.89675105

 $00:30:21.640 \longrightarrow 00:30:24.080$ and what what they care about.

NOTE Confidence: 0.89675105

 $00:30:24.080 \longrightarrow 00:30:26.429$ And so in this way you might use a

NOTE Confidence: 0.89675105

 $00:30:26.429 \longrightarrow 00:30:28.000$ quantitative model and try to fit

NOTE Confidence: 0.89675105

 $00:30:28.000 \longrightarrow 00:30:29.573$ the firing rate of given neurons

NOTE Confidence: 0.89675105

 $00{:}30{:}29.573 \dashrightarrow 00{:}30{:}31.693$ to a spects of your model to try to.

NOTE Confidence: 0.89675105

 $00:30:31.700 \longrightarrow 00:30:34.976$ Understand what those neurons are encoding,

NOTE Confidence: 0.89675105

 $00{:}30{:}34.980 \dashrightarrow 00{:}30{:}37.572$ and so this is what Belal and David

NOTE Confidence: 0.89675105

 $00:30:37.572 \longrightarrow 00:30:39.342$ together in collaboration did to

NOTE Confidence: 0.89675105

 $00:30:39.342 \longrightarrow 00:30:41.799$ ask if there was any impact of

NOTE Confidence: 0.89675105

 $00{:}30{:}41.873 \dashrightarrow 00{:}30{:}44.309$ expectations on firing at the time

 $00:30:44.309 \longrightarrow 00:30:46.606$ the animals drinking the reward and

NOTE Confidence: 0.89675105

 $00:30:46.606 \longrightarrow 00:30:49.270$ the way they did this was to look

NOTE Confidence: 0.89675105

 $00:30:49.346 \longrightarrow 00:30:52.272$ at the to the canonical Rescorla

NOTE Confidence: 0.89675105

 $00:30:52.272 \longrightarrow 00:30:56.274$ Wagner model that that tells us how

NOTE Confidence: 0.89675105

 $00:30:56.274 \longrightarrow 00:30:58.954$ predictions are updated by experience.

NOTE Confidence: 0.89675105

 $00:30:58.960 \longrightarrow 00:31:01.662$ And so this is the reward prediction

NOTE Confidence: 0.89675105

00:31:01.662 --> 00:31:03.924 error framework that many of you

NOTE Confidence: 0.89675105

 $00:31:03.924 \longrightarrow 00:31:05.994$ are familiar with and in this

NOTE Confidence: 0.89675105

 $00:31:05.994 \longrightarrow 00:31:08.195$ framework the expected value and

NOTE Confidence: 0.89675105

 $00:31:08.195 \longrightarrow 00:31:11.020$ animal holds for upcoming reward.

NOTE Confidence: 0.89675105

 $00:31:11.020 \longrightarrow 00:31:12.607$ Is updated iteratively,

NOTE Confidence: 0.89675105

 $00:31:12.607 \longrightarrow 00:31:15.781$ so with every experience based on

NOTE Confidence: 0.89675105

00:31:15.781 --> 00:31:18.492 whether the rewarded receives mattress,

NOTE Confidence: 0.89675105

 $00:31:18.492 \longrightarrow 00:31:20.732$ that or is better, or is worse,

NOTE Confidence: 0.89675105

 $00:31:20.732 \longrightarrow 00:31:22.076$ and so that's where we have

 $00:31:22.076 \longrightarrow 00:31:23.556$ positive prediction error if it's

NOTE Confidence: 0.89675105

 $00{:}31{:}23.556 \dashrightarrow 00{:}31{:}25.014$ better than expected, no change.

NOTE Confidence: 0.89675105

 $00:31:25.014 \longrightarrow 00:31:26.676$ If it's the same as expected,

NOTE Confidence: 0.89675105

 $00:31:26.680 \longrightarrow 00:31:28.312$ negative prediction error is what you

NOTE Confidence: 0.89675105

00:31:28.312 --> 00:31:30.300 get is worse than what you thought,

NOTE Confidence: 0.89675105

 $00:31:30.300 \longrightarrow 00:31:34.367$ and through the use of this canonical.

NOTE Confidence: 0.89675105

 $00:31:34.370 \longrightarrow 00:31:37.260$ Way of explaining what the

NOTE Confidence: 0.89675105

00:31:37.260 --> 00:31:40.590 activity of a neuron might be,

NOTE Confidence: 0.89675105

 $00{:}31{:}40.590 \dashrightarrow 00{:}31{:}44.145$ we can compare that with a much simpler idea,

NOTE Confidence: 0.89675105

 $00:31:44.150 \longrightarrow 00:31:46.310$ which is that the readout at the time,

NOTE Confidence: 0.89675105

 $00:31:46.310 \longrightarrow 00:31:47.912$ the animals drinking that reward just

NOTE Confidence: 0.89675105

 $00:31:47.912 \longrightarrow 00:31:49.370$ reflects a difference in outcome.

NOTE Confidence: 0.89675105

00:31:49.370 --> 00:31:51.209 A binary difference,

NOTE Confidence: 0.89675105

00:31:51.209 --> 00:31:53.048 sucrose versus maltodextrin?

NOTE Confidence: 0.89675105

 $00:31:53.050 \longrightarrow 00:31:55.680$ Or is this bike activity that we see at the

NOTE Confidence: 0.89675105

00:31:55.750 --> 00:31:58.446 time of reward unrelated to either of these?

00:31:58.450 --> 00:31:58.848 Of course,

NOTE Confidence: 0.89675105

 $00{:}31{:}58.848 {\:\raisebox{--}{\text{--}}}{\:\raisebox{--}{\text{--}}} 00{:}32{:}00.440$ we already have an idea that many of

NOTE Confidence: 0.89675105

 $00:32:00.487 \longrightarrow 00:32:01.927$ the neurons do care about rewards,

NOTE Confidence: 0.89675105

 $00:32:01.930 \longrightarrow 00:32:04.600$ so we don't expect that that will be a.

NOTE Confidence: 0.89675105

 $00:32:04.600 \longrightarrow 00:32:06.980$ Huge contributor so you can take this

NOTE Confidence: 0.89675105

 $00:32:06.980 \longrightarrow 00:32:09.389$ bike activity of neurons through time.

NOTE Confidence: 0.89675105

 $00:32:09.390 \longrightarrow 00:32:10.401$ Trial by trial.

NOTE Confidence: 0.89675105

 $00:32:10.401 \dashrightarrow 00:32:13.160$ Look at how the neuron responds to the

NOTE Confidence: 0.89675105

 $00:32:13.160 \dashrightarrow 00:32:15.470$ reward and see if its activity matches

NOTE Confidence: 0.89675105

 $00:32:15.470 \longrightarrow 00:32:18.357$ just a real time difference in outcome.

NOTE Confidence: 0.89675105

 $00:32:18.360 \longrightarrow 00:32:20.453$ Or does it actually account for what

NOTE Confidence: 0.89675105

 $00{:}32{:}20.453 \dashrightarrow 00{:}32{:}22.314$ the animal received a trial before

NOTE Confidence: 0.89675105

 $00:32:22.314 \longrightarrow 00:32:23.819$ trial before the trial before,

NOTE Confidence: 0.89675105

 $00:32:23.820 \longrightarrow 00:32:25.836$ as in a Rescorla Wagner model,

NOTE Confidence: 0.89675105

 $00:32:25.840 \longrightarrow 00:32:28.416$ and I wouldn't be saying this if

 $00:32:28.416 \longrightarrow 00:32:30.636$ we hadn't indeed found a group

NOTE Confidence: 0.89675105

 $00:32:30.636 \longrightarrow 00:32:32.604$ of neurons that does care about

NOTE Confidence: 0.89675105

 $00:32:32.604 \longrightarrow 00:32:34.608$ what reward the animal received.

NOTE Confidence: 0.89675105

 $00:32:34.610 \longrightarrow 00:32:36.782$ The trial before the trial that

NOTE Confidence: 0.89675105

 $00:32:36.782 \longrightarrow 00:32:38.230$ they're experiencing that reward.

NOTE Confidence: 0.89675105

 $00:32:38.230 \longrightarrow 00:32:39.220$ In other words,

NOTE Confidence: 0.89675105

 $00:32:39.220 \longrightarrow 00:32:40.870$ there's an impact of experience.

NOTE Confidence: 0.89675105

 $00:32:40.870 \longrightarrow 00:32:43.299$ So in about 20% of these neurons

NOTE Confidence: 0.89675105

 $00{:}32{:}43.299 \dashrightarrow 00{:}32{:}46.165$ that fire at the time of reward

NOTE Confidence: 0.89675105

00:32:46.165 --> 00:32:48.350 showed an impact of expectation,

NOTE Confidence: 0.89675105

 $00:32:48.350 \longrightarrow 00:32:50.950$ another slightly more than 20%,

NOTE Confidence: 0.89675105

 $00{:}32{:}50.950 \dashrightarrow 00{:}32{:}53.008$ were just encoding the current outcome.

NOTE Confidence: 0.89675105

 $00:32:53.010 \longrightarrow 00:32:54.690$ This ones better, that was worse,

NOTE Confidence: 0.89675105

 $00:32:54.690 \longrightarrow 00:32:56.610$ and that was relatively stable.

NOTE Confidence: 0.89675105

00:32:56.610 --> 00:32:58.794 And then there were neurons that didn't

NOTE Confidence: 0.89675105

 $00:32:58.794 \longrightarrow 00:33:00.668$ care about either of those things,

 $00{:}33{:}00.670 \dashrightarrow 00{:}33{:}03.078$ and you can now look at the neural

NOTE Confidence: 0.9162103975

 $00:33:03.078 \longrightarrow 00:33:04.799$ activity of these different.

NOTE Confidence: 0.9162103975

 $00:33:04.800 \longrightarrow 00:33:07.012$ Classes we now divided up our neurons

NOTE Confidence: 0.9162103975

 $00:33:07.012 \longrightarrow 00:33:09.092$ that respond to rewarding to these

NOTE Confidence: 0.9162103975

 $00{:}33{:}09.092 \dashrightarrow 00{:}33{:}11.628$ two classes and get a nice feel for

NOTE Confidence: 0.9162103975

00:33:11.628 --> 00:33:13.540 what this actually might look like.

NOTE Confidence: 0.9162103975

 $00:33:13.540 \longrightarrow 00:33:15.808$ So here are the neurons that were the reward

NOTE Confidence: 0.9162103975

 $00:33:15.808 \longrightarrow 00:33:17.528$ prediction error neurons they cared about.

NOTE Confidence: 0.9162103975

 $00{:}33{:}17.530 \dashrightarrow 00{:}33{:}19.274$ What happened trial before

NOTE Confidence: 0.9162103975

 $00:33:19.274 \longrightarrow 00:33:21.454$ divide it up just into.

NOTE Confidence: 0.9162103975

00:33:21.460 --> 00:33:23.680 The simplest kind of way of

NOTE Confidence: 0.9162103975

 $00:33:23.680 \longrightarrow 00:33:25.160$ thinking about this trials,

NOTE Confidence: 0.9162103975

 $00{:}33{:}25.160 \dashrightarrow 00{:}33{:}27.930$ in which animals get sucrose.

NOTE Confidence: 0.9162103975

 $00{:}33{:}27.930 \dashrightarrow 00{:}33{:}29.994$ After a trial when they got

NOTE Confidence: 0.9162103975

 $00:33:29.994 \longrightarrow 00:33:32.000$ maltodextrin so better than expected,

 $00:33:32.000 \longrightarrow 00:33:34.340$ that's this very tall yellow peak.

NOTE Confidence: 0.9162103975

 $00{:}33{:}34.340 \dashrightarrow 00{:}33{:}36.368$ Trials when animals got

NOTE Confidence: 0.9162103975

 $00:33:36.368 \longrightarrow 00:33:37.889$ sucrose after sucrose.

NOTE Confidence: 0.9162103975

 $00:33:37.890 \longrightarrow 00:33:39.930$ Trials when animals got multi dextrin.

NOTE Confidence: 0.9162103975

 $00:33:39.930 \longrightarrow 00:33:42.175$ After maltodextrin and trials when

NOTE Confidence: 0.9162103975

 $00:33:42.175 \longrightarrow 00:33:44.420$ animals got maltodextrin after sucrose

NOTE Confidence: 0.9162103975

 $00:33:44.482 \longrightarrow 00:33:46.794$ so much worse than they thought and you

NOTE Confidence: 0.9162103975

 $00:33:46.794 \longrightarrow 00:33:49.280$ can see this big modulation of firing.

NOTE Confidence: 0.9162103975

 $00:33:49.280 \longrightarrow 00:33:51.230$ That depends on what just happened.

NOTE Confidence: 0.9162103975

 $00:33:51.230 \longrightarrow 00:33:52.834$ So that matches this.

NOTE Confidence: 0.9162103975

 $00:33:52.834 \longrightarrow 00:33:54.037$ This quantitative assessment

NOTE Confidence: 0.9162103975

 $00:33:54.037 \longrightarrow 00:33:55.790$ and the current outcome.

NOTE Confidence: 0.9162103975

 $00:33:55.790 \longrightarrow 00:33:58.891$ Neurons show much less or no modulation

NOTE Confidence: 0.9162103975

 $00{:}33{:}58.891 \dashrightarrow 00{:}34{:}01.685$ around expectation and is quite flat

NOTE Confidence: 0.9162103975

00:34:01.685 --> 00:34:04.005 for the unmodulated neurons obviously,

NOTE Confidence: 0.9162103975

 $00{:}34{:}04.010 \dashrightarrow 00{:}34{:}06.106$ and we can do this same analysis in

 $00{:}34{:}06.106 \dashrightarrow 00{:}34{:}08.098$ R3 reward tasks and find the same.

NOTE Confidence: 0.9162103975

 $00:34:08.100 \longrightarrow 00:34:10.330$ Outcome where there's a portion

NOTE Confidence: 0.9162103975

 $00:34:10.330 \longrightarrow 00:34:12.780$ of neurons that encode some kind

NOTE Confidence: 0.9162103975

 $00{:}34{:}12.780 \dashrightarrow 00{:}34{:}14.430$ of expectation signal looks like

NOTE Confidence: 0.9162103975

 $00:34:14.430 \longrightarrow 00:34:15.800$ a reward prediction error,

NOTE Confidence: 0.9162103975

 $00:34:15.800 \longrightarrow 00:34:17.534$ and even more neurons do this

NOTE Confidence: 0.9162103975

 $00:34:17.534 \longrightarrow 00:34:19.553$ when you have this larger dynamic

NOTE Confidence: 0.9162103975

 $00:34:19.553 \longrightarrow 00:34:21.157$ range across the rewards,

NOTE Confidence: 0.9162103975

 $00:34:21.160 \longrightarrow 00:34:22.288$ the animals experiencing,

NOTE Confidence: 0.9162103975

 $00:34:22.288 \longrightarrow 00:34:24.168$ which is kind of interesting,

NOTE Confidence: 0.9162103975

 $00:34:24.170 \longrightarrow 00:34:26.746$ we can again get an intuitive feel

NOTE Confidence: 0.9162103975

 $00:34:26.746 \longrightarrow 00:34:29.616$ for how this maps onto neural activity

NOTE Confidence: 0.9162103975

 $00{:}34{:}29.616 \dashrightarrow 00{:}34{:}32.860$ by looking at subsets of the neurons.

NOTE Confidence: 0.9162103975

 $00:34:32.860 \longrightarrow 00:34:33.721$ In this case,

NOTE Confidence: 0.9162103975

 $00:34:33.721 \longrightarrow 00:34:36.171$ if we just look at the neurons that

 $00:34:36.171 \longrightarrow 00:34:38.409$ are seem to encode and expectation.

NOTE Confidence: 0.9162103975

 $00{:}34{:}38.410 \dashrightarrow 00{:}34{:}40.895$ Signal and categorize them based

NOTE Confidence: 0.9162103975

 $00:34:40.895 \longrightarrow 00:34:42.883$ on the calculated error.

NOTE Confidence: 0.9162103975

00:34:42.890 --> 00:34:44.078 Was it very positive?

NOTE Confidence: 0.9162103975

 $00:34:44.078 \longrightarrow 00:34:46.579$ Was there no error signal on that trial?

NOTE Confidence: 0.9162103975

 $00:34:46.580 \longrightarrow 00:34:48.729$ Was there a negative error signal on

NOTE Confidence: 0.9162103975

 $00:34:48.729 \longrightarrow 00:34:52.026$ that trial and use eight categories for that?

NOTE Confidence: 0.9162103975

 $00:34:52.030 \longrightarrow 00:34:54.290$ We can see this beautiful

NOTE Confidence: 0.9162103975

 $00:34:54.290 \longrightarrow 00:34:56.604$ distribution of signals along the

NOTE Confidence: 0.9162103975

00:34:56.604 --> 00:34:58.989 most positive reward prediction error,

NOTE Confidence: 0.9162103975

 $00{:}34{:}58.990 \dashrightarrow 00{:}35{:}01.450$ little error and negative error.

NOTE Confidence: 0.9162103975

 $00:35:01.450 \longrightarrow 00:35:03.714$ So this gives us a sort of intuitive

NOTE Confidence: 0.9162103975

 $00:35:03.714 \longrightarrow 00:35:05.779$ way to think about how firing

NOTE Confidence: 0.9162103975

 $00{:}35{:}05.779 \dashrightarrow 00{:}35{:}07.915$ happening at the time of reward

NOTE Confidence: 0.9162103975

 $00:35:07.987 \longrightarrow 00:35:09.767$ can actually be telling us.

NOTE Confidence: 0.9162103975

 $00:35:09.770 \longrightarrow 00:35:12.026$ Think about what the animal expects,

 $00:35:12.030 \longrightarrow 00:35:14.970$ not just what is this particular

NOTE Confidence: 0.9162103975

 $00:35:14.970 \longrightarrow 00:35:17.930$ reward as far as identity.

NOTE Confidence: 0.9162103975

 $00:35:17.930 \longrightarrow 00:35:20.121$ So this this was really exciting to

NOTE Confidence: 0.9162103975

 $00:35:20.121 \longrightarrow 00:35:22.979$ us to find this kind of signal and BP.

NOTE Confidence: 0.9162103975

 $00:35:22.980 \dashrightarrow 00:35:24.726$ We're very used to thinking about

NOTE Confidence: 0.9162103975

 $00:35:24.726 \longrightarrow 00:35:25.890$ these kinds of expectations.

NOTE Confidence: 0.9162103975

00:35:25.890 --> 00:35:27.210 Signals in dopamine neurons,

NOTE Confidence: 0.9162103975

 $00:35:27.210 \longrightarrow 00:35:29.190$ and we hadn't been expecting to

NOTE Confidence: 0.9162103975

 $00:35:29.248 \longrightarrow 00:35:31.173$ see this kind of thing in ventral

NOTE Confidence: 0.9162103975

 $00:35:31.173 \longrightarrow 00:35:31.998$ pallidum at all,

NOTE Confidence: 0.9162103975

 $00{:}35{:}32.000 \dashrightarrow 00{:}35{:}33.729$ which we thought would be more just

NOTE Confidence: 0.9162103975

00:35:33.729 --> 00:35:35.730 a basic readout. This is good.

NOTE Confidence: 0.9162103975

 $00{:}35{:}35.730 \dashrightarrow 00{:}35{:}38.418$ This is bad in real time.

NOTE Confidence: 0.9162103975

 $00:35:38.420 \longrightarrow 00:35:40.940$ So because we saw these signals that

NOTE Confidence: 0.9162103975

 $00:35:40.940 \longrightarrow 00:35:43.290$ match what are teaching signals,

 $00:35:43.290 \longrightarrow 00:35:45.786$ signals that are updated over time

NOTE Confidence: 0.9162103975

00:35:45.786 --> 00:35:48.010 and reflect what subjects expect,

NOTE Confidence: 0.9162103975

00:35:48.010 --> 00:35:50.062 David wondered if he could find

NOTE Confidence: 0.9162103975

 $00:35:50.062 \longrightarrow 00:35:52.843$ any way to see if the animal's

NOTE Confidence: 0.9162103975

 $00:35:52.843 \longrightarrow 00:35:55.385$ behavior changed based on what the

NOTE Confidence: 0.9162103975

 $00:35:55.385 \longrightarrow 00:35:58.640$ error signal was on a given trial.

NOTE Confidence: 0.9162103975

 $00{:}35{:}58.640 \dashrightarrow 00{:}36{:}00.944$ So we have to say that the procedure

NOTE Confidence: 0.9162103975

 $00:36:00.944 \longrightarrow 00:36:02.792$ that he designed was not designed

NOTE Confidence: 0.9162103975

 $00:36:02.792 \longrightarrow 00:36:05.086$ to see a lot of rich behavior

NOTE Confidence: 0.9162103975

 $00{:}36{:}05.086 \dashrightarrow 00{:}36{:}07.600$ was designed to have the animals

NOTE Confidence: 0.9162103975

 $00:36:07.600 \longrightarrow 00:36:08.857$ behavior very similar.

NOTE Confidence: 0.871961503571429

00:36:08.860 --> 00:36:11.058 Each trial. Indeed, you saw that when

NOTE Confidence: 0.871961503571429

00:36:11.058 --> 00:36:13.248 we were looking at the licking rate,

NOTE Confidence: 0.871961503571429

00:36:13.250 --> 00:36:15.854 but David did still videotape the animal,

NOTE Confidence: 0.871961503571429

 $00:36:15.860 \longrightarrow 00:36:18.996$ so he went back and analyzed their behavior.

NOTE Confidence: 0.871961503571429

 $00:36:19.000 \longrightarrow 00:36:20.757$ This is what the Chamber looks like.

00:36:20.760 --> 00:36:22.152 A typical rat chamber,

NOTE Confidence: 0.871961503571429

 $00{:}36{:}22.152 \dashrightarrow 00{:}36{:}25.080$ and where the animals can enter reward port.

NOTE Confidence: 0.871961503571429

 $00:36:25.080 \longrightarrow 00:36:27.088$ You know when the queues on and drink

NOTE Confidence: 0.871961503571429

 $00:36:27.088 \longrightarrow 00:36:28.907$ the reward and there's an interest.

NOTE Confidence: 0.871961503571429

 $00:36:28.910 \longrightarrow 00:36:30.114$ Inter trial interval you

NOTE Confidence: 0.871961503571429

 $00:36:30.114 \longrightarrow 00:36:31.318$ know they wander around.

NOTE Confidence: 0.871961503571429

00:36:31.320 --> 00:36:32.616 They might do. A little grooming,

NOTE Confidence: 0.871961503571429

 $00:36:32.620 \longrightarrow 00:36:33.922$ might hang out by the port

NOTE Confidence: 0.871961503571429

 $00:36:33.922 \longrightarrow 00:36:35.120$ waiting for the next month.

NOTE Confidence: 0.871961503571429

 $00:36:35.120 \longrightarrow 00:36:37.388$ Typical behavior that the rat behaviors

NOTE Confidence: 0.871961503571429

00:36:37.388 --> 00:36:39.933 among us are used to seeing what

NOTE Confidence: 0.871961503571429

 $00:36:39.933 \longrightarrow 00:36:42.166$ David found when he looked at the

NOTE Confidence: 0.871961503571429

 $00{:}36{:}42.234 \dashrightarrow 00{:}36{:}44.789$ behavior in detail that on trials when

NOTE Confidence: 0.871961503571429

00:36:44.789 --> 00:36:47.513 animals had just received the reward,

NOTE Confidence: 0.871961503571429

 $00:36:47.513 \longrightarrow 00:36:49.677$ they liked better sucrose.

 $00:36:49.680 \longrightarrow 00:36:50.616$ Right after that,

NOTE Confidence: 0.871961503571429

 $00:36:50.616 \longrightarrow 00:36:53.150$ the animal tended to hang around the port,

NOTE Confidence: 0.871961503571429

 $00:36:53.150 \longrightarrow 00:36:55.470$ waiting presumably for more sucrose.

NOTE Confidence: 0.871961503571429

 $00:36:55.470 \longrightarrow 00:36:57.668$ If the animal had just received maltodextrin,

NOTE Confidence: 0.871961503571429

 $00:36:57.670 \longrightarrow 00:37:00.046$ they tended to wander off more.

NOTE Confidence: 0.871961503571429

 $00:37:00.050 \longrightarrow 00:37:01.710$ It's not a huge effect.

NOTE Confidence: 0.871961503571429

 $00:37:01.710 \longrightarrow 00:37:02.870$ We can look at this.

NOTE Confidence: 0.871961503571429

 $00:37:02.870 \longrightarrow 00:37:04.758$ This is sort of a way to map

NOTE Confidence: 0.871961503571429

 $00:37:04.758 \longrightarrow 00:37:05.750$ individual animal behavior.

NOTE Confidence: 0.871961503571429

 $00:37:05.750 \longrightarrow 00:37:06.995$ Looking at a cartoon of

NOTE Confidence: 0.871961503571429

 $00{:}37{:}06.995 \dashrightarrow 00{:}37{:}08.240$ the square of the chamber,

NOTE Confidence: 0.871961503571429

 $00:37:08.240 \longrightarrow 00:37:09.710$ it's not a huge effect,

NOTE Confidence: 0.871961503571429

 $00:37:09.710 \longrightarrow 00:37:12.118$ but you see more color and more black.

NOTE Confidence: 0.871961503571429

 $00:37:12.120 \longrightarrow 00:37:14.954 \text{ X}$ is away from the port when

NOTE Confidence: 0.871961503571429

00:37:14.954 --> 00:37:17.130 it's post maltodextrin sucrose,

NOTE Confidence: 0.871961503571429

 $00:37:17.130 \longrightarrow 00:37:19.780$ and this is statistically significant.

 $00:37:19.780 \longrightarrow 00:37:21.474$ We see the same pattern when we

NOTE Confidence: 0.871961503571429

 $00:37:21.474 \longrightarrow 00:37:23.160$ have the three rewards together.

NOTE Confidence: 0.871961503571429

00:37:23.160 --> 00:37:24.222 After receiving water,

NOTE Confidence: 0.871961503571429

 $00:37:24.222 \longrightarrow 00:37:25.992$ they're more likely to be

NOTE Confidence: 0.871961503571429

 $00:37:25.992 \longrightarrow 00:37:27.280$ further from the port,

NOTE Confidence: 0.871961503571429

 $00:37:27.280 \longrightarrow 00:37:30.654$ so it's very simple measure of how

NOTE Confidence: 0.871961503571429

 $00:37:30.654 \longrightarrow 00:37:32.942$ their future behavior is impacted

NOTE Confidence: 0.871961503571429

 $00:37:32.942 \longrightarrow 00:37:35.580$ by the the the validation

NOTE Confidence: 0.871961503571429

 $00:37:35.580 \longrightarrow 00:37:38.080$ of their expectation or getting

NOTE Confidence: 0.871961503571429

 $00:37:38.080 \longrightarrow 00:37:40.619$ something less than they expected.

NOTE Confidence: 0.871961503571429

 $00:37:40.620 \longrightarrow 00:37:42.003$ Simple small effect,

NOTE Confidence: 0.871961503571429

 $00:37:42.003 \longrightarrow 00:37:45.230$ but can we manipulate it by changing

NOTE Confidence: 0.871961503571429

 $00:37:45.312 \longrightarrow 00:37:47.898$ these neurons and how they fire?

NOTE Confidence: 0.871961503571429

 $00{:}37{:}47.900 \dashrightarrow 00{:}37{:}50.140$ So this was the setting for an

NOTE Confidence: 0.871961503571429

 $00:37:50.140 \longrightarrow 00:37:51.500$ optogenetic experiment at David,

 $00:37:51.500 \longrightarrow 00:37:52.840$ then performed getting together

NOTE Confidence: 0.871961503571429

 $00{:}37{:}52.840 \dashrightarrow 00{:}37{:}55.157$ with Tabatha Kim and Kurt Fraser to

NOTE Confidence: 0.871961503571429

 $00{:}37{:}55.157 \dashrightarrow 00{:}37{:}56.914$ graduate students in the lab at that

NOTE Confidence: 0.871961503571429

 $00:37:56.914 \longrightarrow 00:37:58.624$ time had now moved on to Charles

NOTE Confidence: 0.871961503571429

 $00:37:58.624 \longrightarrow 00:38:00.725$ River and to post up with stuff on

NOTE Confidence: 0.871961503571429

 $00:38:00.725 \longrightarrow 00:38:02.930$ the mouth and so they simply asked

NOTE Confidence: 0.871961503571429

 $00:38:03.001 \longrightarrow 00:38:05.166$ if they express general adoption.

NOTE Confidence: 0.871961503571429

 $00{:}38{:}05.170 \dashrightarrow 00{:}38{:}06.990$ Excited Tori option in neurons

NOTE Confidence: 0.871961503571429

 $00{:}38{:}06.990 \dashrightarrow 00{:}38{:}09.242$ in the ventral pallidum and then

NOTE Confidence: 0.871961503571429

 $00:38:09.242 \longrightarrow 00:38:10.698$ make those neurons fire.

NOTE Confidence: 0.871961503571429

 $00{:}38{:}10.700 \dashrightarrow 00{:}38{:}13.116$ By shining light on them at the time,

NOTE Confidence: 0.871961503571429

 $00:38:13.120 \longrightarrow 00:38:14.792$ the animals ingesting reward,

NOTE Confidence: 0.871961503571429

 $00:38:14.792 \longrightarrow 00:38:16.882$ can they impact the behavior

NOTE Confidence: 0.871961503571429

 $00:38:16.882 \longrightarrow 00:38:19.070$ after the animal got the reward?

NOTE Confidence: 0.871961503571429 00:38:19.070 --> 00:38:19.346 How? NOTE Confidence: 0.871961503571429

 $00{:}38{:}19.346 \dashrightarrow 00{:}38{:}21.002$ How much the animals likely to

 $00:38:21.002 \longrightarrow 00:38:22.419$ hang out by the port?

NOTE Confidence: 0.871961503571429

 $00{:}38{:}22.420 \dashrightarrow 00{:}38{:}24.534$ That's so if you excite the neurons,

NOTE Confidence: 0.871961503571429

 $00{:}38{:}24.540 \dashrightarrow 00{:}38{:}25.896$ you'd expect to see the animals

NOTE Confidence: 0.871961503571429

 $00:38:25.896 \longrightarrow 00:38:27.429$ hang close to the port right

NOTE Confidence: 0.871961503571429

 $00:38:27.429 \longrightarrow 00:38:28.879$ after you've excited the neurons.

NOTE Confidence: 0.871961503571429

00:38:28.880 --> 00:38:29.370 Alternatively,

NOTE Confidence: 0.871961503571429

00:38:29.370 --> 00:38:32.310 if you express an inhibitory option

NOTE Confidence: 0.871961503571429

 $00{:}38{:}32.310 \dashrightarrow 00{:}38{:}34.846$ in ventral pallidum so that you

NOTE Confidence: 0.871961503571429

 $00{:}38{:}34.846 \dashrightarrow 00{:}38{:}37.149$ can inhibit BP neuron firing at the

NOTE Confidence: 0.871961503571429

 $00:38:37.226 \longrightarrow 00:38:39.536$ time they're drinking the reward,

NOTE Confidence: 0.871961503571429

 $00:38:39.540 \longrightarrow 00:38:41.012$ you'd expect to make.

NOTE Confidence: 0.871961503571429

00:38:41.012 --> 00:38:43.649 The animal more likely to be away

NOTE Confidence: 0.871961503571429

 $00{:}38{:}43.649 \dashrightarrow 00{:}38{:}46.380$ from the court after that sort

NOTE Confidence: 0.871961503571429

00:38:46.380 --> 00:38:48.620 of inhibition of BP,

NOTE Confidence: 0.871961503571429

 $00:38:48.620 \longrightarrow 00:38:50.336$ and so that's the kind of

 $00:38:50.336 \longrightarrow 00:38:51.480$ experiment that they designed.

NOTE Confidence: 0.871961503571429

 $00{:}38{:}51.480 \dashrightarrow 00{:}38{:}53.832$ This is just a cartoon by lateral

NOTE Confidence: 0.871961503571429

 $00{:}38{:}53.832 \dashrightarrow 00{:}38{:}56.176$ inhibition of BP neurons is going to

NOTE Confidence: 0.871961503571429

00:38:56.176 --> 00:38:58.680 occur at the time of reward delivery

NOTE Confidence: 0.871961503571429

 $00:38:58.680 \longrightarrow 00:39:01.260$ or unilateral excitation of DP.

NOTE Confidence: 0.871961503571429

 $00:39:01.260 \longrightarrow 00:39:03.378$ Neurons will occur at the time

NOTE Confidence: 0.871961503571429

 $00:39:03.378 \longrightarrow 00:39:04.437$ of reward delivery.

NOTE Confidence: 0.905507280833333

 $00:39:04.440 \longrightarrow 00:39:05.900$ So for this experiment the

NOTE Confidence: 0.905507280833333

 $00:39:05.900 \longrightarrow 00:39:08.099$ reward is the same in all trials,

NOTE Confidence: 0.905507280833333

 $00:39:08.100 \longrightarrow 00:39:11.150$ it's sucrose, but the optogenetic.

NOTE Confidence: 0.905507280833333

 $00{:}39{:}11.150 \dashrightarrow 00{:}39{:}12.398$ Activation or inhibition will

NOTE Confidence: 0.905507280833333

 $00:39:12.398 \longrightarrow 00:39:14.730$ just occur on half of the trials,

NOTE Confidence: 0.905507280833333

 $00:39:14.730 \longrightarrow 00:39:16.906$ so that lets you see if your change

NOTE Confidence: 0.905507280833333

 $00:39:16.906 \longrightarrow 00:39:19.213$ in neural activity is affecting the

NOTE Confidence: 0.905507280833333

 $00:39:19.213 \longrightarrow 00:39:21.333$ animals behavior independent of the

NOTE Confidence: 0.905507280833333

 $00{:}39{:}21.333 \dashrightarrow 00{:}39{:}23.290$ specific taste of the reward, etc.

 $00:39:23.290 \longrightarrow 00:39:25.110$ So you hold the those aspects of

NOTE Confidence: 0.905507280833333

 $00{:}39{:}25.110 \dashrightarrow 00{:}39{:}26.838$ the reward constant and see if

NOTE Confidence: 0.905507280833333

 $00{:}39{:}26.838 \dashrightarrow 00{:}39{:}28.560$ you're turning up or turning down

NOTE Confidence: 0.905507280833333

 $00:39:28.621 \longrightarrow 00:39:30.191$ of the neural activity impacts

NOTE Confidence: 0.905507280833333

00:39:30.191 --> 00:39:32.123 their behavior as you would expect,

NOTE Confidence: 0.905507280833333

 $00:39:32.123 \longrightarrow 00:39:34.241$ and that is exactly what they

NOTE Confidence: 0.905507280833333

 $00:39:34.241 \longrightarrow 00:39:35.890$ observed in this Chamber.

NOTE Confidence: 0.905507280833333

 $00:39:35.890 \longrightarrow 00:39:38.410$ That reward port is here on the right side,

NOTE Confidence: 0.905507280833333

00:39:38.410 --> 00:39:41.315 on trials in which BP was activated.

NOTE Confidence: 0.905507280833333

 $00:39:41.320 \longrightarrow 00:39:42.964$ The subject tends to hang out

NOTE Confidence: 0.905507280833333

 $00:39:42.964 \longrightarrow 00:39:44.060$ closer to the port,

NOTE Confidence: 0.905507280833333

 $00:39:44.060 \longrightarrow 00:39:46.692$ so a lower value here than on

NOTE Confidence: 0.905507280833333

 $00{:}39{:}46.692 \dashrightarrow 00{:}39{:}48.970$ trials where the subject was not

NOTE Confidence: 0.905507280833333

00:39:48.970 --> 00:39:50.895 did not receive VP activation,

NOTE Confidence: 0.905507280833333

 $00:39:50.900 \longrightarrow 00:39:53.054$ so it's a within subject within

 $00:39:53.054 \longrightarrow 00:39:54.974$ session comparison and we see

NOTE Confidence: 0.905507280833333

 $00:39:54.974 \longrightarrow 00:39:56.638$ the opposite with inhibition.

NOTE Confidence: 0.905507280833333

00:39:56.640 --> 00:39:59.024 They tend to be further away if you

NOTE Confidence: 0.905507280833333

00:39:59.024 --> 00:40:01.448 inhibit while they're consuming the reward,

NOTE Confidence: 0.905507280833333

 $00{:}40{:}01.450 \dashrightarrow 00{:}40{:}04.551$ so this these are reliable effects but

NOTE Confidence: 0.905507280833333

00:40:04.551 --> 00:40:07.219 granted relatively small in this procedure,

NOTE Confidence: 0.905507280833333

 $00:40:07.220 \longrightarrow 00:40:09.040$ not really used to.

NOTE Confidence: 0.905507280833333

00:40:09.040 --> 00:40:11.770 Think about how this impacts decisions,

NOTE Confidence: 0.905507280833333

 $00:40:11.770 \longrightarrow 00:40:13.834$ but because we saw the signal

NOTE Confidence: 0.905507280833333

 $00:40:13.834 \longrightarrow 00:40:14.866$ in this behavior,

NOTE Confidence: 0.905507280833333

 $00{:}40{:}14.870 \dashrightarrow 00{:}40{:}17.118$ we wanted to see if we could find

NOTE Confidence: 0.905507280833333

 $00:40:17.118 \longrightarrow 00:40:19.408$ any evidence that an expectation

NOTE Confidence: 0.905507280833333

00:40:19.408 --> 00:40:21.130 reward prediction, like signal,

NOTE Confidence: 0.905507280833333

 $00:40:21.130 \longrightarrow 00:40:23.130$ impacted future behavior and this

NOTE Confidence: 0.905507280833333

 $00:40:23.130 \longrightarrow 00:40:25.248$ evidence was there and so that

NOTE Confidence: 0.905507280833333

 $00{:}40{:}25.248 \dashrightarrow 00{:}40{:}26.946$ was really exciting to us and

 $00:40:26.946 \longrightarrow 00:40:29.051$ lead the stage for our continued

NOTE Confidence: 0.905507280833333

 $00{:}40{:}29.051 \dashrightarrow 00{:}40{:}31.080$ experiment that I'll tell you about.

NOTE Confidence: 0.905507280833333

00:40:31.080 --> 00:40:33.240 Right now I'm going to take a quick

NOTE Confidence: 0.905507280833333

 $00:40:33.240 \longrightarrow 00:40:35.569$ interim summary and this also be a good time,

NOTE Confidence: 0.905507280833333 00:40:35.570 --> 00:40:37.770 if anyone. NOTE Confidence: 0.905507280833333

 $00:40:37.770 \longrightarrow 00:40:41.506$ Wants me to clarify something that I've said.

NOTE Confidence: 0.905507280833333

 $00:40:41.510 \longrightarrow 00:40:43.510$ So I wanted to just say from these.

NOTE Confidence: 0.905507280833333

 $00:40:43.510 \longrightarrow 00:40:45.729$ So far we've learned that the signal

NOTE Confidence: 0.905507280833333

 $00:40:45.729 \longrightarrow 00:40:48.315$ in VP that responds to reward is

NOTE Confidence: 0.905507280833333

00:40:48.315 --> 00:40:50.270 sensitive to pass reward history

NOTE Confidence: 0.905507280833333

 $00:40:50.270 \longrightarrow 00:40:52.689$ and can provide a reward prediction

NOTE Confidence: 0.905507280833333

 $00:40:52.689 \longrightarrow 00:40:55.290$ error signal to update the animals.

NOTE Confidence: 0.905507280833333

 $00{:}40{:}55.290 \dashrightarrow 00{:}40{:}57.850$ Expected value of reward.

NOTE Confidence: 0.905507280833333

00:40:57.850 --> 00:40:59.290 And So what we would like to know,

NOTE Confidence: 0.905507280833333

 $00:40:59.290 \longrightarrow 00:41:01.775$ of course, is, are these signals used?

00:41:01.780 --> 00:41:04.768 Do they interact with decision processes?

NOTE Confidence: 0.905507280833333

 $00{:}41{:}04.770 \dashrightarrow 00{:}41{:}06.828$ Can they impact the actions animals make?

NOTE Confidence: 0.905507280833333

 $00:41:06.830 \longrightarrow 00:41:08.630$ 'cause that's ultimately what we

NOTE Confidence: 0.905507280833333

00:41:08.630 --> 00:41:11.494 want to explain how our choices made,

NOTE Confidence: 0.905507280833333

 $00:41:11.494 \longrightarrow 00:41:14.831$ what's going on in the brain when

NOTE Confidence: 0.905507280833333

 $00:41:14.831 \longrightarrow 00:41:17.656$ the animal evaluates the options?

NOTE Confidence: 0.905507280833333

00:41:17.660 --> 00:41:19.130 Question yes please.

NOTE Confidence: 0.9494952525

00:41:20.330 --> 00:41:22.230 That's really beautifully done.

NOTE Confidence: 0.9494952525

00:41:22.230 --> 00:41:24.453 I was wondering if you've

NOTE Confidence: 0.9494952525

 $00:41:24.453 \longrightarrow 00:41:26.198$ looked at what happens with.

NOTE Confidence: 0.9494952525

 $00{:}41{:}26.200 \dashrightarrow 00{:}41{:}28.660$ Obviously, this is a learning signal.

NOTE Confidence: 0.9494952525

 $00:41:28.660 \longrightarrow 00:41:31.760$ What if it's a pharmacologic reward that

NOTE Confidence: 0.9494952525

 $00:41:31.760 \longrightarrow 00:41:34.880$ is sensitive to to tolerance effects?

NOTE Confidence: 0.9494952525

00:41:34.880 --> 00:41:38.290 Or you know any sort of reduction?

NOTE Confidence: 0.9321093225

 $00:41:38.300 \longrightarrow 00:41:40.668$ What happens to these neurons and to behavior

NOTE Confidence: 0.849466579

 $00:41:40.680 \longrightarrow 00:41:42.160$ 'cause ya'll animal moved away

 $00:41:42.160 \longrightarrow 00:41:43.640$ when they were not getting,

NOTE Confidence: 0.849466579

00:41:43.640 --> 00:41:45.044 you know, beautiful extinction.

NOTE Confidence: 0.849466579

00:41:45.044 --> 00:41:47.150 But that is not obviously what

NOTE Confidence: 0.849466579

 $00:41:47.216 \longrightarrow 00:41:48.965$ we see when when people are

NOTE Confidence: 0.849466579

00:41:48.965 --> 00:41:50.390 beginning not even addicted.

NOTE Confidence: 0.849466579

 $00:41:50.390 \longrightarrow 00:41:52.720$ Just beginning to come

NOTE Confidence: 0.856978722

 $00:41:52.750 \longrightarrow 00:41:55.456$ to start to really like and

NOTE Confidence: 0.856978722

 $00:41:55.456 \longrightarrow 00:41:57.260$ escalate their their use.

NOTE Confidence: 0.856978722

00:41:57.260 --> 00:41:58.570 Yeah, that's a great question.

NOTE Confidence: 0.856978722

 $00:41:58.570 \longrightarrow 00:42:00.466$ So what we have not done yet is

NOTE Confidence: 0.856978722

00:42:00.466 --> 00:42:02.284 is used Ivy drug, for example,

NOTE Confidence: 0.856978722

 $00:42:02.284 \longrightarrow 00:42:04.853$ or even alcohol in this exact model.

NOTE Confidence: 0.856978722

 $00{:}42{:}04.860 --> 00{:}42{:}06.972$ And we we we need to do that

NOTE Confidence: 0.856978722

 $00:42:06.972 \longrightarrow 00:42:09.080$ because here we are beginning to

NOTE Confidence: 0.856978722

 $00:42:09.080 \longrightarrow 00:42:10.975$ define for ourselves what these

 $00:42:10.975 \longrightarrow 00:42:13.227$ neurons are doing to natural reward.

NOTE Confidence: 0.856978722

 $00{:}42{:}13.230 \dashrightarrow 00{:}42{:}14.815$ And when natural reward choices

NOTE Confidence: 0.856978722

 $00:42:14.815 \longrightarrow 00:42:15.449$ are occurring.

NOTE Confidence: 0.856978722

 $00:42:15.450 \longrightarrow 00:42:18.280$ But the critical question is how are

NOTE Confidence: 0.856978722

 $00:42:18.280 \longrightarrow 00:42:20.611$ these processes altered when that reward?

NOTE Confidence: 0.856978722

 $00{:}42{:}20.611 \longrightarrow 00{:}42{:}22.830$ Is a drug reward that has pharmacological

NOTE Confidence: 0.856978722

 $00:42:22.883 \longrightarrow 00:42:24.573$ properties that are quite different

NOTE Confidence: 0.856978722

 $00:42:24.573 \longrightarrow 00:42:26.590$ and there is important data from,

NOTE Confidence: 0.856978722

00:42:26.590 --> 00:42:27.360 for example,

NOTE Confidence: 0.856978722

00:42:27.360 --> 00:42:29.285 Megan Creed and Christian looser,

NOTE Confidence: 0.856978722

00:42:29.290 --> 00:42:33.292 showing that drugs like cocaine change

NOTE Confidence: 0.856978722

00:42:33.292 --> 00:42:36.082 synaptic efficacy between, for example,

NOTE Confidence: 0.856978722

 $00:42:36.082 \longrightarrow 00:42:38.986$ the accompagnes and ventral pallidal neurons.

NOTE Confidence: 0.856978722

 $00:42:38.990 \longrightarrow 00:42:41.524$ So there are chronic effects of drugs

NOTE Confidence: 0.856978722

 $00:42:41.524 \longrightarrow 00:42:44.472$ on the way that these neurons should

NOTE Confidence: 0.856978722

 $00:42:44.472 \longrightarrow 00:42:47.142$ be activated and should be firing

 $00:42:47.150 \longrightarrow 00:42:48.812$ and and so finding that intersection

NOTE Confidence: 0.856978722

 $00:42:48.812 \longrightarrow 00:42:51.146$ and studying that was, is it?

NOTE Confidence: 0.856978722 00:42:51.146 --> 00:42:51.759 Ago.

NOTE Confidence: 0.935188366842105

00:42:55.170 --> 00:42:58.720 OK so I'm gonna go on and tell you about

NOTE Confidence: 0.935188366842105

 $00:42:58.808 \longrightarrow 00:43:01.863$ the next reward behavioral procedure

NOTE Confidence: 0.935188366842105

 $00:43:01.863 \longrightarrow 00:43:05.222$ where David extended this work to try

NOTE Confidence: 0.935188366842105

 $00:43:05.222 \longrightarrow 00:43:07.046$ to understand how these signals are.

NOTE Confidence: 0.935188366842105

 $00{:}43{:}07.050 \dashrightarrow 00{:}43{:}09.310$ Even if these signals matter.

NOTE Confidence: 0.935188366842105

 $00:43:09.310 \longrightarrow 00:43:11.086$ As far as the choices the animals make,

NOTE Confidence: 0.935188366842105

 $00:43:11.090 \longrightarrow 00:43:14.204$ and so he again turns to the notion of.

NOTE Confidence: 0.935188366842105

 $00:43:14.210 \longrightarrow 00:43:15.758$ Reward choice, so you need more

NOTE Confidence: 0.935188366842105

 $00{:}43{:}15.758 \dashrightarrow 00{:}43{:}17.579$ than one reward at the same time

NOTE Confidence: 0.935188366842105

 $00{:}43{:}17.579 \dashrightarrow 00{:}43{:}19.280$ and to provide a setting where he

NOTE Confidence: 0.935188366842105

 $00{:}43{:}19.335 \dashrightarrow 00{:}43{:}20.940$ could try to understand choice.

NOTE Confidence: 0.935188366842105

 $00:43:20.940 \longrightarrow 00:43:22.794$ He thought it would make the

 $00:43:22.794 \longrightarrow 00:43:24.941$ most sense to make the animals

NOTE Confidence: 0.935188366842105

00:43:24.941 --> 00:43:27.046 choices change through the session

NOTE Confidence: 0.935188366842105

 $00:43:27.046 \longrightarrow 00:43:29.100$ by changing their motivation.

NOTE Confidence: 0.935188366842105

 $00:43:29.100 \longrightarrow 00:43:31.896$ He was interested in understanding how

NOTE Confidence: 0.935188366842105

 $00:43:31.896 \longrightarrow 00:43:33.760$ motivational state impacts choice.

NOTE Confidence: 0.935188366842105

00:43:33.760 --> 00:43:36.350 And I was very interested in this

NOTE Confidence: 0.935188366842105

 $00:43:36.350 \longrightarrow 00:43:38.145$ because motivational state and how

NOTE Confidence: 0.935188366842105

 $00:43:38.145 \longrightarrow 00:43:40.301$ it might be relieved by rewards you

NOTE Confidence: 0.935188366842105

00:43:40.301 --> 00:43:42.756 choose is a nice analogy for eventually

NOTE Confidence: 0.935188366842105

00:43:42.756 --> 00:43:44.874 thinking about how drug craving may

NOTE Confidence: 0.935188366842105

 $00:43:44.874 \longrightarrow 00:43:47.076$ operate within the system and how

NOTE Confidence: 0.935188366842105

00:43:47.076 --> 00:43:49.080 taking drugs may reduce craving.

NOTE Confidence: 0.935188366842105

 $00:43:49.080 \longrightarrow 00:43:50.907$ And then what happens in the brain.

NOTE Confidence: 0.935188366842105

 $00:43:50.910 \longrightarrow 00:43:52.246$ So that's the Longview.

NOTE Confidence: 0.935188366842105

 $00:43:52.246 \longrightarrow 00:43:53.916$ But in the short view,

NOTE Confidence: 0.935188366842105

 $00:43:53.920 \longrightarrow 00:43:56.435$ David wanted to ask specifically

 $00:43:56.435 \longrightarrow 00:43:58.447$ about motivational state shifts

NOTE Confidence: 0.935188366842105

 $00:43:58.447 \longrightarrow 00:44:01.400$ and how they may impact animals

NOTE Confidence: 0.935188366842105

00:44:01.400 --> 00:44:03.770 decision making through this system.

NOTE Confidence: 0.935188366842105

 $00:44:03.770 \longrightarrow 00:44:06.118$ Where we're recording expected

NOTE Confidence: 0.935188366842105

00:44:06.118 --> 00:44:08.466 reward and reward preference,

NOTE Confidence: 0.935188366842105

 $00:44:08.470 \longrightarrow 00:44:10.346$ and so he's doing this in the

NOTE Confidence: 0.935188366842105

 $00:44:10.346 \longrightarrow 00:44:12.328$ face of a shift in thirst.

NOTE Confidence: 0.935188366842105

 $00:44:12.330 \longrightarrow 00:44:13.685$ So obviously whether you choose

NOTE Confidence: 0.935188366842105

 $00:44:13.685 \longrightarrow 00:44:15.339$ food or water will depend on

NOTE Confidence: 0.935188366842105

00:44:15.339 --> 00:44:16.569 if you're hungry or thirsty,

NOTE Confidence: 0.935188366842105

 $00:44:16.570 \longrightarrow 00:44:18.546$ so this is a a kind of motivational

NOTE Confidence: 0.935188366842105

 $00:44:18.546 \longrightarrow 00:44:20.433$ shift that has great relevance to

NOTE Confidence: 0.935188366842105

 $00{:}44{:}20.433 \dashrightarrow 00{:}44{:}22.098$ the natural functioning of this

NOTE Confidence: 0.935188366842105

00:44:22.098 --> 00:44:23.999 circuit and that we thought would

NOTE Confidence: 0.935188366842105

 $00:44:23.999 \longrightarrow 00:44:25.504$ help us understand the natural

 $00:44:25.510 \longrightarrow 00:44:27.082$ functioning of this circuit.

NOTE Confidence: 0.935188366842105

 $00{:}44{:}27.082 \dashrightarrow 00{:}44{:}29.440$ So David developed what he called

NOTE Confidence: 0.935188366842105

00:44:29.505 --> 00:44:31.277 the dynamic preference task.

NOTE Confidence: 0.935188366842105

 $00:44:31.280 \longrightarrow 00:44:33.836$ So this task is very simple.

NOTE Confidence: 0.935188366842105

 $00:44:33.840 \longrightarrow 00:44:36.948$ Some animals are choosing between sucrose

NOTE Confidence: 0.935188366842105

 $00:44:36.948 \longrightarrow 00:44:40.549$ and water reward by pressing a lever.

NOTE Confidence: 0.935188366842105

 $00:44:40.550 \longrightarrow 00:44:42.338$ They begin each day,

NOTE Confidence: 0.935188366842105

 $00:44:42.338 \longrightarrow 00:44:44.573$ thirsty and within the session

NOTE Confidence: 0.935188366842105

 $00:44:44.573 \longrightarrow 00:44:46.858$ they assuaged their thirst.

NOTE Confidence: 0.935188366842105

00:44:46.860 --> 00:44:48.708 And within this session,

NOTE Confidence: 0.935188366842105

 $00:44:48.708 \longrightarrow 00:44:50.556$ besides the choice trials,

NOTE Confidence: 0.935188366842105

 $00:44:50.560 \longrightarrow 00:44:53.212$ it's critical that David also had

NOTE Confidence: 0.935188366842105

 $00:44:53.212 \longrightarrow 00:44:55.548$ forced choice trials where throughout

NOTE Confidence: 0.935188366842105

 $00:44:55.548 \longrightarrow 00:44:58.506$ time the animals had to experience

NOTE Confidence: 0.935188366842105

 $00:44:58.506 \longrightarrow 00:45:00.434$ water and experienced sucrose

NOTE Confidence: 0.935188366842105

 $00{:}45{:}00.434 \dashrightarrow 00{:}45{:}03.010$ so that he could monitor the VP

 $00:45:03.010 \longrightarrow 00:45:05.094$ signals to that through the session,

NOTE Confidence: 0.935188366842105

 $00{:}45{:}05.094 \dashrightarrow 00{:}45{:}07.878$ and so this will become clear when I

NOTE Confidence: 0.935188366842105

 $00:45:07.878 \longrightarrow 00:45:09.984$ explain again how this procedure works.

NOTE Confidence: 0.935188366842105

 $00:45:09.990 \longrightarrow 00:45:13.140$ So animals rats are in the

NOTE Confidence: 0.935188366842105

00:45:13.140 --> 00:45:14.982 behavioral chamber, their electrodes,

NOTE Confidence: 0.935188366842105

 $00:45:14.982 \longrightarrow 00:45:16.850$ and their ventral pallidum 60.

NOTE Confidence: 0.935188366842105

 $00:45:16.850 \longrightarrow 00:45:18.530$ Percent of the trials they receive

NOTE Confidence: 0.935188366842105

 $00{:}45{:}18.530 \dashrightarrow 00{:}45{:}20.557$ over an hour and a half are the

NOTE Confidence: 0.935188366842105

 $00:45:20.557 \longrightarrow 00:45:22.460$ same as what we talked about before.

NOTE Confidence: 0.935188366842105

 $00:45:22.460 \longrightarrow 00:45:24.595$ There's a queue that tells them go

NOTE Confidence: 0.935188366842105

 $00:45:24.595 \longrightarrow 00:45:26.826$ to the reward port and 50% of the

NOTE Confidence: 0.935188366842105

 $00:45:26.826 \longrightarrow 00:45:28.444$ time they get sucrose. 50% water.

NOTE Confidence: 0.935188366842105

 $00{:}45{:}28.444 \dashrightarrow 00{:}45{:}29.088$ It's randomized.

NOTE Confidence: 0.935188366842105

00:45:29.088 --> 00:45:31.250 They don't know what it will be.

NOTE Confidence: 0.935188366842105

 $00:45:31.250 \longrightarrow 00:45:33.284$ These are the forced choice trials

 $00:45:33.284 \longrightarrow 00:45:35.428$ they have to complete this to go

NOTE Confidence: 0.935188366842105

00:45:35.428 --> 00:45:36.638 on to the next trial.

NOTE Confidence: 0.935188366842105

 $00:45:36.640 \longrightarrow 00:45:38.782$ 40% of the time they hear a cue that

NOTE Confidence: 0.935188366842105

 $00:45:38.782 \longrightarrow 00:45:40.496$ tells them it's a choice trial.

NOTE Confidence: 0.935188366842105

 $00:45:40.500 \longrightarrow 00:45:42.418$ They get to pick if they get.

NOTE Confidence: 0.935188366842105

00:45:42.420 --> 00:45:44.640 If they receive sucrose or water

NOTE Confidence: 0.935188366842105

 $00:45:44.640 \longrightarrow 00:45:46.580$ by pressing the relevant lover.

NOTE Confidence: 0.935188366842105

 $00:45:46.580 \longrightarrow 00:45:48.070$ So we have a mix.

NOTE Confidence: 0.935188366842105

 $00:45:48.070 \longrightarrow 00:45:49.726$ Of these outcome choice trials where

NOTE Confidence: 0.935188366842105

 $00:45:49.726 \longrightarrow 00:45:52.030$ we can see their behavioral preference,

NOTE Confidence: 0.935188366842105

 $00:45:52.030 \longrightarrow 00:45:54.802$ what do they want at that moment in time?

NOTE Confidence: 0.935188366842105

 $00:45:54.810 \longrightarrow 00:45:56.562$ And we also have the forest

NOTE Confidence: 0.935188366842105

00:45:56.562 --> 00:45:57.730 trials where we can't

NOTE Confidence: 0.941335286875

 $00{:}45{:}57.799 \to 00{:}46{:}00.247$ see their preference from their behavior,

NOTE Confidence: 0.941335286875

 $00:46:00.250 \longrightarrow 00:46:02.914$ but instead we can look at how their

NOTE Confidence: 0.941335286875

 $00:46:02.914 \longrightarrow 00:46:05.067$ neurons respond to the two rewards and

 $00:46:05.067 \longrightarrow 00:46:07.494$ see if it changes as their choices change.

NOTE Confidence: 0.941335286875

 $00:46:07.494 \longrightarrow 00:46:10.276$ So we use both of these kinds of

NOTE Confidence: 0.941335286875

00:46:10.276 --> 00:46:12.640 trials to get important behavioral and

NOTE Confidence: 0.941335286875

 $00:46:12.640 \longrightarrow 00:46:15.036$ neural data that we want to relate.

NOTE Confidence: 0.941335286875

 $00:46:15.040 \longrightarrow 00:46:16.948$ And what you see behaviourally when

NOTE Confidence: 0.941335286875

 $00:46:16.948 \longrightarrow 00:46:19.459$ you look at the responses of a rat

NOTE Confidence: 0.941335286875

 $00:46:19.459 \longrightarrow 00:46:21.648$ in this kind of procedure is that

NOTE Confidence: 0.941335286875

 $00:46:21.648 \longrightarrow 00:46:23.468$ they start out choosing water.

NOTE Confidence: 0.941335286875

00:46:23.470 --> 00:46:25.000 That's the long purple lines.

NOTE Confidence: 0.941335286875

 $00:46:25.000 \longrightarrow 00:46:26.788$ This is session time and the

NOTE Confidence: 0.941335286875

 $00:46:26.788 \longrightarrow 00:46:28.524$ number of trials which makes sense.

NOTE Confidence: 0.941335286875

 $00:46:28.524 \longrightarrow 00:46:29.701$ They're thirsty, they're going to

NOTE Confidence: 0.941335286875

 $00{:}46{:}29.701 \dashrightarrow 00{:}46{:}31.370$ press on the water level quite a bit,

NOTE Confidence: 0.941335286875

 $00:46:31.370 \longrightarrow 00:46:32.558$ and as they get less thirsty,

NOTE Confidence: 0.941335286875

 $00:46:32.560 \longrightarrow 00:46:35.409$ they'll press on the water level less.

 $00:46:35.410 \longrightarrow 00:46:37.300$ They'll press on the sucrose level

NOTE Confidence: 0.941335286875

 $00:46:37.300 \longrightarrow 00:46:39.660$ lever a few times in the beginning,

NOTE Confidence: 0.941335286875

 $00:46:39.660 \longrightarrow 00:46:41.788$ but that increases overtime

NOTE Confidence: 0.941335286875

 $00:46:41.788 \longrightarrow 00:46:44.384$ as they become less thirsty,

NOTE Confidence: 0.941335286875

 $00:46:44.384 \longrightarrow 00:46:47.863$ so there's a shift in their choices,

NOTE Confidence: 0.941335286875

00:46:47.870 --> 00:46:50.183 and you can graph that with this green line,

NOTE Confidence: 0.941335286875

 $00:46:50.190 \longrightarrow 00:46:52.310$ which shows their relative preference.

NOTE Confidence: 0.941335286875

00:46:52.310 --> 00:46:54.242 The short black lines tell us

NOTE Confidence: 0.941335286875

 $00{:}46{:}54.242 \dashrightarrow 00{:}46{:}56.150$ when the forced trials occurred,

NOTE Confidence: 0.941335286875

 $00:46:56.150 \longrightarrow 00:46:58.398$ so you see they're forced to sample sucrose

NOTE Confidence: 0.941335286875

00:46:58.398 --> 00:47:00.549 and water throughout the whole session,

NOTE Confidence: 0.941335286875

 $00:47:00.550 \longrightarrow 00:47:03.049$ and we see there be choice behavior

NOTE Confidence: 0.941335286875

 $00:47:03.049 \longrightarrow 00:47:04.630$ through these choice trials.

NOTE Confidence: 0.941335286875

 $00:47:04.630 \longrightarrow 00:47:06.280$ So in all of the subjects.

NOTE Confidence: 0.941335286875

 $00:47:06.280 \longrightarrow 00:47:08.440$ Used in this study that I'll talk about.

NOTE Confidence: 0.941335286875

 $00:47:08.440 \longrightarrow 00:47:11.569$ We see a similar shift in preference

 $00:47:11.569 \longrightarrow 00:47:13.600$ through the behavioral session,

NOTE Confidence: 0.941335286875

 $00:47:13.600 \longrightarrow 00:47:15.736$ so as they become less thirsty,

NOTE Confidence: 0.941335286875

 $00:47:15.740 \longrightarrow 00:47:17.860$ they tend to just respond for the supers,

NOTE Confidence: 0.941335286875

 $00:47:17.860 \longrightarrow 00:47:20.029$ which makes sense.

NOTE Confidence: 0.941335286875

 $00{:}47{:}20.030 \dashrightarrow 00{:}47{:}23.246$ So we see this behavioral shift.

NOTE Confidence: 0.941335286875

 $00:47:23.250 \longrightarrow 00:47:25.476$ What about the neurons in the

NOTE Confidence: 0.941335286875

 $00:47:25.476 \longrightarrow 00:47:27.746$ VP and so to to address this,

NOTE Confidence: 0.941335286875

 $00:47:27.750 \longrightarrow 00:47:30.792$ David looked again at this response

NOTE Confidence: 0.941335286875

 $00:47:30.792 \longrightarrow 00:47:33.924$ to reward that VP neurons emit,

NOTE Confidence: 0.941335286875

 $00:47:33.930 \longrightarrow 00:47:36.054$ so he's looking at this time

NOTE Confidence: 0.941335286875

00:47:36.054 --> 00:47:38.140 period just after reward delivery.

NOTE Confidence: 0.941335286875

00:47:38.140 --> 00:47:41.318 When many neurons fire spikes when they

NOTE Confidence: 0.941335286875

 $00{:}47{:}41.318 \dashrightarrow 00{:}47{:}44.992$ get reward and he's using now a general

NOTE Confidence: 0.941335286875

 $00:47:44.992 \longrightarrow 00:47:48.170$ little mini linear excuse me model to

NOTE Confidence: 0.941335286875

 $00:47:48.170 \longrightarrow 00:47:51.180$ try to understand which aspect of of.

00:47:51.180 --> 00:47:51.693 Uhm,

NOTE Confidence: 0.941335286875

 $00:47:51.693 \longrightarrow 00:47:55.284$ the the design best captures how neurons

NOTE Confidence: 0.941335286875

 $00:47:55.284 \longrightarrow 00:47:58.877$ fire through session time trial by trial.

NOTE Confidence: 0.941335286875

 $00:47:58.880 \longrightarrow 00:48:00.568$ Do they just tend to show a difference?

NOTE Confidence: 0.941335286875

 $00:48:00.570 \longrightarrow 00:48:03.198$ Reflective of the difference in outcome?

NOTE Confidence: 0.941335286875

00:48:03.200 --> 00:48:04.792 Sucrose versus water that's

NOTE Confidence: 0.941335286875

 $00:48:04.792 \longrightarrow 00:48:06.384$ relatively stable over time.

NOTE Confidence: 0.941335286875

 $00:48:06.390 \longrightarrow 00:48:08.466$ Do they just show a decrement

NOTE Confidence: 0.941335286875

 $00:48:08.466 \longrightarrow 00:48:09.850$ or increase in activity?

NOTE Confidence: 0.941335286875

 $00:48:09.850 \longrightarrow 00:48:12.490$ They start satisty as you move through time.

NOTE Confidence: 0.941335286875

 $00:48:12.490 \longrightarrow 00:48:14.255$ Or is there an interaction

NOTE Confidence: 0.941335286875

 $00:48:14.255 \longrightarrow 00:48:15.667$ between these two processes?

NOTE Confidence: 0.941335286875

00:48:15.670 --> 00:48:17.740 And by looking at this statistically,

NOTE Confidence: 0.941335286875

 $00{:}48{:}17.740 \dashrightarrow 00{:}48{:}19.408$ David founded sizeable proportion

NOTE Confidence: 0.941335286875

 $00:48:19.408 \longrightarrow 00:48:21.493$ of neurons that care about.

NOTE Confidence: 0.941335286875

 $00:48:21.500 \longrightarrow 00:48:23.019$ Both of these at the same time,

 $00:48:23.020 \longrightarrow 00:48:25.120$ so their activity fits best.

NOTE Confidence: 0.941335286875

 $00{:}48{:}25.120 \dashrightarrow 00{:}48{:}27.104$ Changing preference through time,

NOTE Confidence: 0.941335286875

 $00:48:27.104 \longrightarrow 00:48:30.800$ so some something to do with satiety.

NOTE Confidence: 0.941335286875

 $00:48:30.800 \longrightarrow 00:48:34.478$ Presumably something to do with preference.

NOTE Confidence: 0.941335286875

 $00:48:34.480 \longrightarrow 00:48:35.568$ And that makes sense.

NOTE Confidence: 0.941335286875

00:48:35.568 --> 00:48:36.928 'cause that's what happens to

NOTE Confidence: 0.941335286875

 $00:48:36.928 \longrightarrow 00:48:38.464$ the behavior with the behavior

NOTE Confidence: 0.941335286875

 $00{:}48{:}38.464 \dashrightarrow 00{:}48{:}40.276$ switches as you move through time.

NOTE Confidence: 0.941335286875

 $00:48:40.280 \longrightarrow 00:48:41.972$ The animals preference for

NOTE Confidence: 0.941335286875

 $00:48:41.972 \longrightarrow 00:48:44.087$ water versus sucrose switches as

NOTE Confidence: 0.941335286875

00:48:44.087 --> 00:48:45.898 they become less thirsty,

NOTE Confidence: 0.941335286875

 $00:48:45.900 \longrightarrow 00:48:47.340$ and so here on the left,

NOTE Confidence: 0.941335286875

 $00:48:47.340 \longrightarrow 00:48:49.628$ if you can see this might be hard,

NOTE Confidence: 0.941335286875

 $00:48:49.630 \longrightarrow 00:48:52.814$ but I'll describe it for you is just

NOTE Confidence: 0.941335286875

 $00:48:52.814 \longrightarrow 00:48:55.547$ an example to show 1 neuron firing

00:48:55.547 --> 00:48:58.803 in a very typical way for the whole

NOTE Confidence: 0.941335286875

 $00{:}48{:}58.803 \dashrightarrow 00{:}49{:}00.828$ population through the session.

NOTE Confidence: 0.941335286875

 $00:49:00.830 \longrightarrow 00:49:03.170$ So at the beginning we have

NOTE Confidence: 0.941335286875

 $00:49:03.170 \longrightarrow 00:49:04.730$ neuron spiking at session.

NOTE Confidence: 0.8900720348

 $00:49:04.730 \longrightarrow 00:49:06.781$ The first trials and at the end

NOTE Confidence: 0.8900720348

 $00:49:06.781 \longrightarrow 00:49:08.712$ the last trials and these shaded

NOTE Confidence: 0.8900720348

 $00:49:08.712 \longrightarrow 00:49:11.022$ areas are the times when the animals

NOTE Confidence: 0.8900720348

 $00:49:11.088 \longrightarrow 00:49:13.268$ drinking sucrose or drinking water.

NOTE Confidence: 0.8900720348

 $00:49:13.270 \longrightarrow 00:49:15.742$ These are the times analyzed and you can

NOTE Confidence: 0.8900720348

 $00:49:15.742 \longrightarrow 00:49:18.616$ see that as the animal first gets sucrose,

NOTE Confidence: 0.8900720348

 $00{:}49{:}18.616 \dashrightarrow 00{:}49{:}20.588$ you see moderate spiking.

NOTE Confidence: 0.8900720348

 $00:49:20.590 \longrightarrow 00:49:22.330$ That increases overtime when

NOTE Confidence: 0.8900720348

 $00:49:22.330 \longrightarrow 00:49:24.505$ the animal first gets water,

NOTE Confidence: 0.8900720348

 $00:49:24.510 \longrightarrow 00:49:26.250$ you see a lot of spiking.

NOTE Confidence: 0.8900720348

 $00:49:26.250 \longrightarrow 00:49:28.062$ That really decreases overtime.

NOTE Confidence: 0.8900720348

 $00:49:28.062 \longrightarrow 00:49:31.726$ If you look at this same kind of

 $00:49:31.726 \longrightarrow 00:49:34.743$ feature overtime for all of the neurons.

NOTE Confidence: 0.8900720348

 $00:49:34.750 \longrightarrow 00:49:37.048$ Plotted here in these two figures,

NOTE Confidence: 0.8900720348

 $00{:}49{:}37.050 \dashrightarrow 00{:}49{:}39.336$ with the sessions divided into quarters,

NOTE Confidence: 0.8900720348

00:49:39.340 --> 00:49:40.278 quarter 1234,

NOTE Confidence: 0.8900720348

 $00:49:40.278 \longrightarrow 00:49:44.030$ you see that the mean reward response to

NOTE Confidence: 0.8900720348

 $00:49:44.124 \longrightarrow 00:49:47.687$ sucrose is moderate and then gets bigger.

NOTE Confidence: 0.8900720348

 $00:49:47.690 \longrightarrow 00:49:50.175$ You see that the mean response to

NOTE Confidence: 0.8900720348

00:49:50.175 --> 00:49:52.463 water starts big and positive and

NOTE Confidence: 0.8900720348

 $00{:}49{:}52.463 \dashrightarrow 00{:}49{:}54.378$ gets smaller and more negative

NOTE Confidence: 0.8900720348

 $00:49:54.378 \longrightarrow 00:49:56.360$ as the session goes on.

NOTE Confidence: 0.8900720348

 $00{:}49{:}56.360 \dashrightarrow 00{:}49{:}58.440$ So you can see this much more easily

NOTE Confidence: 0.8900720348

 $00:49:58.440 \longrightarrow 00:50:00.822$ if we think about the mean here

NOTE Confidence: 0.8900720348

 $00{:}50{:}00.822 \dashrightarrow 00{:}50{:}02.617$ over quarters for sucrose versus

NOTE Confidence: 0.8900720348

00:50:02.687 --> 00:50:04.920 water in this final graph down here,

NOTE Confidence: 0.8900720348

 $00:50:04.920 \longrightarrow 00:50:06.580$ the bend firing rate,

 $00:50:06.580 \longrightarrow 00:50:10.242$ we can see the increase in activity for

NOTE Confidence: 0.8900720348

 $00:50:10.242 \longrightarrow 00:50:13.433$ sucrose overtime in one session and

NOTE Confidence: 0.8900720348

 $00:50:13.433 \longrightarrow 00:50:16.744$ the decrease in mean activity for water.

NOTE Confidence: 0.8900720348

 $00:50:16.750 \longrightarrow 00:50:18.544$ So this is interesting because we

NOTE Confidence: 0.8900720348

 $00:50:18.544 \longrightarrow 00:50:20.396$ see that the neural activity sort

NOTE Confidence: 0.8900720348

 $00:50:20.396 \longrightarrow 00:50:22.190$ of shifts more excited for water

NOTE Confidence: 0.8900720348

 $00:50:22.190 \longrightarrow 00:50:23.410$ in the beginning,

NOTE Confidence: 0.8900720348

 $00:50:23.410 \longrightarrow 00:50:26.448$ more excited for sucrose at the end.

NOTE Confidence: 0.8900720348

00:50:26.450 --> 00:50:28.397 And so does the animals preference, right?

NOTE Confidence: 0.8900720348

 $00:50:28.397 \longrightarrow 00:50:30.966$ The preference of the animus shift similarly.

NOTE Confidence: 0.8900720348

 $00{:}50{:}30.970 \dashrightarrow 00{:}50{:}32.764$ But another thing to note is

NOTE Confidence: 0.8900720348

 $00:50:32.764 \longrightarrow 00:50:34.869$ this isn't like a mirror image.

NOTE Confidence: 0.8900720348

 $00:50:34.870 \longrightarrow 00:50:37.678$ These two curves are exactly symmetrical.

NOTE Confidence: 0.8900720348

 $00:50:37.680 \longrightarrow 00:50:40.120$ This water line really decreases,

NOTE Confidence: 0.8900720348

 $00:50:40.120 \longrightarrow 00:50:42.360$ and the sucrose mine is kind of flat,

NOTE Confidence: 0.8900720348

 $00:50:42.360 \longrightarrow 00:50:44.375$ so this was pretty interesting

00:50:44.375 --> 00:50:46.187 and I just thought, well,

NOTE Confidence: 0.8900720348

 $00:50:46.187 \longrightarrow 00:50:47.369$ that's the way the data are,

NOTE Confidence: 0.8900720348

 $00:50:47.370 \longrightarrow 00:50:50.100$ but given David's beautiful more sort

NOTE Confidence: 0.8900720348

00:50:50.100 --> 00:50:52.094 of quantitative mind, he thought,

NOTE Confidence: 0.8900720348

 $00:50:52.094 \longrightarrow 00:50:52.718$ well, what?

NOTE Confidence: 0.8900720348

 $00:50:52.718 \longrightarrow 00:50:54.980$ How can I explain that particular shape?

NOTE Confidence: 0.8900720348

 $00:50:54.980 \longrightarrow 00:50:56.765$ Is there a way I can characterize?

NOTE Confidence: 0.8900720348

 $00:50:56.770 \longrightarrow 00:50:59.175$ That quantitatively and he started

NOTE Confidence: 0.8900720348

 $00{:}50{:}59.175 \to 00{:}51{:}02.136$ thinking about whether this reward signal

NOTE Confidence: 0.8900720348

 $00:51:02.136 \longrightarrow 00:51:05.004$ that signaling a reward prediction error.

NOTE Confidence: 0.8900720348

00:51:05.010 --> 00:51:06.910 Could it contain more information

NOTE Confidence: 0.8900720348

 $00:51:06.910 \longrightarrow 00:51:09.137$ than just something related to what

NOTE Confidence: 0.8900720348

 $00{:}51{:}09.137 \dashrightarrow 00{:}51{:}11.249$ reward did I get on the last trial?

NOTE Confidence: 0.8900720348

 $00:51:11.250 \longrightarrow 00:51:14.466$ Could it also reflect the value of the

NOTE Confidence: 0.8900720348

 $00:51:14.466 \longrightarrow 00:51:17.816$ whole task as the animals becoming sated?

00:51:17.820 --> 00:51:18.169 So,

NOTE Confidence: 0.8900720348

00:51:18.169 --> 00:51:20.263 so every every both rewards will

NOTE Confidence: 0.8900720348

00:51:20.263 --> 00:51:22.239 become less valuable in some sense,

NOTE Confidence: 0.8900720348

 $00:51:22.240 \longrightarrow 00:51:25.510$ as the animals becoming less thirsty.

NOTE Confidence: 0.8900720348

 $00:51:25.510 \longrightarrow 00:51:28.190$ So I I really thought this was a

NOTE Confidence: 0.8900720348

 $00:51:28.190 \longrightarrow 00:51:30.503$ beautiful insight that he had and he

NOTE Confidence: 0.8900720348

 $00:51:30.503 \longrightarrow 00:51:33.150$ developed based on his prior work with Bill.

NOTE Confidence: 0.8900720348

 $00:51:33.150 \longrightarrow 00:51:35.593$ All models again to fit to the

NOTE Confidence: 0.8900720348

 $00{:}51{:}35.593 \dashrightarrow 00{:}51{:}38.529$ activity of each neuron to see which

NOTE Confidence: 0.8900720348

00:51:38.529 --> 00:51:40.794 kind of quantitative model best

NOTE Confidence: 0.8900720348

 $00{:}51{:}40.794 \dashrightarrow 00{:}51{:}42.995$ explained the way the neurons fired

NOTE Confidence: 0.8900720348

 $00:51:42.995 \longrightarrow 00:51:45.543$ and on the left you see the firing

NOTE Confidence: 0.8900720348

00:51:45.543 --> 00:51:47.580 rate in a session of an example

NOTE Confidence: 0.8900720348

 $00{:}51{:}47.649 \dashrightarrow 00{:}51{:}49.842$ neuron just to remind us it's the

NOTE Confidence: 0.8900720348

00:51:49.842 --> 00:51:51.858 increase in response to sucrose at

NOTE Confidence: 0.8900720348

 $00:51:51.858 \longrightarrow 00:51:54.310$ at the top that orangey red line

 $00:51:54.310 \longrightarrow 00:51:56.500$ is moderate and there's a sharp.

NOTE Confidence: 0.8900720348

 $00:51:56.500 \longrightarrow 00:51:58.560$ More dramatic decrease in responding

NOTE Confidence: 0.8900720348

 $00:51:58.560 \longrightarrow 00:52:01.058$ to water in this blueish purple

NOTE Confidence: 0.8900720348

 $00:52:01.058 \longrightarrow 00:52:03.565$ line and so you can ask if just

NOTE Confidence: 0.8900720348

 $00{:}52{:}03.565 \dashrightarrow 00{:}52{:}05.449$ a simple straight line satiety.

NOTE Confidence: 0.8900720348

 $00:52:05.450 \longrightarrow 00:52:07.630$ Does that explain best the

NOTE Confidence: 0.8900720348

 $00:52:07.630 \longrightarrow 00:52:09.374$ way the firing changes?

NOTE Confidence: 0.8900720348

 $00:52:09.380 \longrightarrow 00:52:11.265$ Is the firing explained best

NOTE Confidence: 0.8900720348

 $00:52:11.265 \longrightarrow 00:52:13.150$ by a preference switch that

NOTE Confidence: 0.887304820869565

 $00:52:13.219 \longrightarrow 00:52:15.374$ it would be perfectly symmetrical

NOTE Confidence: 0.887304820869565

 $00{:}52{:}15.374 \dashrightarrow 00{:}52{:}18.269$ preferences just from you know zero to 1?

NOTE Confidence: 0.887304820869565

 $00:52:18.270 \longrightarrow 00:52:20.790$ Or what about both of these together?

NOTE Confidence: 0.887304820869565

 $00:52:20.790 \longrightarrow 00:52:22.558$ And so for David,

NOTE Confidence: 0.887304820869565

 $00:52:22.558 \longrightarrow 00:52:24.768$ that's just a linear combination.

NOTE Confidence: 0.887304820869565

 $00:52:24.770 \longrightarrow 00:52:27.086$ Of models describing both of these,

 $00:52:27.090 \longrightarrow 00:52:29.596$ and then when you combine these literally,

NOTE Confidence: 0.887304820869565

 $00:52:29.600 \longrightarrow 00:52:31.670$ it looks like this and you know you can

NOTE Confidence: 0.887304820869565

 $00:52:31.670 \longrightarrow 00:52:33.910$ see where I'm going because already the

NOTE Confidence: 0.887304820869565

 $00:52:33.910 \longrightarrow 00:52:36.127$ model shape looks similar to the neural

NOTE Confidence: 0.887304820869565

00:52:36.127 --> 00:52:37.837 shapes that we've been looking at.

NOTE Confidence: 0.887304820869565

 $00:52:37.840 \longrightarrow 00:52:39.415$ And so here again, is the model.

NOTE Confidence: 0.887304820869565

 $00:52:39.420 \longrightarrow 00:52:41.895$ When he looked at each neuron and fit its

NOTE Confidence: 0.887304820869565

00:52:41.895 --> 00:52:43.999 activity to these three different models,

NOTE Confidence: 0.887304820869565

 $00:52:44.000 \longrightarrow 00:52:46.682$ he finds that the best fit

NOTE Confidence: 0.887304820869565

 $00:52:46.682 \longrightarrow 00:52:49.160$ model is this mixed model.

NOTE Confidence: 0.887304820869565

 $00{:}52{:}49.160 \dashrightarrow 00{:}52{:}51.098$ What this means is most neurons

NOTE Confidence: 0.887304820869565

00:52:51.098 --> 00:52:53.299 seem to care about both satiety,

NOTE Confidence: 0.887304820869565

 $00:52:53.300 \longrightarrow 00:52:56.177$ so movement through the session in time

NOTE Confidence: 0.887304820869565

00:52:56.177 --> 00:52:59.078 and their current preference for reward,

NOTE Confidence: 0.887304820869565

 $00:52:59.080 \longrightarrow 00:53:01.940$ which one they're liking better.

NOTE Confidence: 0.887304820869565

 $00:53:01.940 \longrightarrow 00:53:03.524$ So that's pretty cool.

 $00:53:03.524 \longrightarrow 00:53:05.630$ So, So what David showed us is

NOTE Confidence: 0.887304820869565

 $00:53:05.630 \longrightarrow 00:53:07.650$ that it's not just the immediate

NOTE Confidence: 0.887304820869565

 $00:53:07.650 \longrightarrow 00:53:10.086$ difference in reward value that is

NOTE Confidence: 0.887304820869565

00:53:10.086 --> 00:53:12.039 being reflected in this activity,

NOTE Confidence: 0.887304820869565

 $00:53:12.040 \longrightarrow 00:53:15.645$ but there's also an impact of satiety.

NOTE Confidence: 0.887304820869565

 $00:53:15.650 \longrightarrow 00:53:18.950$ So this was all analyzed based

NOTE Confidence: 0.887304820869565

00:53:18.950 --> 00:53:21.454 on forced trial data, right?

NOTE Confidence: 0.887304820869565

 $00{:}53{:}21.454 \dashrightarrow 00{:}53{:}23.582$ 'cause we're looking at how the animal

NOTE Confidence: 0.887304820869565

 $00{:}53{:}23.582 \to 00{:}53{:}25.948$ responds to the reward through session time.

NOTE Confidence: 0.887304820869565

 $00:53:25.950 \longrightarrow 00:53:28.050$ But we have all of these choice

NOTE Confidence: 0.887304820869565

 $00{:}53{:}28.050 \dashrightarrow 00{:}53{:}30.173$ trials for the animals making its

NOTE Confidence: 0.887304820869565

 $00{:}53{:}30.173 \dashrightarrow 00{:}53{:}32.098$ own decision about which reward

NOTE Confidence: 0.887304820869565

00:53:32.098 --> 00:53:34.530 it wants at that given time,

NOTE Confidence: 0.887304820869565

 $00:53:34.530 \longrightarrow 00:53:36.605$ and what David wondered is,

NOTE Confidence: 0.887304820869565

 $00:53:36.610 \longrightarrow 00:53:39.340$ do these responses of the neurons

 $00:53:39.340 \longrightarrow 00:53:42.270$ that tell us how much the animal,

NOTE Confidence: 0.887304820869565

 $00:53:42.270 \longrightarrow 00:53:44.610$ what the animal thinks about the

NOTE Confidence: 0.887304820869565

00:53:44.690 --> 00:53:47.270 reward relative to its expectation,

NOTE Confidence: 0.887304820869565

 $00:53:47.270 \longrightarrow 00:53:49.178$ does that have anything to do

NOTE Confidence: 0.887304820869565

 $00:53:49.178 \longrightarrow 00:53:50.132$ with their behavior?

NOTE Confidence: 0.887304820869565

 $00.53.50.140 \longrightarrow 00.53.51.108$ Because in the end,

NOTE Confidence: 0.887304820869565

 $00:53:51.108 \longrightarrow 00:53:53.352$ we'd like to try to get an understanding

NOTE Confidence: 0.887304820869565

 $00:53:53.352 \longrightarrow 00:53:55.548$ of how these systems impact choice.

NOTE Confidence: 0.887304820869565

 $00{:}53{:}55.550 {\:\dashrightarrow\:} 00{:}53{:}57.406$ That's our eventual goal.

NOTE Confidence: 0.887304820869565

 $00:53:57.406 \longrightarrow 00:54:00.190$ So the way that David decided

NOTE Confidence: 0.887304820869565

 $00:54:00.281 \longrightarrow 00:54:01.969$ to think about that.

NOTE Confidence: 0.887304820869565

 $00:54:01.970 \longrightarrow 00:54:04.784$ Was to look at the animals behavior.

NOTE Confidence: 0.887304820869565

 $00:54:04.790 \longrightarrow 00:54:06.150$ Good idea to do first.

NOTE Confidence: 0.887304820869565

 $00:54:06.150 \longrightarrow 00:54:08.790$ Here are three rat examples on the left.

NOTE Confidence: 0.887304820869565

00:54:08.790 --> 00:54:11.106 You're looking at the animals choice

NOTE Confidence: 0.887304820869565

00:54:11.106 --> 00:54:12.650 behavior through the session.

 $00:54:12.650 \longrightarrow 00:54:14.582$ The purple bars on the bottom are

NOTE Confidence: 0.887304820869565

 $00{:}54{:}14.582 \dashrightarrow 00{:}54{:}16.035$ when the animal chooses water

NOTE Confidence: 0.887304820869565

 $00.54:16.035 \longrightarrow 00:54:18.009$ they do a lot at the beginning,

NOTE Confidence: 0.887304820869565

00:54:18.010 --> 00:54:18.856 last moving forward,

NOTE Confidence: 0.887304820869565

 $00:54:18.856 \longrightarrow 00:54:21.182$ and then they shift and tend to choose

NOTE Confidence: 0.887304820869565

00:54:21.182 --> 00:54:23.268 sucrose more as session time goes on.

NOTE Confidence: 0.887304820869565

 $00:54:23.270 \longrightarrow 00:54:25.160$ So you can plot the preference curve.

NOTE Confidence: 0.887304820869565 00:54:25.160 --> 00:54:25.870 For sucrose,

NOTE Confidence: 0.887304820869565

 $00:54:25.870 \longrightarrow 00:54:27.645$ the preference curve for water

NOTE Confidence: 0.887304820869565

 $00:54:27.645 \longrightarrow 00:54:29.100$ and you see this.

NOTE Confidence: 0.887304820869565

 $00{:}54{:}29.100 \dashrightarrow 00{:}54{:}31.820$ This kind of function and you can see

NOTE Confidence: 0.887304820869565

 $00:54:31.820 \longrightarrow 00:54:34.280$ that example for three different rats.

NOTE Confidence: 0.887304820869565

 $00{:}54{:}34.280 \dashrightarrow 00{:}54{:}37.346$ When David looked at the neural

NOTE Confidence: 0.887304820869565

 $00:54:37.346 \longrightarrow 00:54:40.509$ estimates for this mixed model that

NOTE Confidence: 0.887304820869565

 $00:54:40.509 \longrightarrow 00:54:43.587$ came from what he calculated here,

 $00:54:43.590 \longrightarrow 00:54:45.378$ it came from these forced trials

NOTE Confidence: 0.887304820869565

 $00:54:45.378 \longrightarrow 00:54:47.759$ and tried on a trial by trial

NOTE Confidence: 0.887304820869565

 $00:54:47.759 \longrightarrow 00:54:49.629$ to estimate what the animal's

NOTE Confidence: 0.887304820869565

00:54:49.629 --> 00:54:51.790 preference was just based on the

NOTE Confidence: 0.887304820869565

00:54:51.790 --> 00:54:53.806 neural activity for a given neuron.

NOTE Confidence: 0.887304820869565

 $00:54:53.810 \longrightarrow 00:54:55.310$ Then averaging that across

NOTE Confidence: 0.887304820869565

00:54:55.310 --> 00:54:57.560 all neurons from a given rat,

NOTE Confidence: 0.887304820869565

00:54:57.560 --> 00:54:59.140 you get these preference curves,

NOTE Confidence: 0.887304820869565

 $00{:}54{:}59.140 \dashrightarrow 00{:}55{:}01.840$ and they're just remarkably similar.

NOTE Confidence: 0.887304820869565

 $00:55:01.840 \longrightarrow 00:55:04.108$ You don't even need all the beautiful

NOTE Confidence: 0.887304820869565

 $00{:}55{:}04.108 --> 00{:}55{:}05.922$ statistics to tell you the neurons

NOTE Confidence: 0.887304820869565

 $00:55:05.922 \longrightarrow 00:55:07.728$ are giving the same readout of

NOTE Confidence: 0.887304820869565

 $00{:}55{:}07.728 \dashrightarrow 00{:}55{:}09.569$ what the animals preferences,

NOTE Confidence: 0.887304820869565

00:55:09.570 --> 00:55:12.006 moment by moment as the decision

NOTE Confidence: 0.887304820869565

 $00:55:12.006 \longrightarrow 00:55:13.224$ the animal makes.

NOTE Confidence: 0.887304820869565

 $00:55:13.230 \longrightarrow 00:55:16.282$ So this is a really nice correlation

 $00:55:16.282 \longrightarrow 00:55:18.982$ that helps us build on the idea

NOTE Confidence: 0.887304820869565

 $00{:}55{:}18.982 \dashrightarrow 00{:}55{:}21.070$ that this signal is important for

NOTE Confidence: 0.93921235

00:55:21.145 --> 00:55:23.417 the animals future decision, and.

NOTE Confidence: 0.93921235

 $00:55:23.417 \longrightarrow 00:55:24.845$ That's what these graphs

NOTE Confidence: 0.93921235

 $00:55:24.845 \longrightarrow 00:55:26.630$ here on the right support.

NOTE Confidence: 0.93921235

 $00:55:26.630 \longrightarrow 00:55:29.246$ If you look at the correlation for each

NOTE Confidence: 0.93921235

 $00:55:29.246 \longrightarrow 00:55:32.071$ neuron of its activity with the animals

NOTE Confidence: 0.93921235

00:55:32.071 --> 00:55:34.171 preference for neurons like that.

NOTE Confidence: 0.93921235

 $00{:}55{:}34.180 \dashrightarrow 00{:}55{:}36.322$ Were weighted within this mixed model

NOTE Confidence: 0.93921235

 $00:55:36.322 \dashrightarrow 00:55:38.279$ that care about outcome in time.

NOTE Confidence: 0.93921235

 $00:55:38.280 \dashrightarrow 00:55:39.978$ We see that correlation is very

NOTE Confidence: 0.93921235

 $00:55:39.978 \longrightarrow 00:55:42.240$ close to one for very many of them.

NOTE Confidence: 0.93921235

00:55:42.240 --> 00:55:44.580 If you ask from the neural

NOTE Confidence: 0.93921235

00:55:44.580 --> 00:55:47.098 activity when in time this switch

NOTE Confidence: 0.93921235

00:55:47.098 --> 00:55:49.720 point might be for each neuron.

 $00:55:49.720 \longrightarrow 00:55:53.020$ Many neurons that care about outcome

NOTE Confidence: 0.93921235

 $00{:}55{:}53.020 \longrightarrow 00{:}55{:}55.935$ by time give you a very close

NOTE Confidence: 0.93921235

00:55:55.935 --> 00:55:57.750 estimation of the actual trial.

NOTE Confidence: 0.93921235

 $00:55:57.750 \longrightarrow 00:55:59.460$ So the switch point is zero.

NOTE Confidence: 0.93921235

 $00:55:59.460 \longrightarrow 00:56:02.516$ You can see many or within 20 trials.

NOTE Confidence: 0.93921235

00:56:02.520 --> 00:56:03.488 So, quantitatively,

NOTE Confidence: 0.93921235

 $00:56:03.488 \longrightarrow 00:56:06.392$ we've got a really nice agreement

NOTE Confidence: 0.93921235

 $00:56:06.392 \longrightarrow 00:56:08.714$ between neural activity with the

NOTE Confidence: 0.93921235

 $00{:}56{:}08.714 \dashrightarrow 00{:}56{:}10.849$ animal actually decides to do.

NOTE Confidence: 0.93921235

 $00:56:10.850 \longrightarrow 00:56:13.482$ So that led save it again to turn

NOTE Confidence: 0.93921235

00:56:13.482 --> 00:56:15.683 to optogenetics to see if he

NOTE Confidence: 0.93921235

 $00:56:15.683 \longrightarrow 00:56:17.528$ could manipulate the system and

NOTE Confidence: 0.93921235

 $00:56:17.528 \longrightarrow 00:56:19.308$ manipulate the subjects choice.

NOTE Confidence: 0.93921235

00:56:19.310 --> 00:56:21.596 And this is the final little bit of data,

NOTE Confidence: 0.93921235

 $00:56:21.600 \longrightarrow 00:56:24.156$ bit of data that I'll be showing you and

NOTE Confidence: 0.93921235

 $00:56:24.156 \longrightarrow 00:56:26.530$ then we can discuss this as you wish.

 $00:56:26.530 \longrightarrow 00:56:29.482$ So now David wants to see if by

NOTE Confidence: 0.93921235

 $00{:}56{:}29.482 \dashrightarrow 00{:}56{:}31.242$ controlling ventral pallidal neuron

NOTE Confidence: 0.93921235

00:56:31.242 --> 00:56:34.068 activity he can impact their choice,

NOTE Confidence: 0.93921235

 $00:56:34.070 \longrightarrow 00:56:37.166$ so so he has optimal control.

NOTE Confidence: 0.93921235

00:56:37.170 --> 00:56:39.508 He's now not going to make the

NOTE Confidence: 0.93921235

 $00{:}56{:}39.508 \dashrightarrow 00{:}56{:}41.268$ animals thirsty, he's just going to.

NOTE Confidence: 0.93921235

00:56:41.268 --> 00:56:43.416 Go back to the situation where they're

NOTE Confidence: 0.93921235

 $00:56:43.416 \longrightarrow 00:56:45.686$ choosing between sucrose and maltodextrin.

NOTE Confidence: 0.93921235

 $00:56:45.690 \longrightarrow 00:56:47.720$ This is a situation when they're not

NOTE Confidence: 0.93921235

 $00:56:47.720 \longrightarrow 00:56:49.336$ thirsty and their behavior through

NOTE Confidence: 0.93921235

 $00{:}56{:}49.336 \dashrightarrow 00{:}56{:}51.412$ the session tends to be relatively

NOTE Confidence: 0.93921235

 $00{:}56{:}51.412 \dashrightarrow 00{:}56{:}53.550$ stable and he wants the behavior

NOTE Confidence: 0.93921235

 $00{:}56{:}53.550 \dashrightarrow 00{:}56{:}55.644$ to be relatively stable because now

NOTE Confidence: 0.93921235

 $00:56:55.644 \longrightarrow 00:56:57.786$ he must to go in and try to change

NOTE Confidence: 0.93921235

00:56:57.857 --> 00:57:00.363 it by messing with the BP reward

 $00:57:00.363 \longrightarrow 00:57:01.437$ prediction error signal.

NOTE Confidence: 0.93921235

 $00:57:01.440 \longrightarrow 00:57:04.360$ So what he's going to do is express

NOTE Confidence: 0.93921235

 $00:57:04.360 \longrightarrow 00:57:05.907$ channelrhodopsin in ventral palatal

NOTE Confidence: 0.93921235

 $00{:}57{:}05.907 \dashrightarrow 00{:}57{:}08.139$ neurons and shine light on them

NOTE Confidence: 0.93921235

 $00:57:08.139 \longrightarrow 00:57:10.983$ to force them to fire every time

NOTE Confidence: 0.93921235

00:57:10.983 --> 00:57:12.607 the animal drinks maltodextrin.

NOTE Confidence: 0.93921235

 $00:57:12.610 \longrightarrow 00:57:14.605$ And in the procedure that he uses,

NOTE Confidence: 0.93921235

00.57:14.610 --> 00.57:17.472 there's going to be forced choice

NOTE Confidence: 0.93921235

 $00:57:17.472 \longrightarrow 00:57:20.266$ trials so he can continue to make

NOTE Confidence: 0.93921235

00:57:20.266 --> 00:57:22.427 them drink maltodextrin paired with

NOTE Confidence: 0.93921235

 $00{:}57{:}22.427 \dashrightarrow 00{:}57{:}24.950$ stimulation and choice trials so we

NOTE Confidence: 0.93921235

 $00:57:24.950 \longrightarrow 00:57:27.388$ know what the animal actually would

NOTE Confidence: 0.93921235

 $00:57:27.388 \longrightarrow 00:57:30.006$ prefer to drink at any given time.

NOTE Confidence: 0.93921235

00:57:30.010 --> 00:57:32.110 First. Animals are well trained.

NOTE Confidence: 0.93921235

 $00:57:32.110 \longrightarrow 00:57:34.780$ Then there is a session of

NOTE Confidence: 0.93921235

 $00:57:34.780 \longrightarrow 00:57:35.670$ optogenetic manipulation,

 $00:57:35.670 \longrightarrow 00:57:36.898$ so well trained animals.

NOTE Confidence: 0.93921235

 $00:57:36.898 \longrightarrow 00:57:39.549$ And this is just a diagram of when

NOTE Confidence: 0.93921235

 $00:57:39.549 \longrightarrow 00:57:40.668$ the stimulation occurs.

NOTE Confidence: 0.93921235

 $00:57:40.670 \longrightarrow 00:57:43.254$ So at the time that the animals actually

NOTE Confidence: 0.93921235

 $00{:}57{:}43.254 \dashrightarrow 00{:}57{:}44.852$ drinking maltod extrin and we can talk

NOTE Confidence: 0.93921235

 $00:57:44.852 \longrightarrow 00:57:46.719$ about that more if you want to later.

NOTE Confidence: 0.93921235

00:57:46.720 --> 00:57:49.890 So if you look at baseline here on the right,

NOTE Confidence: 0.93921235

 $00{:}57{:}49.890 \dashrightarrow 00{:}57{:}52.326$ a well trained animals prefer sucrose

NOTE Confidence: 0.93921235

 $00:57:52.326 \longrightarrow 00:57:54.510$ based on their choice trials.

NOTE Confidence: 0.93921235

 $00{:}57{:}54.510 \dashrightarrow 00{:}57{:}56.162$ The press the lever to get sucrose

NOTE Confidence: 0.93921235

00:57:56.162 --> 00:57:58.412 most of the time and the blue dots are

NOTE Confidence: 0.93921235

 $00:57:58.412 \longrightarrow 00:58:00.202$ the subjects in which were expressing

NOTE Confidence: 0.93921235

 $00{:}58{:}00.202 \dashrightarrow 00{:}58{:}02.077$ channel rhodopsin the Gray dots or

NOTE Confidence: 0.93921235

 $00:58:02.077 \longrightarrow 00:58:04.314$ subjects expressing the empty vector GFP.

NOTE Confidence: 0.93921235

00:58:04.314 --> 00:58:06.522 Both animals get laser shining in

 $00:58:06.522 \longrightarrow 00:58:08.950$ their brain, but the great author.

NOTE Confidence: 0.93921235

 $00:58:08.950 \longrightarrow 00:58:09.750$ Our control.

NOTE Confidence: 0.93921235

 $00:58:09.750 \longrightarrow 00:58:11.078$ In the test session,

NOTE Confidence: 0.93921235

00:58:11.078 --> 00:58:13.070 the next day after baseline were

NOTE Confidence: 0.93921235

 $00:58:13.138 \longrightarrow 00:58:15.922$ stimulating every time the subject gets

NOTE Confidence: 0.93921235

 $00{:}58{:}15.922 \dashrightarrow 00{:}58{:}18.225$ less preferred reward maltod extrin and

NOTE Confidence: 0.93921235

 $00:58:18.225 \longrightarrow 00:58:20.899$ what you see is it shifts preference

NOTE Confidence: 0.93921235

 $00:58:20.899 \longrightarrow 00:58:23.182$ towards maltodextrin on the choice trials.

NOTE Confidence: 0.93921235

 $00{:}58{:}23.182 \dashrightarrow 00{:}58{:}25.450$ So what does that actually look

NOTE Confidence: 0.92219158

00:58:25.525 --> 00:58:26.809 like through time?

NOTE Confidence: 0.92219158

 $00{:}58{:}26.810 \dashrightarrow 00{:}58{:}29.624$ Here is one session, the test session.

NOTE Confidence: 0.92219158

 $00:58:29.630 \longrightarrow 00:58:31.770$ And here's the smoothed preference

NOTE Confidence: 0.92219158

 $00{:}58{:}31.770 \dashrightarrow 00{:}58{:}34.840$ based on liver choice for each rat

NOTE Confidence: 0.92219158

 $00:58:34.840 \longrightarrow 00:58:37.408$ Gray or the controls relatively stable.

NOTE Confidence: 0.92219158

 $00:58:37.410 \longrightarrow 00:58:39.696$ They mostly want sucrose and blue.

NOTE Confidence: 0.92219158

 $00:58:39.700 \longrightarrow 00:58:41.782$ Are these experimental animals where we've

 $00:58:41.782 \longrightarrow 00:58:43.890$ shifted the preference to maltodextrin,

NOTE Confidence: 0.92219158

 $00{:}58{:}43.890 \dashrightarrow 00{:}58{:}46.658$ and you can see it's a gradual effect

NOTE Confidence: 0.92219158

00:58:46.658 --> 00:58:48.770 that accrues through experience,

NOTE Confidence: 0.92219158

 $00:58:48.770 \longrightarrow 00:58:49.922$ so it's not that the first

NOTE Confidence: 0.92219158

 $00:58:49.922 \longrightarrow 00:58:50.690$ time you stimulate it,

NOTE Confidence: 0.92219158

 $00:58:50.690 \longrightarrow 00:58:51.782$ they immediately shift.

NOTE Confidence: 0.92219158

 $00:58:51.782 \longrightarrow 00:58:53.966$ This is congruent with the idea.

NOTE Confidence: 0.92219158

 $00{:}58{:}53.970 \dashrightarrow 00{:}58{:}55.540$ But it's a learning response.

NOTE Confidence: 0.92219158

 $00{:}58{:}55.540 \dashrightarrow 00{:}58{:}57.190$ You're sending them a signal that

NOTE Confidence: 0.92219158

 $00:58:57.190 \dashrightarrow 00:58:59.040$ that reward is better than expected.

NOTE Confidence: 0.92219158

 $00:58:59.040 \longrightarrow 00:59:01.110$ So maybe you should change what

NOTE Confidence: 0.92219158

 $00:59:01.110 \longrightarrow 00:59:03.050$ you do on upcoming trials.

NOTE Confidence: 0.92219158

 $00{:}59{:}03.050 \dashrightarrow 00{:}59{:}04.988$ And also congruent with the idea

NOTE Confidence: 0.92219158

 $00:59:04.988 \longrightarrow 00:59:07.450$ that this is a learning signal.

NOTE Confidence: 0.92219158

00:59:07.450 --> 00:59:10.015 This behavior change does last

 $00:59:10.015 \longrightarrow 00:59:12.067$ until the next day.

NOTE Confidence: 0.92219158

 $00:59:12.070 \longrightarrow 00:59:15.805$ So the test day is the day of optogenetic

NOTE Confidence: 0.92219158

00:59:15.805 --> 00:59:17.610 manipulation Recovery day one.

NOTE Confidence: 0.92219158

 $00:59:17.610 \longrightarrow 00:59:19.416$ We just see what their choices are

NOTE Confidence: 0.92219158

 $00:59:19.416 \longrightarrow 00:59:21.421$ and you can see that their choices

NOTE Confidence: 0.92219158

 $00:59:21.421 \longrightarrow 00:59:23.173$ still tend to be more towards

NOTE Confidence: 0.92219158

 $00:59:23.233 \longrightarrow 00:59:24.737$ maltodextrins then they weren't

NOTE Confidence: 0.92219158

 $00:59:24.737 \longrightarrow 00:59:26.993$ baseline and this changes over time

NOTE Confidence: 0.92219158

 $00{:}59{:}27.000 \dashrightarrow 00{:}59{:}28.962$ as we no longer ascending that

NOTE Confidence: 0.92219158

 $00{:}59{:}28.962 \dashrightarrow 00{:}59{:}31.324$ fake signal that we could send

NOTE Confidence: 0.92219158

 $00:59:31.324 \longrightarrow 00:59:33.288$ with the optogenetic manipulation.

NOTE Confidence: 0.92219158

 $00:59:33.290 \longrightarrow 00:59:36.034$ So this is a first step to providing

NOTE Confidence: 0.92219158

 $00:59:36.034 \longrightarrow 00:59:38.818$ some evidence that this signal can

NOTE Confidence: 0.92219158

00:59:38.818 --> 00:59:41.320 impact the animals future choice.

NOTE Confidence: 0.92219158

 $00:59:41.320 \longrightarrow 00:59:43.318$ So if you look at their.

NOTE Confidence: 0.92219158

 $00:59:43.320 \longrightarrow 00:59:46.410$ Latency's to choose levers in the

 $00:59:46.410 \longrightarrow 00:59:47.832$ optogenetic stimulation experiment,

NOTE Confidence: 0.92219158

00:59:47.832 --> 00:59:50.538 you find that overtime they tend

NOTE Confidence: 0.92219158

 $00{:}59{:}50.538 \dashrightarrow 00{:}59{:}52.740$ to choose maltod extrin more,

NOTE Confidence: 0.92219158

 $00:59:52.740 \longrightarrow 00:59:54.516$ and on trials choice trials when

NOTE Confidence: 0.92219158

00:59:54.516 --> 00:59:56.440 they're going to choose maltodextrin,

NOTE Confidence: 0.92219158

 $00:59:56.440 \longrightarrow 00:59:58.225$ their latency to go to the lever

NOTE Confidence: 0.92219158

 $00:59:58.225 \longrightarrow 01:00:00.010$ to make that choice is faster,

NOTE Confidence: 0.92219158

 $01:00:00.010 \longrightarrow 01:00:02.649$ so we see this change in behavior

NOTE Confidence: 0.92219158

 $01:00:02.649 \longrightarrow 01:00:05.118$ that matches what you would expect.

NOTE Confidence: 0.92219158

 $01:00:05.120 \longrightarrow 01:00:07.766$ For this. This kind of signal.

NOTE Confidence: 0.92219158

01:00:07.770 --> 01:00:09.970 So what I told you is that initially

NOTE Confidence: 0.92219158

 $01:00:09.970 \longrightarrow 01:00:12.446$ we see a signal in the ventral

NOTE Confidence: 0.92219158

 $01:00:12.446 \longrightarrow 01:00:14.301$ pallidum when animals are actually

NOTE Confidence: 0.92219158

 $01{:}00{:}14.374 \dashrightarrow 01{:}00{:}16.490$ experiencing reward ingesting it.

NOTE Confidence: 0.92219158

01:00:16.490 --> 01:00:18.794 That seems to match their relative

01:00:18.794 --> 01:00:20.710 preference at that current time,

NOTE Confidence: 0.92219158

 $01:00:20.710 \longrightarrow 01:00:23.174$ and if you analyze that spike activity

NOTE Confidence: 0.92219158

 $01:00:23.174 \longrightarrow 01:00:25.426$ relative to the current time and

NOTE Confidence: 0.92219158

 $01:00:25.426 \longrightarrow 01:00:27.682$ the reward period just before that,

NOTE Confidence: 0.92219158

 $01:00:27.690 \longrightarrow 01:00:28.650$ and just before that,

NOTE Confidence: 0.92219158

01:00:28.650 --> 01:00:29.610 and just before that,

NOTE Confidence: 0.92219158

01:00:29.610 --> 01:00:30.579 IE reward history,

NOTE Confidence: 0.92219158

 $01:00:30.579 \longrightarrow 01:00:33.649$ you see that at least a subset of these

NOTE Confidence: 0.92219158

 $01{:}00{:}33.649 \dashrightarrow 01{:}00{:}35.863$ care about reward history and what

NOTE Confidence: 0.92219158

 $01:00:35.863 \longrightarrow 01:00:38.389$ they instead are signaling is a reward.

NOTE Confidence: 0.92219158

01:00:38.390 --> 01:00:40.298 Prediction error is what I just

NOTE Confidence: 0.92219158

 $01:00:40.298 \longrightarrow 01:00:42.220$ got better than I expected.

NOTE Confidence: 0.92219158

 $01:00:42.220 \longrightarrow 01:00:44.424$ The same or worse.

NOTE Confidence: 0.92219158

 $01:00:44.424 \longrightarrow 01:00:47.179$ And so these same signals,

NOTE Confidence: 0.92219158

 $01:00:47.180 \longrightarrow 01:00:49.625$ these moment by moment reward

NOTE Confidence: 0.92219158

 $01{:}00{:}49.625 \dashrightarrow 01{:}00{:}52.070$ prediction error signals also care

 $01:00:52.151 \longrightarrow 01:00:54.976$ about the current motivational state.

NOTE Confidence: 0.92219158

 $01:00:54.980 \longrightarrow 01:00:57.194$ They're also able to integrate the

NOTE Confidence: 0.92219158

01:00:57.194 --> 01:00:59.504 larger change in value that might

NOTE Confidence: 0.92219158

01:00:59.504 --> 01:01:01.796 happen as your motivational state is

NOTE Confidence: 0.92219158

 $01{:}01{:}01{:}01{:}03{:}989$ changing as you get less thirsty.

NOTE Confidence: 0.92219158

01:01:03.990 --> 01:01:04.752 And hypothetically,

NOTE Confidence: 0.92219158

01:01:04.752 --> 01:01:07.419 in other situations we haven't tried yet,

NOTE Confidence: 0.92219158

 $01:01:07.420 \longrightarrow 01:01:07.825$ right?

NOTE Confidence: 0.92219158

 $01:01:07.825 \longrightarrow 01:01:10.255$ As your craving might be reduced

NOTE Confidence: 0.92219158

 $01:01:10.255 \longrightarrow 01:01:12.010$ as you ingest drugs.

NOTE Confidence: 0.92219158

 $01:01:12.010 \longrightarrow 01:01:15.489$ As hunger changes as you eat etc and

NOTE Confidence: 0.92219158

 $01:01:15.489 \longrightarrow 01:01:17.512$ so these signals that occur at the

NOTE Confidence: 0.92219158

 $01{:}01{:}17.512 \dashrightarrow 01{:}01{:}19.688$ time the animals ingesting reward,

NOTE Confidence: 0.92219158

 $01:01:19.690 \longrightarrow 01:01:21.640$ they affect their future behavior.

NOTE Confidence: 0.92219158

 $01:01:21.640 \longrightarrow 01:01:25.259$ As we saw in that very simple.

01:01:25.260 --> 01:01:27.500 A measure of how close you are to the port,

NOTE Confidence: 0.92219158

 $01:01:27.500 \longrightarrow 01:01:28.616$ and as we saw in this,

NOTE Confidence: 0.92219158

01:01:28.620 --> 01:01:30.712 perhaps more informative choice

NOTE Confidence: 0.92219158

 $01:01:30.712 \longrightarrow 01:01:33.327$ procedure where animals are choosing

NOTE Confidence: 0.92219158

 $01:01:33.327 \longrightarrow 01:01:35.517$ which lever to push in order

NOTE Confidence: 0.92219158

01:01:35.517 --> 01:01:37.107 to get the reward that

NOTE Confidence: 0.9222073612

 $01:01:37.182 \longrightarrow 01:01:38.807$ they want at that time.

NOTE Confidence: 0.9222073612

01:01:38.810 --> 01:01:40.430 And so the big question,

NOTE Confidence: 0.9222073612

 $01:01:40.430 \longrightarrow 01:01:41.850$ of course, is what?

NOTE Confidence: 0.9222073612

 $01:01:41.850 \longrightarrow 01:01:43.625$ What do these signals mean

NOTE Confidence: 0.9222073612

 $01:01:43.625 \longrightarrow 01:01:45.690$ for the circuit as a whole?

NOTE Confidence: 0.9222073612

 $01:01:45.690 \longrightarrow 01:01:47.679$ So if I go back to the statement that

NOTE Confidence: 0.9222073612

 $01:01:47.679 \longrightarrow 01:01:49.974$ I made at the beginning that usually

NOTE Confidence: 0.9222073612

 $01:01:49.974 \longrightarrow 01:01:51.955$ the ventral pallidum was the more

NOTE Confidence: 0.9222073612

 $01:01:51.955 \longrightarrow 01:01:53.985$ boring area that was just the output.

NOTE Confidence: 0.9222073612

 $01:01:53.990 \longrightarrow 01:01:55.890$ For the fantastically interesting

01:01:55.890 --> 01:01:57.550 nucleus incumbents, of course,

NOTE Confidence: 0.9222073612

 $01:01:57.550 \longrightarrow 01:01:59.325$ the nucleus of Cummins is

NOTE Confidence: 0.9222073612

 $01:01:59.325 \longrightarrow 01:02:00.035$ fantastically interesting.

NOTE Confidence: 0.9222073612

 $01:02:00.040 \longrightarrow 01:02:01.912$ But these are big excitatory signals

NOTE Confidence: 0.9222073612

 $01:02:01.912 \longrightarrow 01:02:03.813$ in the ventral pallidum unlikely to

NOTE Confidence: 0.9222073612

 $01:02:03.813 \longrightarrow 01:02:05.607$ be driven by the Gabaergic medium.

NOTE Confidence: 0.9222073612

01:02:05.610 --> 01:02:08.388 Spiny neurons of the nucleus accumbens,

NOTE Confidence: 0.9222073612

 $01:02:08.390 \longrightarrow 01:02:09.306$ and when we look.

NOTE Confidence: 0.9222073612

 $01:02:09.306 \longrightarrow 01:02:10.451$ And David didn't record in

NOTE Confidence: 0.9222073612

 $01:02:10.451 \longrightarrow 01:02:11.539$ the comments as well.

NOTE Confidence: 0.9222073612

 $01{:}02{:}11.540 \dashrightarrow 01{:}02{:}12.660$ When we look there,

NOTE Confidence: 0.9222073612

 $01:02:12.660 \longrightarrow 01:02:14.759$ we don't see the large numbers of

NOTE Confidence: 0.9222073612

 $01{:}02{:}14.759 \dashrightarrow 01{:}02{:}16.195$ neurons representing this reward

NOTE Confidence: 0.9222073612

01:02:16.195 --> 01:02:18.270 prediction error in the same way,

NOTE Confidence: 0.9222073612

 $01:02:18.270 \longrightarrow 01:02:20.878$ so it's a it's a signal can built

 $01:02:20.878 \longrightarrow 01:02:24.006$ here in the VP most likely by

NOTE Confidence: 0.9222073612

 $01{:}02{:}24.006 \dashrightarrow 01{:}02{:}25.800$ integrating various inputs important

NOTE Confidence: 0.9222073612

01:02:25.800 --> 01:02:28.320 new work from Megan Creed's lab at

NOTE Confidence: 0.9222073612

01:02:28.320 --> 01:02:30.753 Saint Louis in Saint Louis showed

NOTE Confidence: 0.9222073612

 $01:02:30.753 \longrightarrow 01:02:32.808$ that projections from the ventral

NOTE Confidence: 0.9222073612

 $01{:}02{:}32.808 \dashrightarrow 01{:}02{:}34.577$ pallidum back to the nucleus.

NOTE Confidence: 0.9222073612

01:02:34.580 --> 01:02:35.834 Incumbents in fact,

NOTE Confidence: 0.9222073612

 $01:02:35.834 \longrightarrow 01:02:38.342$ might be really important when animals

NOTE Confidence: 0.9222073612

 $01{:}02{:}38.342 \dashrightarrow 01{:}02{:}40.719$ are making decisions to consume.

NOTE Confidence: 0.9222073612

01:02:40.720 --> 01:02:41.530 Rewards so,

NOTE Confidence: 0.9222073612

 $01{:}02{:}41.530 \dashrightarrow 01{:}02{:}44.365$ so the VP has a really interesting

NOTE Confidence: 0.9222073612

 $01:02:44.365 \longrightarrow 01:02:45.956$ relationship with the rest

NOTE Confidence: 0.9222073612

01:02:45.956 --> 01:02:47.444 of the reward circuitry,

NOTE Confidence: 0.9222073612

01:02:47.450 --> 01:02:50.346 their inputs to to BTI and VTA projects,

NOTE Confidence: 0.9222073612

01:02:50.350 --> 01:02:51.208 to ventral pallidum,

NOTE Confidence: 0.9222073612

01:02:51.208 --> 01:02:51.780 and so,

 $01:02:51.780 \longrightarrow 01:02:53.916$ so how these signals that we

NOTE Confidence: 0.9222073612

 $01{:}02{:}53.916 \dashrightarrow 01{:}02{:}56.095$ identified fit in with the rest

NOTE Confidence: 0.9222073612

 $01:02:56.095 \longrightarrow 01:02:58.147$ of the activity of the reward

NOTE Confidence: 0.9222073612

 $01:02:58.147 \longrightarrow 01:03:00.229$ circuit is a really important.

NOTE Confidence: 0.9222073612

01:03:00.230 --> 01:03:02.372 Future direction as well as trying

NOTE Confidence: 0.9222073612

 $01:03:02.372 \longrightarrow 01:03:04.747$ to map how the circuit response

NOTE Confidence: 0.9222073612

01:03:04.747 --> 01:03:07.369 to natural reward with how it

NOTE Confidence: 0.9222073612

 $01:03:07.369 \longrightarrow 01:03:09.409$ might respond to drug reward.

NOTE Confidence: 0.9222073612

01:03:09.410 --> 01:03:10.454 Because, as mentioned,

NOTE Confidence: 0.9222073612

 $01:03:10.454 \longrightarrow 01:03:12.890$ your eventual goal is to try to

NOTE Confidence: 0.9222073612

 $01:03:12.963 \longrightarrow 01:03:14.319$ understand these interactive

NOTE Confidence: 0.9222073612

 $01:03:14.319 \longrightarrow 01:03:16.579$ processes and how they modulate

NOTE Confidence: 0.9222073612

 $01:03:16.579 \longrightarrow 01:03:18.969$ in humans are seeking of things.

NOTE Confidence: 0.9222073612

 $01:03:18.970 \longrightarrow 01:03:21.721$ We should seek our food rewards and

NOTE Confidence: 0.9222073612

 $01:03:21.721 \longrightarrow 01:03:24.690$ think and seeking of rewards that in

 $01:03:24.690 \longrightarrow 01:03:28.370$ some individuals can become unhealthy.

NOTE Confidence: 0.9222073612

 $01:03:28.370 \longrightarrow 01:03:30.750$ So I think I'll I'll stop there

NOTE Confidence: 0.9222073612

 $01:03:30.750 \longrightarrow 01:03:32.250$ with just thinking again.

NOTE Confidence: 0.9222073612

 $01:03:32.250 \longrightarrow 01:03:35.060$ The lab members that participated

NOTE Confidence: 0.9222073612

 $01:03:35.060 \longrightarrow 01:03:36.672$ in this work,

NOTE Confidence: 0.9222073612

01:03:36.672 --> 01:03:40.046 and I identified David and Jocelyn early.

NOTE Confidence: 0.9222073612

 $01:03:40.050 \longrightarrow 01:03:42.206$ They're really the main drivers of this.

NOTE Confidence: 0.9222073612

01:03:42.210 --> 01:03:43.974 I also showed Jude members of

NOTE Confidence: 0.9222073612

 $01:03:43.974 \longrightarrow 01:03:45.720$ Jeremiah coincide that were important,

NOTE Confidence: 0.9222073612

 $01:03:45.720 \longrightarrow 01:03:47.856$ but I want to thank the lab in

NOTE Confidence: 0.9222073612

 $01:03:47.856 \longrightarrow 01:03:50.019$ general for all of their input for

NOTE Confidence: 0.9222073612

 $01:03:50.019 \longrightarrow 01:03:52.014$ this work and lab meetings and

NOTE Confidence: 0.9222073612

01:03:52.014 --> 01:03:53.859 and helping one another conduct

NOTE Confidence: 0.9222073612

 $01:03:53.859 \longrightarrow 01:03:55.786$ all of the experiments that I

NOTE Confidence: 0.9222073612

01:03:55.786 --> 01:03:57.124 want to thank funding from NIH,

NOTE Confidence: 0.9222073612

01:03:57.130 --> 01:03:57.670 of course,

 $01:03:57.670 \longrightarrow 01:03:58.750$ and I want to.

NOTE Confidence: 0.9222073612

 $01:03:58.750 \longrightarrow 01:04:01.123$ Thank you all very much for giving

NOTE Confidence: 0.9222073612

 $01:04:01.123 \longrightarrow 01:04:03.303$ me the opportunity to talk about

NOTE Confidence: 0.9222073612

 $01:04:03.303 \longrightarrow 01:04:04.763$ this basic neuroscience research

NOTE Confidence: 0.9222073612

 $01:04:04.763 \longrightarrow 01:04:07.202$ and I hope it gives us all some

NOTE Confidence: 0.9222073612

 $01:04:07.202 \longrightarrow 01:04:09.337$ ideas about how we can think about

NOTE Confidence: 0.9222073612

01:04:09.337 --> 01:04:11.027 basic nice neuroscience work and

NOTE Confidence: 0.9222073612

01:04:11.027 --> 01:04:13.730 how it can tell us about the human

NOTE Confidence: 0.9222073612

01:04:13.730 --> 01:04:15.650 condition and how we doing basic

NOTE Confidence: 0.9222073612

 $01{:}04{:}15.650 \dashrightarrow 01{:}04{:}17.768$ neuroscience work can learn and shape

NOTE Confidence: 0.9222073612

 $01:04:17.768 \longrightarrow 01:04:20.750$ what we do based on the human condition.

NOTE Confidence: 0.9222073612

 $01:04:20.750 \longrightarrow 01:04:21.670$ So thanks very much.

NOTE Confidence: 0.913032607222222

 $01:04:24.180 \longrightarrow 01:04:26.520$ Thank you so much, I'm really

NOTE Confidence: 0.913032607222222

 $01{:}04{:}26.520 \dashrightarrow 01{:}04{:}29.205$ enjoyed that and I have encouraged

NOTE Confidence: 0.913032607222222

01:04:29.205 --> 01:04:32.313 the trainees to ask questions first

 $01:04:32.320 \longrightarrow 01:04:34.540$ if there's any trainees out there.

NOTE Confidence: 0.913032607222222

01:04:34.540 --> 01:04:37.366 Doctor Taylor is a trainee of

NOTE Confidence: 0.913032607222222

01:04:37.366 --> 01:04:39.779 course lifelong but may not qualify,

NOTE Confidence: 0.913032607222222

01:04:39.779 --> 01:04:43.280 so I would like to start with with a trainee,

NOTE Confidence: 0.913032607222222

 $01:04:43.280 \longrightarrow 01:04:45.952$ but if not we can we can get

NOTE Confidence: 0.913032607222222

 $01{:}04{:}45.952 \dashrightarrow 01{:}04{:}48.090$ to questions from UN trainees.

NOTE Confidence: 0.6206374

 $01:04:50.780 \longrightarrow 01:04:52.230$ Doctor Taylor's training

NOTE Confidence: 0.84640145

01:04:54.610 --> 01:04:56.856 all right? Well then that seems appropriate.

NOTE Confidence: 0.87042520375

 $01:04:56.860 \longrightarrow 01:04:57.350$ Doctor Taylor.

NOTE Confidence: 0.87042520375

01:04:57.350 --> 01:04:58.820 Why don't you kick us off?

NOTE Confidence: 0.87042520375

01:04:58.820 --> 01:05:03.201 I have questions too. Sorry, I come.

NOTE Confidence: 0.87042520375

01:05:03.201 --> 01:05:05.868 If a trainee wants to interrupt me,

NOTE Confidence: 0.87042520375

 $01:05:05.870 \longrightarrow 01:05:07.520$ please go ahead.

NOTE Confidence: 0.87042520375

 $01:05:07.520 \longrightarrow 01:05:11.925$ That was a beautiful talk as always Patricia.

NOTE Confidence: 0.87042520375

 $01:05:11.925 \longrightarrow 01:05:15.810$ So I have a question which is

NOTE Confidence: 0.87042520375

01:05:15.810 --> 01:05:20.374 sort of how dynamic do you think

 $01{:}05{:}20.374 \dashrightarrow 01{:}05{:}24.160$ these VP responses are in that.

NOTE Confidence: 0.87042520375

 $01{:}05{:}24.160 \dashrightarrow 01{:}05{:}27.730$ I wonder whether you would see

NOTE Confidence: 0.87042520375

 $01:05:27.730 \longrightarrow 01:05:30.522$ similar VP signals related to

NOTE Confidence: 0.87042520375

 $01:05:30.522 \longrightarrow 01:05:33.460$ expectation and prediction error.

NOTE Confidence: 0.87042520375

 $01:05:33.460 \longrightarrow 01:05:36.484$ If you in your experiment initially

NOTE Confidence: 0.87042520375

 $01:05:36.484 \longrightarrow 01:05:38.500$ looked at sucrose preference

NOTE Confidence: 0.87042520375

 $01:05:38.586 \longrightarrow 01:05:41.441$ compared to something that was

NOTE Confidence: 0.87042520375

01:05:41.441 --> 01:05:43.154 actually slightly aversive,

NOTE Confidence: 0.87042520375

01:05:43.160 --> 01:05:45.660 like a salt solution,

NOTE Confidence: 0.87042520375

 $01:05:45.660 \longrightarrow 01:05:48.785$ but then made the animals

NOTE Confidence: 0.87042520375

01:05:48.790 --> 01:05:53.230 physiologically salt induced the salt,

NOTE Confidence: 0.87042520375

 $01:05:53.230 \longrightarrow 01:05:56.119$ a salt state for that reward.

NOTE Confidence: 0.87042520375

 $01{:}05{:}56.120 \dashrightarrow 01{:}06{:}00.026$ Would you see the VP neurons suddenly

NOTE Confidence: 0.87042520375

 $01{:}06{:}00.026 \dashrightarrow 01{:}06{:}03.209$ switch over to tracking the?

NOTE Confidence: 0.87042520375

01:06:03.210 --> 01:06:05.058 The salt, uhm?

 $01:06:05.500 \longrightarrow 01:06:08.540$ Yeah, yeah, I I think so and so.

NOTE Confidence: 0.9554099425

01:06:08.540 --> 01:06:11.840 So although this is a history

NOTE Confidence: 0.9554099425

01:06:11.840 --> 01:06:13.630 dependent signal that follows this

NOTE Confidence: 0.9554099425

01:06:13.689 --> 01:06:15.915 reward prediction error kind of model,

NOTE Confidence: 0.9554099425

 $01:06:15.920 \longrightarrow 01:06:18.768$ we think it's not model free but it's

NOTE Confidence: 0.9554099425

01:06:18.768 --> 01:06:21.414 more model based in that the subject

NOTE Confidence: 0.9554099425

 $01:06:21.414 \longrightarrow 01:06:25.740$ can update it on the fly and so so you

NOTE Confidence: 0.9554099425

 $01{:}06{:}25.740 \dashrightarrow 01{:}06{:}28.787$ know in in this procedure animals start

NOTE Confidence: 0.9554099425

 $01{:}06{:}28.787 \longrightarrow 01{:}06{:}32.224$ each day thirsty and then become sated.

NOTE Confidence: 0.9554099425

 $01:06:32.230 \longrightarrow 01:06:34.240$ Their response to in a different

NOTE Confidence: 0.9554099425

 $01{:}06{:}34.240 \dashrightarrow 01{:}06{:}36.000$ procedure where there's a water

NOTE Confidence: 0.9554099425

 $01:06:36.000 \longrightarrow 01:06:38.178$ predictive cue their response to the

NOTE Confidence: 0.9554099425

 $01:06:38.178 \longrightarrow 01:06:40.218$ water predictive cue is very high

NOTE Confidence: 0.9554099425

 $01{:}06{:}40.218 \dashrightarrow 01{:}06{:}41.988$ at the beginning of each session,

NOTE Confidence: 0.9554099425

01:06:41.990 --> 01:06:44.132 even though it's very low by the

NOTE Confidence: 0.9554099425

 $01:06:44.132 \longrightarrow 01:06:46.522$ end of the session when thirst is

 $01:06:46.522 \longrightarrow 01:06:48.262$ is no longer a drive.

NOTE Confidence: 0.9554099425

 $01:06:48.270 \longrightarrow 01:06:48.936$ So so there.

NOTE Confidence: 0.9554099425

01:06:48.936 --> 01:06:50.857 I'm not sure if I'm getting to exactly

NOTE Confidence: 0.9554099425

 $01:06:50.857 \longrightarrow 01:06:53.319$ what what your question was, but it's a.

NOTE Confidence: 0.9554099425

01:06:53.319 --> 01:06:55.137 It's a super dynamic system and

NOTE Confidence: 0.9554099425

01:06:55.137 --> 01:06:56.920 I think immediately impacted

NOTE Confidence: 0.9554099425

 $01:06:56.920 \longrightarrow 01:06:59.060$ by the animals expectations,

NOTE Confidence: 0.9554099425

 $01:06:59.060 \longrightarrow 01:07:02.525$ and it doesn't necessarily have to accrue.

NOTE Confidence: 0.9554099425

01:07:02.530 --> 01:07:04.094 Overtime it's impacted by

NOTE Confidence: 0.9554099425

 $01:07:04.094 \longrightarrow 01:07:05.658$ what happens over time,

NOTE Confidence: 0.9554099425

 $01:07:05.660 \longrightarrow 01:07:08.372$ but also can be directed by a more

NOTE Confidence: 0.9554099425

 $01:07:08.372 \longrightarrow 01:07:12.214$ sort of cognitive, a goal directed.

NOTE Confidence: 0.9554099425

 $01:07:12.214 \longrightarrow 01:07:13.730$ Uh, evaluation.

NOTE Confidence: 0.6166265

01:07:15.900 --> 01:07:16.810 List Europe.

NOTE Confidence: 0.929409916

 $01:07:18.820 \longrightarrow 01:07:20.428$ Thank you for that amazing talk

 $01:07:20.428 \longrightarrow 01:07:21.919$ and I'm curious 'cause we've

NOTE Confidence: 0.929409916

 $01{:}07{:}21.919 \dashrightarrow 01{:}07{:}23.387$ been talking about expectation,

NOTE Confidence: 0.929409916

 $01:07:23.390 \longrightarrow 01:07:24.696$ but most of the signal that

NOTE Confidence: 0.929409916

 $01:07:24.696 \longrightarrow 01:07:26.260$ you've been presenting is

NOTE Confidence: 0.929409916

 $01:07:26.260 \longrightarrow 01:07:28.080$ during the reward consumption.

NOTE Confidence: 0.929409916

 $01:07:28.080 \longrightarrow 01:07:29.627$ So do you see anything when the

NOTE Confidence: 0.929409916

01:07:29.627 --> 01:07:31.244 queue is present that indicates the

NOTE Confidence: 0.929409916

01:07:31.244 --> 01:07:32.744 outcome that they're expecting before

NOTE Confidence: 0.929409916

 $01{:}07{:}32.744 \dashrightarrow 01{:}07{:}34.408$ they can compute the prediction

NOTE Confidence: 0.87532847

01:07:34.420 --> 01:07:36.700 error? Yeah, that's a great question.

NOTE Confidence: 0.87532847

 $01{:}07{:}36.700 \dashrightarrow 01{:}07{:}38.518$ So in this particular set of

NOTE Confidence: 0.87532847

01:07:38.518 --> 01:07:40.560 studies that I told you about

NOTE Confidence: 0.87532847

 $01:07:40.560 \longrightarrow 01:07:42.455$ the queues were not informative.

NOTE Confidence: 0.87532847

 $01:07:42.460 \longrightarrow 01:07:44.580$ As far as the identity of the reward,

NOTE Confidence: 0.87532847

 $01:07:44.580 \longrightarrow 01:07:47.015$ so we didn't find very

NOTE Confidence: 0.87532847

 $01{:}07{:}47.015 \dashrightarrow 01{:}07{:}48.476$ much expectations signal.

 $01:07:48.480 \longrightarrow 01:07:49.593$ In those cues,

NOTE Confidence: 0.87532847

 $01:07:49.593 \longrightarrow 01:07:52.190$ but David ran a variation in this

NOTE Confidence: 0.87532847

 $01:07:52.268 \longrightarrow 01:07:54.623$ dynamic thirst task where there

NOTE Confidence: 0.87532847

 $01:07:54.623 \longrightarrow 01:07:56.978$ were accused that came before

NOTE Confidence: 0.87532847

 $01:07:57.061 \longrightarrow 01:07:59.490$ each of the rewards in the forest

NOTE Confidence: 0.87532847

 $01:07:59.490 \longrightarrow 01:08:01.824$ trials of a water Q&A sucrose Q,

NOTE Confidence: 0.87532847

 $01:08:01.824 \longrightarrow 01:08:04.241$ and in that case the reward prediction

NOTE Confidence: 0.87532847

 $01:08:04.241 \longrightarrow 01:08:05.925$ error like signaling transferred

NOTE Confidence: 0.87532847

 $01{:}08{:}05.925 \dashrightarrow 01{:}08{:}09.039$ to the queue as you would predict,

NOTE Confidence: 0.87532847

 $01:08:09.040 \longrightarrow 01:08:11.350$ and so that expectation related

NOTE Confidence: 0.87532847

 $01:08:11.350 \longrightarrow 01:08:13.440$ response was mostly there in

NOTE Confidence: 0.87532847

 $01:08:13.440 \longrightarrow 01:08:15.890$ the in the way that the animal

NOTE Confidence: 0.87532847

 $01:08:15.890 \longrightarrow 01:08:17.238$ responded to the queue.

NOTE Confidence: 0.87532847

 $01:08:17.240 \longrightarrow 01:08:20.000$ So that's that's a great question.

NOTE Confidence: 0.87532847 01:08:20.000 --> 01:08:20.530 Al

 $01:08:22.060 \longrightarrow 01:08:23.796$ hi, thanks so much for the talk.

NOTE Confidence: 0.913256071428571

 $01{:}08{:}23.800 \dashrightarrow 01{:}08{:}25.660$ I was really struck by your result

NOTE Confidence: 0.913256071428571

01:08:25.660 --> 01:08:28.128 that about heterogeneity in terms

NOTE Confidence: 0.913256071428571

01:08:28.128 --> 01:08:30.950 of value versus RPE signals in

NOTE Confidence: 0.913256071428571

 $01:08:30.950 \longrightarrow 01:08:33.190$ the ventral pallidum, especially

NOTE Confidence: 0.933677535

 $01:08:33.200 \longrightarrow 01:08:35.615$ given this kind of ongoing

NOTE Confidence: 0.933677535

 $01:08:35.615 \longrightarrow 01:08:37.594$ discussion about whether a Cummins

NOTE Confidence: 0.933677535

01:08:37.594 --> 01:08:39.106 concentration of dopamine represents

NOTE Confidence: 0.716693086

01:08:39.120 --> 01:08:41.260 a value or RPE signal,

NOTE Confidence: 0.716693086

 $01:08:41.260 \longrightarrow 01:08:44.500$ and in particular. Some

NOTE Confidence: 0.774157

 $01{:}08{:}44.510 \dashrightarrow 01{:}08{:}45.820$ recent work of Sam Gershman,

NOTE Confidence: 0.774157

01:08:45.820 --> 01:08:47.338 and now, which is showing that

NOTE Confidence: 0.774157

 $01:08:47.338 \longrightarrow 01:08:48.770$ you can maybe reconcile these

NOTE Confidence: 0.774157

 $01:08:48.770 \longrightarrow 01:08:51.542$ approaches by having a kind of RP

NOTE Confidence: 0.774157

 $01:08:51.542 \longrightarrow 01:08:53.432$ signal with sensory feedback that

NOTE Confidence: 0.89370191

 $01:08:53.550 \longrightarrow 01:08:55.510$ looks like a value signal.

 $01:08:55.510 \longrightarrow 01:08:57.534$ And so I guess I was just served as an

NOTE Confidence: 0.883696619090909

 $01:08:57.550 \longrightarrow 01:08:58.609$ open ended question.

NOTE Confidence: 0.883696619090909

01:08:58.609 --> 01:09:00.727 Wondering whether you think that this

NOTE Confidence: 0.883696619090909

 $01:09:00.727 \longrightarrow 01:09:02.640$ heterogeneity in the ventral pallidum

NOTE Confidence: 0.883696619090909

 $01:09:02.640 \longrightarrow 01:09:05.050$ might somehow either resolve this

NOTE Confidence: 0.883696619090909

01:09:05.050 --> 01:09:07.264 discrepancy or correspond to as well

NOTE Confidence: 0.883696619090909

 $01:09:07.264 \longrightarrow 01:09:10.100$ the concentration of dopamine dynamics.

NOTE Confidence: 0.883696619090909

01:09:10.100 --> 01:09:12.308 Yeah, that's a great question and

NOTE Confidence: 0.883696619090909

 $01:09:12.308 \longrightarrow 01:09:14.869$ my short answer is is I I don't

NOTE Confidence: 0.883696619090909

 $01:09:14.869 \longrightarrow 01:09:17.490$ know so it is true that this system

NOTE Confidence: 0.883696619090909

01:09:17.490 --> 01:09:20.035 is highly interconnected with the

NOTE Confidence: 0.883696619090909

 $01:09:20.035 \longrightarrow 01:09:22.515$ canonical dopamine neurons in the VTA.

NOTE Confidence: 0.883696619090909

 $01{:}09{:}22.515 \dashrightarrow 01{:}09{:}25.070$ So VP neurons project back to the VTA,

NOTE Confidence: 0.883696619090909

 $01:09:25.070 \longrightarrow 01:09:27.982$ both to dopamine neurons but also to

NOTE Confidence: 0.883696619090909

 $01:09:27.982 \longrightarrow 01:09:30.169$ the GABA interneurons and the VTA.

01:09:30.170 --> 01:09:32.626 Dopamine neurons do send a projection to VP,

NOTE Confidence: 0.883696619090909

 $01{:}09{:}32.630 \dashrightarrow 01{:}09{:}34.688$ so there's some interaction in the

NOTE Confidence: 0.883696619090909

 $01:09:34.688 \longrightarrow 01:09:36.732$ creation of these kinds of dopamine

NOTE Confidence: 0.883696619090909

 $01:09:36.732 \longrightarrow 01:09:39.148$ signals that that we think about as far

NOTE Confidence: 0.883696619090909

 $01:09:39.210 \longrightarrow 01:09:41.664$ as our PE and. What exactly are they?

NOTE Confidence: 0.883696619090909

01:09:41.664 --> 01:09:44.100 Are they communicating to the incumbents?

NOTE Confidence: 0.883696619090909 01:09:44.100 --> 01:09:44.835 Uh, so I. NOTE Confidence: 0.883696619090909

 $01:09:44.835 \longrightarrow 01:09:46.550$ I don't know how that's all gonna

NOTE Confidence: 0.883696619090909

 $01{:}09{:}46.607 \dashrightarrow 01{:}09{:}48.607$ work out as far as trying to see if

NOTE Confidence: 0.883696619090909

01:09:48.607 --> 01:09:50.468 there's a separable value signal or

NOTE Confidence: 0.883696619090909

 $01:09:50.468 \longrightarrow 01:09:52.804$ whether it's really going to be able

NOTE Confidence: 0.883696619090909

 $01:09:52.804 \longrightarrow 01:09:55.876$ to be understood all as a readout of

NOTE Confidence: 0.883696619090909

 $01:09:55.876 \longrightarrow 01:09:58.929$ online changes in what the animal

NOTE Confidence: 0.883696619090909

01:09:58.929 --> 01:10:00.993 is actually actually receiving.

NOTE Confidence: 0.883696619090909

 $01:10:01.000 \longrightarrow 01:10:02.656$ So I think that's something that's

NOTE Confidence: 0.883696619090909

 $01:10:02.656 \longrightarrow 01:10:04.020$ still left to work out.

01:10:04.020 --> 01:10:07.060 So really interesting sort of not a problem,

NOTE Confidence: 0.883696619090909

 $01:10:07.060 \longrightarrow 01:10:08.995$ and the communication between the

NOTE Confidence: 0.883696619090909

 $01:10:08.995 \longrightarrow 01:10:10.543$ VP and the incumbents.

NOTE Confidence: 0.883696619090909

 $01:10:10.550 \longrightarrow 01:10:12.244$ Is also going to be a factor.

NOTE Confidence: 0.883696619090909

 $01:10:12.250 \longrightarrow 01:10:13.849$ Presumably so, uhm.

NOTE Confidence: 0.883696619090909

01:10:13.849 --> 01:10:17.580 I like the direction of your question

NOTE Confidence: 0.883696619090909

 $01:10:17.675 \longrightarrow 01:10:21.766$ because it forces these results have

NOTE Confidence: 0.883696619090909

 $01{:}10{:}21.766 \dashrightarrow 01{:}10{:}24.756$ forced me and all this forces us to

NOTE Confidence: 0.883696619090909

 $01:10:24.756 \longrightarrow 01:10:26.389$ not think of reward prediction error

NOTE Confidence: 0.883696619090909

 $01:10:26.389 \longrightarrow 01:10:28.791$ is just here in this group of dopamine

NOTE Confidence: 0.883696619090909

01:10:28.791 --> 01:10:30.945 neurons and then medium spiny neurons.

NOTE Confidence: 0.883696619090909

 $01:10:30.950 \longrightarrow 01:10:32.900$ Maybe all are transmitting expected value

NOTE Confidence: 0.883696619090909

 $01:10:32.900 \longrightarrow 01:10:35.239$ and this area does this in this area.

NOTE Confidence: 0.883696619090909

01:10:35.240 --> 01:10:37.274 Does that and instead you know

NOTE Confidence: 0.883696619090909

 $01:10:37.274 \longrightarrow 01:10:39.292$ these systems are all interconnected

 $01:10:39.292 \longrightarrow 01:10:41.600$ and these variables seem to be

NOTE Confidence: 0.883696619090909

 $01{:}10{:}41.600 \dashrightarrow 01{:}10{:}43.320$ represented to greater or lesser

NOTE Confidence: 0.883696619090909

01:10:43.381 --> 01:10:45.189 degrees throughout the circuit.

NOTE Confidence: 0.883696619090909

01:10:45.190 --> 01:10:48.270 So beautiful work from the U2 lab,

NOTE Confidence: 0.883696619090909 01:10:48.270 --> 01:10:49.076 in fact, NOTE Confidence: 0.883696619090909

 $01:10:49.076 \longrightarrow 01:10:51.494$ has shown has shown that really

NOTE Confidence: 0.883696619090909

01:10:51.494 --> 01:10:52.890 painstaking recording studies,

NOTE Confidence: 0.883696619090909

 $01:10:52.890 \longrightarrow 01:10:55.886$ so it's it's going to be harder

NOTE Confidence: 0.883696619090909

 $01:10:55.886 \longrightarrow 01:10:58.220$ to figure out than then.

NOTE Confidence: 0.883696619090909

 $01:10:58.220 \longrightarrow 01:11:00.596$ I would dream when we want to make a

NOTE Confidence: 0.883696619090909

 $01{:}11{:}00.596 \dashrightarrow 01{:}11{:}02.800$ nice model where everything is only

NOTE Confidence: 0.883696619090909

 $01:11:02.800 \longrightarrow 01:11:05.076$ just very very separable and in a

NOTE Confidence: 0.883696619090909

 $01:11:05.076 \longrightarrow 01:11:07.559$ separate kind of neuron and separate place.

NOTE Confidence: 0.883696619090909

01:11:07.560 --> 01:11:09.180 But that's a nice challenge.

NOTE Confidence: 0.883696619090909

01:11:09.180 --> 01:11:10.764 There's work to do it for the future.

NOTE Confidence: 0.8874942125

01:11:12.700 --> 01:11:14.716 Ralph, I'm going to ask a quick question

 $01:11:14.716 \longrightarrow 01:11:16.617$ from the chat and then over to you.

NOTE Confidence: 0.8874942125

01:11:16.620 --> 01:11:19.602 Media Naseer says rats prefer water

NOTE Confidence: 0.8874942125

 $01:11:19.602 \longrightarrow 01:11:21.590$ over sucrose when dehydrated.

NOTE Confidence: 0.8874942125

01:11:21.590 --> 01:11:23.620 However, some people prefer sugary

NOTE Confidence: 0.8874942125

 $01:11:23.620 \longrightarrow 01:11:26.250$ soft drinks over water when thirsty.

NOTE Confidence: 0.8874942125

 $01:11:26.250 \longrightarrow 01:11:27.550$ Is there a mechanistic

NOTE Confidence: 0.8874942125

01:11:27.550 --> 01:11:28.850 difference in this situation?

NOTE Confidence: 0.92638313625

 $01{:}11{:}30.320 \dashrightarrow 01{:}11{:}32.288$ That's a great question, and one I I

NOTE Confidence: 0.92638313625

01:11:32.288 --> 01:11:34.199 don't know the answer to and I would.

NOTE Confidence: 0.92638313625

 $01:11:34.200 \longrightarrow 01:11:36.816$ I guess I would immediately wonder

NOTE Confidence: 0.92638313625

01:11:36.816 --> 01:11:40.072 about the role of long term experience

NOTE Confidence: 0.92638313625

 $01:11:40.072 \longrightarrow 01:11:43.436$ in humans for going for the sugary

NOTE Confidence: 0.92638313625

 $01{:}11{:}43.436 \dashrightarrow 01{:}11{:}46.248$ soft drink to relieve thirst and

NOTE Confidence: 0.92638313625

 $01:11:46.248 \longrightarrow 01:11:48.576$ and whether we could model that.

NOTE Confidence: 0.92638313625

 $01:11:48.580 \longrightarrow 01:11:51.261$ By the way we expose our rats

 $01:11:51.261 \longrightarrow 01:11:53.130$ to these different rewards.

NOTE Confidence: 0.92638313625

 $01:11:53.130 \longrightarrow 01:11:54.798$ Overtime probably someone working

NOTE Confidence: 0.92638313625

 $01:11:54.798 \longrightarrow 01:11:56.883$ more in the nutrition field

NOTE Confidence: 0.92638313625

01:11:56.883 --> 01:11:58.737 maybe has already done that,

NOTE Confidence: 0.92638313625

 $01:11:58.740 \longrightarrow 01:12:00.060$ so that answer might be known

NOTE Confidence: 0.92638313625

01:12:00.060 --> 01:12:01.369 and I just don't know it.

NOTE Confidence: 0.92638313625

 $01:12:01.370 \longrightarrow 01:12:04.136$ I think that's a great question.

NOTE Confidence: 0.92638313625

01:12:04.140 --> 01:12:06.138 Ralph, go ahead.

NOTE Confidence: 0.92638313625

 $01:12:06.140 \longrightarrow 01:12:07.704$ Trisha really like these

NOTE Confidence: 0.92638313625

 $01:12:07.704 \longrightarrow 01:12:09.130$ experiments and great data.

NOTE Confidence: 0.8771184825

 $01:12:09.400 \longrightarrow 01:12:11.128$ I have a question

NOTE Confidence: 0.8797689

 $01:12:11.140 \longrightarrow 01:12:14.204$ I guess about the maltodextrin and and what.

NOTE Confidence: 0.896369645

01:12:14.870 --> 01:12:16.670 Is this is this purely sensory?

NOTE Confidence: 0.896369645

01:12:16.670 --> 01:12:17.622 Is it post ingestive?

NOTE Confidence: 0.896369645

01:12:17.622 --> 01:12:19.340 I know the answer is probably both,

NOTE Confidence: 0.896369645

 $01:12:19.340 \longrightarrow 01:12:21.115$ but I guess I'm interested

01:12:21.115 --> 01:12:22.046 because maltodextrins are

NOTE Confidence: 0.896369645

01:12:22.046 --> 01:12:23.126 sort of a tricky thing.

NOTE Confidence: 0.896369645

01:12:23.130 --> 01:12:24.170 It's not very sweet,

NOTE Confidence: 0.896369645

 $01:12:24.170 \longrightarrow 01:12:25.386$ but of course you can't detect

NOTE Confidence: 0.896369645

 $01:12:25.386 \longrightarrow 01:12:26.850$ it at high concentrations.

NOTE Confidence: 0.896369645

 $01:12:26.850 \longrightarrow 01:12:27.540$ I can't remember

NOTE Confidence: 0.79494786

01:12:27.550 --> 01:12:30.286 from from David early paper how

NOTE Confidence: 0.79494786

 $01:12:30.290 \longrightarrow 01:12:31.850$ high it was, and if you know

NOTE Confidence: 0.79494786

 $01:12:31.850 \longrightarrow 01:12:33.500$ that Jamaican detector at least

NOTE Confidence: 0.870325631764706

 $01:12:33.510 \longrightarrow 01:12:35.124$ taste it. And of course some

NOTE Confidence: 0.870325631764706

 $01{:}12{:}35.124 \dashrightarrow 01{:}12{:}36.547$ of the signals you're seeing

NOTE Confidence: 0.870325631764706

 $01:12:36.547 \longrightarrow 01:12:38.299$ or so fast is clearly something

NOTE Confidence: 0.841295649

 $01{:}12{:}38.310 \dashrightarrow 01{:}12{:}40.494$ century. But I wonder if over the session

NOTE Confidence: 0.841295649

 $01:12:40.494 \longrightarrow 01:12:42.140$ they're learning. It has a really

NOTE Confidence: 0.86341530375

 $01:12:42.270 \longrightarrow 01:12:44.150$ it gets broken down so quickly into glucose.

 $01:12:44.150 \longrightarrow 01:12:45.090$ It's guys seeing indexes.

NOTE Confidence: 0.86341530375

 $01:12:45.090 \longrightarrow 01:12:46.582$ Is higher than sucrose, right?

NOTE Confidence: 0.86341530375

 $01:12:46.582 \longrightarrow 01:12:48.444$ So it should be really fast post

NOTE Confidence: 0.86341530375

 $01:12:48.450 \longrightarrow 01:12:50.418$ ingestive signals and so I wonder

NOTE Confidence: 0.86341530375

 $01:12:50.418 \longrightarrow 01:12:52.750$ how you how you think about that.

NOTE Confidence: 0.838073193076923

01:12:52.750 --> 01:12:55.690 I think, uh, so David more than

NOTE Confidence: 0.838073193076923

01:12:55.690 --> 01:12:58.829 me really did the important work.

NOTE Confidence: 0.838073193076923

 $01:12:58.830 \longrightarrow 01:13:00.342$ All you train is out there of

NOTE Confidence: 0.838073193076923

 $01{:}13{:}00.342 \dashrightarrow 01{:}13{:}01.978$ looking at all the old literature,

NOTE Confidence: 0.838073193076923

 $01:13:01.980 \longrightarrow 01:13:04.085$ including all the old animal

NOTE Confidence: 0.838073193076923

 $01{:}13{:}04.085 \dashrightarrow 01{:}13{:}05.769$ behavior literature where people

NOTE Confidence: 0.838073193076923

01:13:05.769 --> 01:13:08.242 have done a lot of work comparing

NOTE Confidence: 0.838073193076923

 $01:13:08.242 \longrightarrow 01:13:10.340$ tastants and looking at preference.

NOTE Confidence: 0.838073193076923

 $01:13:10.340 \longrightarrow 01:13:12.116$ Ways to measure preference in assessed

NOTE Confidence: 0.838073193076923

 $01:13:12.116 \longrightarrow 01:13:13.490$ preference in rodents and so.

NOTE Confidence: 0.838073193076923

 $01:13:13.490 \longrightarrow 01:13:15.594$ So he he looked carefully at that when

 $01:13:15.594 \longrightarrow 01:13:17.807$ he chose maltodextrin as a comparison.

NOTE Confidence: 0.838073193076923

01:13:17.810 --> 01:13:19.714 So I think it's a good comparison,

NOTE Confidence: 0.838073193076923

 $01:13:19.720 \longrightarrow 01:13:20.551$ but we can't.

NOTE Confidence: 0.838073193076923

 $01:13:20.551 \longrightarrow 01:13:22.213$ It's very difficult to get two

NOTE Confidence: 0.838073193076923

 $01:13:22.213 \longrightarrow 01:13:23.957$ things that are exactly the same,

NOTE Confidence: 0.838073193076923

01:13:23.960 --> 01:13:26.336 but different, and so in that you know

NOTE Confidence: 0.838073193076923

01:13:26.336 --> 01:13:28.543 in fact that's impossible, right?

NOTE Confidence: 0.838073193076923

 $01:13:28.543 \longrightarrow 01:13:31.527$ So one thing I can say is that,

NOTE Confidence: 0.838073193076923

 $01:13:31.530 \longrightarrow 01:13:33.245$ at least for the signal in the

NOTE Confidence: 0.838073193076923

01:13:33.245 --> 01:13:34.770 way that David's looking at it,

NOTE Confidence: 0.838073193076923

 $01{:}13{:}34.770 \dashrightarrow 01{:}13{:}38.144$ there isn't a change through the session

NOTE Confidence: 0.838073193076923

 $01:13:38.150 \longrightarrow 01:13:40.560$ in the circumstance when animals.

NOTE Confidence: 0.838073193076923

 $01:13:40.560 \longrightarrow 01:13:42.294$ Thirsty so there.

NOTE Confidence: 0.838073193076923

01:13:42.294 --> 01:13:43.450 Waters stated,

NOTE Confidence: 0.838073193076923

01:13:43.450 --> 01:13:45.235 and they're just choosing between

 $01:13:45.235 \longrightarrow 01:13:47.636$ the two rewards that might show that

NOTE Confidence: 0.838073193076923

 $01:13:47.636 \longrightarrow 01:13:49.828$ there's a big impact of the way that

NOTE Confidence: 0.838073193076923

 $01:13:49.891 \longrightarrow 01:13:51.961$ that these are is post ingestive

NOTE Confidence: 0.838073193076923

 $01:13:51.961 \longrightarrow 01:13:53.727$ effects of sucrose versus maltodextrin.

NOTE Confidence: 0.838073193076923

 $01:13:53.727 \longrightarrow 01:13:57.120$ So so at least on the face of it,

NOTE Confidence: 0.838073193076923

 $01:13:57.120 \longrightarrow 01:13:59.353$ based on the analysis of this signal

NOTE Confidence: 0.838073193076923

01:13:59.353 --> 01:14:01.220 there during the reward period,

NOTE Confidence: 0.838073193076923

 $01:14:01.220 \longrightarrow 01:14:02.249$ there's nothing obvious.

NOTE Confidence: 0.838073193076923

 $01:14:02.249 \longrightarrow 01:14:04.307$ Maybe there are other ways that

NOTE Confidence: 0.838073193076923

 $01:14:04.307 \longrightarrow 01:14:06.459$ he could look at the signals more

NOTE Confidence: 0.838073193076923

 $01{:}14{:}06.459 \dashrightarrow 01{:}14{:}08.729$ carefully to see if there is feedback,

NOTE Confidence: 0.838073193076923 01:14:08.730 --> 01:14:09.206 because. NOTE Confidence: 0.838073193076923

01:14:09.206 --> 01:14:11.110 And as you know,

NOTE Confidence: 0.838073193076923

 $01:14:11.110 \longrightarrow 01:14:13.054$ you know there's a big literature

NOTE Confidence: 0.838073193076923

01:14:13.054 --> 01:14:17.000 on how post ingestive impacts of

NOTE Confidence: 0.838073193076923

01:14:17.000 --> 01:14:18.984 food impact functioning within

 $01:14:18.984 \longrightarrow 01:14:20.319$ the striedl circuits that that

NOTE Confidence: 0.838073193076923

 $01:14:20.319 \longrightarrow 01:14:21.909$ we all know and love so much.

NOTE Confidence: 0.838073193076923

01:14:21.910 --> 01:14:23.038 So I I would,

NOTE Confidence: 0.838073193076923

 $01:14:23.038 \longrightarrow 01:14:25.590$ I wouldn't say there's not an interaction.

NOTE Confidence: 0.838073193076923

01:14:25.590 --> 01:14:25.890 I.

NOTE Confidence: 0.838073193076923

01:14:25.890 --> 01:14:27.390 I think there probably is,

NOTE Confidence: 0.838073193076923

 $01:14:27.390 \longrightarrow 01:14:28.566$ but at least for the signal

NOTE Confidence: 0.838073193076923

 $01:14:28.566 \longrightarrow 01:14:29.350$ that he's looking at,

NOTE Confidence: 0.838073193076923

01:14:29.350 --> 01:14:32.000 he didn't note anything obvious.

NOTE Confidence: 0.838073193076923

 $01{:}14{:}32.000 \dashrightarrow 01{:}14{:}32.480 \ \mathrm{Text}.$

NOTE Confidence: 0.9806579

01:14:34.550 --> 01:14:36.278 I have a quick question

NOTE Confidence: 0.9806579

 $01:14:36.278 \longrightarrow 01:14:38.188$ or maybe not so quick.

NOTE Confidence: 0.818468543846154

 $01{:}14{:}38.190 \dashrightarrow 01{:}14{:}41.892$ The VP is a surprisingly large

NOTE Confidence: 0.818468543846154

 $01:14:41.892 \longrightarrow 01:14:44.360$ and heterogeneous structure and

NOTE Confidence: 0.818468543846154

 $01:14:44.464 \longrightarrow 01:14:46.792$ so I have a couple of questions.

 $01:14:46.792 \longrightarrow 01:14:48.582$ One is from your recordings.

NOTE Confidence: 0.818468543846154

 $01{:}14{:}48.590 \dashrightarrow 01{:}14{:}52.682$ Can you determine the cell types

NOTE Confidence: 0.818468543846154

01:14:52.682 --> 01:14:55.876 that are responding based on their

NOTE Confidence: 0.818468543846154

01:14:55.876 --> 01:14:58.168 firing patterns or some kind of

NOTE Confidence: 0.818468543846154

01:14:58.168 --> 01:15:00.230 algorithms that let you know about,

NOTE Confidence: 0.818468543846154

 $01:15:00.230 \longrightarrow 01:15:02.686$ let's say cholinergic neurons

NOTE Confidence: 0.818468543846154

 $01:15:02.686 \longrightarrow 01:15:05.756$ versus other types of neurons?

NOTE Confidence: 0.818468543846154

 $01:15:05.760 \longrightarrow 01:15:08.566$ And then here on top and then

NOTE Confidence: 0.818468543846154

 $01{:}15{:}08.566 \dashrightarrow 01{:}15{:}10.706$ the other question is around

NOTE Confidence: 0.818468543846154

 $01:15:10.706 \longrightarrow 01:15:12.898$ a sub sections of the VP.

NOTE Confidence: 0.818468543846154

 $01:15:12.900 \longrightarrow 01:15:14.993$ Whether you see these kinds of responses

NOTE Confidence: 0.818468543846154

01:15:14.993 --> 01:15:16.400 uniformly throughout the structure,

NOTE Confidence: 0.818468543846154

 $01:15:16.400 \longrightarrow 01:15:18.360$ or whether the anterior versus

NOTE Confidence: 0.818468543846154

 $01:15:18.360 \longrightarrow 01:15:20.320$ the posterior is more responsive

NOTE Confidence: 0.818468543846154

 $01:15:20.386 \longrightarrow 01:15:22.130$ to these value measurements.

NOTE Confidence: 0.934005198666667

 $01:15:23.670 \longrightarrow 01:15:25.662$ Those are great questions because it's

01:15:25.662 --> 01:15:28.329 known that as as many of you may know,

NOTE Confidence: 0.934005198666667

 $01:15:28.330 \longrightarrow 01:15:30.130$ the VP contains a lot of

NOTE Confidence: 0.934005198666667

 $01:15:30.130 \longrightarrow 01:15:31.330$ kinds of different neurons.

NOTE Confidence: 0.934005198666667

01:15:31.330 --> 01:15:33.600 It's it's mostly Gabaergic neurons,

NOTE Confidence: 0.934005198666667

 $01:15:33.600 \longrightarrow 01:15:36.180$ but there are neurons that release

NOTE Confidence: 0.934005198666667

 $01:15:36.180 \longrightarrow 01:15:37.470$ glutamate cholinergic neurons.

NOTE Confidence: 0.934005198666667

01:15:37.470 --> 01:15:39.806 It's a mix, and it's an area that

NOTE Confidence: 0.934005198666667

 $01{:}15{:}39.806 \dashrightarrow 01{:}15{:}41.793$ doesn't have easily discernible

NOTE Confidence: 0.934005198666667

01:15:41.793 --> 01:15:43.127 boundaries necessarily.

NOTE Confidence: 0.934005198666667

01:15:43.130 --> 01:15:44.768 So when people work with it

NOTE Confidence: 0.934005198666667

01:15:44.768 --> 01:15:46.800 in in the rat and the road,

NOTE Confidence: 0.934005198666667

 $01:15:46.800 \longrightarrow 01:15:48.230$ it's a little bit difficult.

NOTE Confidence: 0.934005198666667

 $01:15:48.230 \longrightarrow 01:15:51.272$ So one way that David went about that that

NOTE Confidence: 0.934005198666667

 $01:15:51.272 \longrightarrow 01:15:54.279$ gets to your second point is he tried to.

NOTE Confidence: 0.934005198666667

 $01:15:54.280 \longrightarrow 01:15:55.330$ Positional has electrodes

 $01:15:55.330 \longrightarrow 01:15:57.080$ in sort of a central,

NOTE Confidence: 0.934005198666667

 $01:15:57.080 \longrightarrow 01:15:58.181$ not very anterior,

NOTE Confidence: 0.934005198666667

01:15:58.181 --> 01:16:00.383 not very posterior and operating medial,

NOTE Confidence: 0.934005198666667

 $01:16:00.390 \longrightarrow 01:16:01.222$ not very lateral area.

NOTE Confidence: 0.934005198666667

 $01:16:01.222 \longrightarrow 01:16:02.762$ So he could just be in the

NOTE Confidence: 0.934005198666667

01:16:02.762 --> 01:16:03.957 middle of the canonical BP,

NOTE Confidence: 0.934005198666667

 $01:16:03.960 \longrightarrow 01:16:05.934$ at least as described on Atlas is.

NOTE Confidence: 0.934005198666667

 $01:16:05.940 \longrightarrow 01:16:07.228$ So that's not necessarily

NOTE Confidence: 0.934005198666667

01:16:07.228 --> 01:16:08.516 what you might want,

NOTE Confidence: 0.934005198666667

 $01:16:08.520 \longrightarrow 01:16:12.112$ but he did not see variation based on

NOTE Confidence: 0.934005198666667

 $01:16:12.112 \longrightarrow 01:16:14.439$ electrode placement with his signals.

NOTE Confidence: 0.934005198666667

01:16:14.440 --> 01:16:16.648 Had he gone very far anterior

NOTE Confidence: 0.934005198666667

 $01:16:16.648 \longrightarrow 01:16:18.540$ posterior to search for that,

NOTE Confidence: 0.934005198666667

 $01:16:18.540 \longrightarrow 01:16:21.242$ he might have seen that because there

NOTE Confidence: 0.934005198666667

 $01:16:21.242 \longrightarrow 01:16:23.396$ is work emerging from other labs,

NOTE Confidence: 0.934005198666667

 $01:16:23.396 \longrightarrow 01:16:24.724$ including the Berridge lab.

01:16:24.730 --> 01:16:27.142 Showing that there the VP plays

NOTE Confidence: 0.934005198666667

 $01:16:27.142 \longrightarrow 01:16:29.661$ different roles in behavior when you

NOTE Confidence: 0.934005198666667

 $01:16:29.661 \longrightarrow 01:16:32.169$ are more anterior versus more posterior.

NOTE Confidence: 0.934005198666667

 $01:16:32.170 \longrightarrow 01:16:33.815$ So that's an important issue

NOTE Confidence: 0.934005198666667

 $01:16:33.815 \longrightarrow 01:16:35.901$ that that we've not looked at as

NOTE Confidence: 0.934005198666667

 $01:16:35.901 \longrightarrow 01:16:37.728$ far as cell types in the rat.

NOTE Confidence: 0.934005198666667

01:16:37.730 --> 01:16:40.145 So not having the the beauty of

NOTE Confidence: 0.934005198666667

 $01{:}16{:}40.145 \dashrightarrow 01{:}16{:}42.697$ the transgenic mouse to let let us

NOTE Confidence: 0.934005198666667

 $01:16:42.697 \longrightarrow 01:16:44.502$ access different cell types easily,

NOTE Confidence: 0.934005198666667

01:16:44.510 --> 01:16:45.776 like Gabaergic cells,

NOTE Confidence: 0.934005198666667

01:16:45.776 --> 01:16:47.111 glutamatergic cells, etc.

NOTE Confidence: 0.934005198666667

 $01:16:47.111 \longrightarrow 01:16:49.566$ We can't say for sure.

NOTE Confidence: 0.934005198666667

 $01{:}16{:}49.570 \dashrightarrow 01{:}16{:}52.438$ The area that he implanted his

NOTE Confidence: 0.934005198666667

 $01:16:52.438 \longrightarrow 01:16:55.300$ electrodes and his mostly Gabaergic.

NOTE Confidence: 0.934005198666667

 $01:16:55.300 \longrightarrow 01:16:57.694$ When we you there is a small

01:16:57.694 --> 01:16:58.720 a glutamatergic population,

NOTE Confidence: 0.934005198666667 01:16:58.720 --> 01:16:59.025 though, NOTE Confidence: 0.934005198666667

 $01:16:59.025 \longrightarrow 01:17:00.550$ for example that people studied

NOTE Confidence: 0.934005198666667

 $01:17:00.550 \longrightarrow 01:17:02.649$ that has a critical behavioral role.

NOTE Confidence: 0.934005198666667

 $01:17:02.650 \longrightarrow 01:17:05.569$ When we use waveform shape and other

NOTE Confidence: 0.934005198666667

 $01:17:05.569 \longrightarrow 01:17:07.224$ neuro physiological characteristics to

NOTE Confidence: 0.934005198666667

01:17:07.224 --> 01:17:09.604 try to cluster neurons into different types,

NOTE Confidence: 0.934005198666667

 $01:17:09.610 \longrightarrow 01:17:10.600$ we get mostly,

NOTE Confidence: 0.934005198666667 01:17:10.600 --> 01:17:11.260 you know, NOTE Confidence: 0.934005198666667

 $01:17:11.260 \longrightarrow 01:17:13.780$ big amorphous cluster that we presume

NOTE Confidence: 0.934005198666667

 $01:17:13.780 \longrightarrow 01:17:15.918$ is largely these canonical Gabaergic

NOTE Confidence: 0.934005198666667

 $01:17:15.918 \longrightarrow 01:17:18.627$ neurons and a smaller cluster that we

NOTE Confidence: 0.934005198666667

 $01:17:18.627 \longrightarrow 01:17:21.378$ guess could be the glutamatergic neurons,

NOTE Confidence: 0.934005198666667

01:17:21.380 --> 01:17:24.020 because it tends to signal very

NOTE Confidence: 0.934005198666667

 $01:17:24.020 \longrightarrow 01:17:25.780$ differently from the Gabaergic.

NOTE Confidence: 0.934005198666667 01:17:25.780 --> 01:17:26.091 Iran,

 $01:17:26.091 \longrightarrow 01:17:28.579$ so it doesn't show any of the same

NOTE Confidence: 0.934005198666667

01:17:28.579 --> 01:17:30.144 kinds of signals that I've just

NOTE Confidence: 0.934005198666667

 $01:17:30.144 \longrightarrow 01:17:31.952$ talked to you about, but we can't.

NOTE Confidence: 0.934005198666667

 $01:17:31.952 \longrightarrow 01:17:33.222$ And unless we do something

NOTE Confidence: 0.934005198666667

01:17:33.222 --> 01:17:34.510 you know more rigorous,

NOTE Confidence: 0.934005198666667

 $01:17:34.510 \longrightarrow 01:17:35.754$ something genetic like using

NOTE Confidence: 0.934005198666667

 $01:17:35.754 \longrightarrow 01:17:37.620$ viruses somehow in the rat to

NOTE Confidence: 0.934005198666667

01:17:37.682 --> 01:17:38.886 access the different populations

NOTE Confidence: 0.934005198666667

 $01:17:38.886 \longrightarrow 01:17:41.070$ or redo the work in the house.

NOTE Confidence: 0.934005198666667

 $01:17:41.070 \longrightarrow 01:17:42.480$ We really can't say for sure,

NOTE Confidence: 0.934005198666667

 $01:17:42.480 \longrightarrow 01:17:44.839$ and we've not even begun to think

NOTE Confidence: 0.934005198666667

 $01:17:44.839 \longrightarrow 01:17:47.530$ about or touch on the cholinergic.

NOTE Confidence: 0.934005198666667

 $01:17:47.530 \longrightarrow 01:17:48.499$ Aspects of this?

NOTE Confidence: 0.934005198666667

01:17:48.499 --> 01:17:51.290 I'm ashamed to admit in front of Marina,

NOTE Confidence: 0.729416466666667

01:17:51.640 --> 01:17:56.090 oh, that's OK. Are there any other questions?

01:17:56.090 --> 01:17:58.178 I don't see anything else in the track chat.

NOTE Confidence: 0.75193547125

01:18:00.820 --> 01:18:04.044 If not one more please or I'm Joe.

NOTE Confidence: 0.592878958

 $01:18:06.980 \longrightarrow 01:18:09.860$ So I actually have to.

NOTE Confidence: 0.592878958

 $01:18:09.860 \longrightarrow 01:18:11.685$ I guess it's more related

NOTE Confidence: 0.592878958

 $01:18:11.685 \longrightarrow 01:18:12.780$ to technical questions.

NOTE Confidence: 0.592878958

 $01:18:12.780 \longrightarrow 01:18:16.590$ One was that when you used

NOTE Confidence: 0.592878958

 $01:18:16.590 \longrightarrow 01:18:19.429$ optogenetics that you injected.

NOTE Confidence: 0.592878958

01:18:19.430 --> 01:18:21.530 Inhibitory virus bilaterally.

NOTE Confidence: 0.592878958

 $01{:}18{:}21.530 \dashrightarrow 01{:}18{:}25.660$ But you injected unilateral for the stimulus.

NOTE Confidence: 0.592878958

 $01:18:25.660 \longrightarrow 01:18:27.739$ So I was wondering why there is

NOTE Confidence: 0.592878958

 $01{:}18{:}27.739 \dashrightarrow 01{:}18{:}30.124$ difference used by letter for

NOTE Confidence: 0.592878958

 $01:18:30.124 \longrightarrow 01:18:32.614$ inhibitory and Union letter for

NOTE Confidence: 0.592878958

 $01{:}18{:}32.614 \dashrightarrow 01{:}18{:}35.338$ stimulus and the second one was.

NOTE Confidence: 0.592878958

 $01:18:35.340 \longrightarrow 01:18:38.280$ So all the dopamine signals were measured

NOTE Confidence: 0.8600155625

 $01:18:38.290 \longrightarrow 01:18:40.890$ during the first trial for the

NOTE Confidence: 0.8600155625

 $01:18:40.890 \longrightarrow 01:18:42.685$ second half of the experiments.

 $01:18:42.685 \longrightarrow 01:18:45.006$ And do you think there will be

NOTE Confidence: 0.8600155625

 $01:18:45.006 \longrightarrow 01:18:46.338$ difference if you measure the

NOTE Confidence: 0.825760248

01:18:46.540 --> 01:18:48.750 signals during the free trials?

NOTE Confidence: 0.825760248

 $01:18:48.750 \longrightarrow 01:18:50.019$ Because Brett doesn't.

NOTE Confidence: 0.825760248

 $01:18:50.019 \longrightarrow 01:18:52.134$ No, what they're going to

NOTE Confidence: 0.825760248

 $01:18:52.134 \longrightarrow 01:18:54.169$ get when they press the.

NOTE Confidence: 0.825760248

01:18:54.170 --> 01:18:57.338 Or ignore spoke or pistol ever?

NOTE Confidence: 0.825760248

 $01:18:57.340 \longrightarrow 01:18:59.110$ Yes, those are both good questions

NOTE Confidence: 0.910630422857143

 $01:18:59.140 \longrightarrow 01:19:00.274 \text{ I I think I would do that.}$

NOTE Confidence: 0.910630422857143

 $01:19:00.280 \longrightarrow 01:19:02.170$ The second one first and that is

NOTE Confidence: 0.910630422857143

 $01:19:02.170 \longrightarrow 01:19:04.400$ so in the in the forced trials.

NOTE Confidence: 0.910630422857143

 $01:19:04.400 \longrightarrow 01:19:06.596$ The rat doesn't know what reward it will get,

NOTE Confidence: 0.910630422857143

 $01{:}19{:}06.600 \longrightarrow 01{:}19{:}09.600$ so that's when we see these big signals.

NOTE Confidence: 0.910630422857143

 $01:19:09.600 \longrightarrow 01:19:10.728$ That are very different,

NOTE Confidence: 0.910630422857143

01:19:10.728 --> 01:19:12.138 but in the free trials,

 $01:19:12.140 \longrightarrow 01:19:13.592$ when the animal presses

NOTE Confidence: 0.910630422857143

 $01:19:13.592 \longrightarrow 01:19:15.044$ the lever for sucrose.

NOTE Confidence: 0.910630422857143

 $01:19:15.050 \longrightarrow 01:19:17.346$ He or she knows that that it's

NOTE Confidence: 0.910630422857143

 $01:19:17.346 \longrightarrow 01:19:20.148$ about to drink sucrose in the in the port.

NOTE Confidence: 0.910630422857143

 $01:19:20.150 \longrightarrow 01:19:22.302$ So expectation already reduces

NOTE Confidence: 0.910630422857143

 $01:19:22.302 \longrightarrow 01:19:24.346$ that signal because he knows

NOTE Confidence: 0.910630422857143

01:19:24.346 --> 01:19:25.956 already what's going to happen,

NOTE Confidence: 0.910630422857143

 $01:19:25.960 \longrightarrow 01:19:27.898$ so that that's a great question,

NOTE Confidence: 0.910630422857143

 $01:19:27.900 \longrightarrow 01:19:29.860$ and one reason why we didn't look

NOTE Confidence: 0.910630422857143

 $01:19:29.860 \longrightarrow 01:19:31.648$ at those signals in great detail

NOTE Confidence: 0.910630422857143

 $01:19:31.648 \longrightarrow 01:19:33.133$ through the session is because

NOTE Confidence: 0.910630422857143

 $01:19:33.133 \longrightarrow 01:19:35.386$ by the time you get to the end of

NOTE Confidence: 0.910630422857143

 $01:19:35.386 \longrightarrow 01:19:37.151$ the session is really only enough

NOTE Confidence: 0.910630422857143

 $01:19:37.151 \longrightarrow 01:19:38.936$ data from the sucrose trials.

NOTE Confidence: 0.910630422857143

 $01:19:38.940 \longrightarrow 01:19:40.120$ The beginning of the session.

NOTE Confidence: 0.910630422857143

 $01:19:40.120 \longrightarrow 01:19:41.184$ There's really only enough

01:19:41.184 --> 01:19:42.514 data for the water trials,

NOTE Confidence: 0.910630422857143

 $01:19:42.520 \longrightarrow 01:19:44.005$ but it would be interesting

NOTE Confidence: 0.910630422857143

 $01:19:44.005 \longrightarrow 01:19:45.193$ still to show those.

NOTE Confidence: 0.910630422857143

 $01:19:45.200 \longrightarrow 01:19:46.844$ Think about those more and I

NOTE Confidence: 0.910630422857143

01:19:46.844 --> 01:19:48.520 think that's a nice question,

NOTE Confidence: 0.910630422857143

 $01:19:48.520 \longrightarrow 01:19:50.375$ and it might be interesting to look

NOTE Confidence: 0.910630422857143

 $01:19:50.375 \longrightarrow 01:19:52.153$ at the neural activity around the

NOTE Confidence: 0.910630422857143

 $01:19:52.153 \longrightarrow 01:19:54.309$ lever press for example so that there

NOTE Confidence: 0.910630422857143

 $01:19:54.372 \longrightarrow 01:19:56.160$ are other other time periods that

NOTE Confidence: 0.910630422857143

 $01{:}19{:}56.160 \dashrightarrow 01{:}19{:}57.959$ I think I would be interested in.

NOTE Confidence: 0.910630422857143

 $01{:}19{:}57.960 \dashrightarrow 01{:}19{:}59.958$ Knowing what are those neurons doing

NOTE Confidence: 0.910630422857143

 $01:19:59.958 \longrightarrow 01:20:01.800$ during those other time periods?

NOTE Confidence: 0.910630422857143

 $01{:}20{:}01.800 \dashrightarrow 01{:}20{:}03.672$ The first question you asked is

NOTE Confidence: 0.910630422857143

 $01{:}20{:}03.672 \dashrightarrow 01{:}20{:}04.920$ about the optogenetics procedure

NOTE Confidence: 0.910630422857143

 $01:20:04.969 \longrightarrow 01:20:06.577$ and so this is just something

 $01:20:06.577 \longrightarrow 01:20:08.090$ away that we've typically done it,

NOTE Confidence: 0.910630422857143

 $01:20:08.090 \longrightarrow 01:20:10.344$ and it's probably mostly out of convenience.

NOTE Confidence: 0.910630422857143

01:20:10.350 --> 01:20:12.175 So when you're activating neurons

NOTE Confidence: 0.910630422857143

01:20:12.175 --> 01:20:14.535 you usually can change the animal's

NOTE Confidence: 0.910630422857143

 $01:20:14.535 \longrightarrow 01:20:15.469$ behavior with.

NOTE Confidence: 0.910630422857143

01:20:15.470 --> 01:20:17.258 Optogenetics because it's such

NOTE Confidence: 0.910630422857143

 $01{:}20{:}17.258 \dashrightarrow 01{:}20{:}19.493$ a strong and artificial approach

NOTE Confidence: 0.910630422857143

 $01:20:19.493 \longrightarrow 01:20:21.707$ usually can change the behavior by

NOTE Confidence: 0.910630422857143

 $01{:}20{:}21.707 \dashrightarrow 01{:}20{:}24.039$ impacting the brain just on one side,

NOTE Confidence: 0.910630422857143

 $01:20:24.040 \longrightarrow 01:20:25.882$ and it's easier for the experimenter

NOTE Confidence: 0.910630422857143

 $01:20:25.882 \longrightarrow 01:20:27.677$ because then there's only one connection

NOTE Confidence: 0.910630422857143

 $01:20:27.677 \longrightarrow 01:20:29.483$ that they need to make with the.

NOTE Confidence: 0.910630422857143

01:20:29.490 --> 01:20:31.426 With the Rotary joint and all of this,

NOTE Confidence: 0.910630422857143

 $01:20:31.430 \longrightarrow 01:20:33.440$ but when you're inhibiting in a

NOTE Confidence: 0.910630422857143

 $01:20:33.440 \longrightarrow 01:20:35.476$ particular brain region a lot of

NOTE Confidence: 0.910630422857143

01:20:35.476 --> 01:20:37.378 times behavior is still not greatly

 $01:20:37.378 \longrightarrow 01:20:39.517$ impacted or even almost normal if the

NOTE Confidence: 0.910630422857143

01:20:39.517 --> 01:20:42.346 other side of the brain is still functioning,

NOTE Confidence: 0.910630422857143

 $01:20:42.346 \longrightarrow 01:20:46.070$ so to avoid a possible outcome like that.

NOTE Confidence: 0.910630422857143

01:20:46.070 --> 01:20:46.844 For inhibition,

NOTE Confidence: 0.910630422857143

 $01:20:46.844 \longrightarrow 01:20:49.940$ people often will do it bilaterally and try

NOTE Confidence: 0.910630422857143

 $01:20:50.008 \longrightarrow 01:20:52.825$ to inhibit in both both sides of the brain.

NOTE Confidence: 0.910630422857143 01:20:52.830 --> 01:20:53.943 So it's it's. NOTE Confidence: 0.910630422857143

01:20:53.943 --> 01:20:55.427 It's really just a.

NOTE Confidence: 0.910630422857143

 $01:20:55.430 \longrightarrow 01:20:57.180$ Experimental practicality

NOTE Confidence: 0.942304115

01:20:58.120 --> 01:20:59.940 thank you so much, sure.

NOTE Confidence: 0.8202534

 $01:21:02.960 \longrightarrow 01:21:06.835$ OK. Well, I I want to thank

NOTE Confidence: 0.8202534

 $01:21:06.835 \longrightarrow 01:21:08.630$ you again for a great lecture.

NOTE Confidence: 0.8202534

01:21:08.630 --> 01:21:10.592 Thank you all for being here

NOTE Confidence: 0.8202534

 $01:21:10.592 \longrightarrow 01:21:12.382$ and for your great questions

NOTE Confidence: 0.8202534

 $01:21:12.382 \longrightarrow 01:21:14.830$ and let's thank Dr Janik again.

 $01:21:14.830 \longrightarrow 01:21:16.356$ And for those of you who are

NOTE Confidence: 0.8202534

01:21:16.356 --> 01:21:17.633 having who are trainees who

NOTE Confidence: 0.8202534

01:21:17.633 --> 01:21:19.048 are having lunch with her,

NOTE Confidence: 0.8202534

 $01:21:19.050 \longrightarrow 01:21:22.270$ you should have the link to

NOTE Confidence: 0.8202534

 $01:21:22.270 \longrightarrow 01:21:25.830$ the other zoom room and I hope

NOTE Confidence: 0.8202534

 $01:21:25.830 \longrightarrow 01:21:28.606$ you'll move over to that room to

NOTE Confidence: 0.8202534

01:21:28.606 --> 01:21:30.690 talk with Doctor Genich further.

NOTE Confidence: 0.8202534

 $01:21:30.690 \longrightarrow 01:21:31.798$ Thanks I wanna thank.

NOTE Confidence: 0.9208700175

 $01:21:32.130 \longrightarrow 01:21:33.426$ I want to thank you all so much.

NOTE Confidence: 0.9208700175

 $01{:}21{:}33.430 \dashrightarrow 01{:}21{:}34.702$ This was really a pleasure and

NOTE Confidence: 0.9208700175

 $01{:}21{:}34.702 \dashrightarrow 01{:}21{:}36.000$ thanks for your great questions.