WEBVTT

NOTE duration: "00:58:46.8050000"

NOTE language:en-us

NOTE Confidence: 0.701370894908905

 $00:00:00.000 \longrightarrow 00:00:02.508$ Fortnight yeah we can see out

NOTE Confidence: 0.701370894908905

 $00:00:02.510 \longrightarrow 00:00:04.540$ the window behind you looks

NOTE Confidence: 0.847600843225207

 $00:00:04.540 \longrightarrow 00:00:08.187$ nice. So why don't we get started?

NOTE Confidence: 0.847600843225207

00:00:08.190 --> 00:00:11.246 And I know there's still people joining us,

NOTE Confidence: 0.847600843225207

 $00:00:11.250 \longrightarrow 00:00:13.536$ so to all those in attendance,

NOTE Confidence: 0.847600843225207

 $00:00:13.540 \dashrightarrow 00:00:16.186$ since those logging in welcome to a

NOTE Confidence: 0.847600843225207

00:00:16.186 --> 00:00:18.129 special Cancer Center grand rounds

NOTE Confidence: 0.847600843225207

00:00:18.129 --> 00:00:20.355 where you know very privileged to

NOTE Confidence: 0.847600843225207

00:00:20.355 --> 00:00:23.087 honor the legacy of Doctor John Murnan,

NOTE Confidence: 0.847600843225207

 $00:00:23.090 \longrightarrow 00:00:25.393$ have a very special guest as our

NOTE Confidence: 0.847600843225207

 $00:00:25.393 \longrightarrow 00:00:27.669$ 2000 two 1020 Memorial Lecture,

NOTE Confidence: 0.847600843225207

00:00:27.670 --> 00:00:30.078 and I want to introduce where I

NOTE Confidence: 0.847600843225207

00:00:30.078 --> 00:00:31.966 herbs whose everyone knows is

NOTE Confidence: 0.847600843225207

 $00:00:31.966 \dashrightarrow 00:00:34.312$ our chief of Medical Oncology and

 $00:00:34.312 \longrightarrow 00:00:35.973$ Associate Cancer Center director

NOTE Confidence: 0.847600843225207

 $00{:}00{:}35.973 \dashrightarrow 00{:}00{:}38.409$ for a translation of science and.

NOTE Confidence: 0.847600843225207

 $00:00:38.410 \longrightarrow 00:00:39.858$ Leader of our lungs.

NOTE Confidence: 0.847600843225207

 $00:00:39.858 \longrightarrow 00:00:41.297$ For, among other things,

NOTE Confidence: 0.847600843225207

 $00{:}00{:}41.297 \dashrightarrow 00{:}00{:}43.092$ to speak about the lectureship

NOTE Confidence: 0.847600843225207

00:00:43.092 --> 00:00:44.549 and introduce our speaker.

NOTE Confidence: 0.847600843225207

00:00:44.550 --> 00:00:45.918 So Roy, thank you,

NOTE Confidence: 0.847600843225207

00:00:45.918 --> 00:00:47.286 thanks Charlie and thanks

NOTE Confidence: 0.847600843225207

 $00:00:47.286 \longrightarrow 00:00:48.520$ everyone for joining.

NOTE Confidence: 0.847600843225207

 $00:00:48.520 \longrightarrow 00:00:51.416$ This is a very special lecture because it

NOTE Confidence: 0.847600843225207

 $00:00:51.416 \longrightarrow 00:00:54.448$ honors our colleague and friend John Urine.

NOTE Confidence: 0.847600843225207

 $00:00:54.450 \longrightarrow 00:00:56.074$ If we can go to the next slide,

NOTE Confidence: 0.847600843225207 00:00:56.080 --> 00:00:56.290 please. NOTE Confidence: 0.860820174217224

00:00:59.160 --> 00:01:04.898 Emily. So many people who are on the line,

NOTE Confidence: 0.860820174217224

00:01:04.900 --> 00:01:06.867 new John I see John Sister Mary

00:01:06.867 --> 00:01:09.598 Kay is on and his wife Nancy and

NOTE Confidence: 0.860820174217224

 $00{:}01{:}09.598 \dashrightarrow 00{:}01{:}11.413$ perhaps others are the family.

NOTE Confidence: 0.860820174217224

00:01:11.420 --> 00:01:13.292 One of the special things about

NOTE Confidence: 0.860820174217224

 $00:01:13.292 \longrightarrow 00:01:15.608$ this lecture is we usually can get

NOTE Confidence: 0.860820174217224

00:01:15.608 --> 00:01:17.606 together and talk and in person.

NOTE Confidence: 0.860820174217224

00:01:17.610 --> 00:01:19.885 Hopefully that will happen again next year.

NOTE Confidence: 0.860820174217224

 $00:01:19.890 \longrightarrow 00:01:22.530$ But no, John was a very special person

NOTE Confidence: 0.860820174217224

 $00:01:22.530 \longrightarrow 00:01:25.675$ and as many of us know he passed away in

NOTE Confidence: 0.860820174217224

 $00:01:25.675 \longrightarrow 00:01:28.696$ 2005 of Melanoma at the young age of 47.

NOTE Confidence: 0.860820174217224

 $00:01:28.700 \longrightarrow 00:01:30.315$ He was an associate professor

NOTE Confidence: 0.860820174217224

 $00:01:30.315 \longrightarrow 00:01:32.320$ of medicine at the at Yale.

NOTE Confidence: 0.860820174217224

 $00:01:32.320 \longrightarrow 00:01:34.238$ He was the director of the outpatient

NOTE Confidence: 0.860820174217224

 $00:01:34.238 \longrightarrow 00:01:36.237$ center director of the Lung Cancer Clinic.

NOTE Confidence: 0.860820174217224

00:01:36.240 --> 00:01:38.200 I wouldn't probably be here at Yelp,

NOTE Confidence: 0.860820174217224

 $00:01:38.200 \longrightarrow 00:01:39.600$ not for John and Co.

NOTE Confidence: 0.860820174217224

 $00:01:39.600 \longrightarrow 00:01:41.395$ Leader of the Experimental Therapeutics

 $00:01:41.395 \longrightarrow 00:01:43.630$ program at the Yale Cancer Center.

NOTE Confidence: 0.860820174217224

00:01:43.630 --> 00:01:46.430 He was survived by his wife Nancy,

NOTE Confidence: 0.860820174217224

00:01:46.430 --> 00:01:47.606 his son John,

NOTE Confidence: 0.860820174217224

 $00:01:47.606 \longrightarrow 00:01:50.350$ his mother Jean and I know that

NOTE Confidence: 0.860820174217224

 $00:01:50.440 \longrightarrow 00:01:53.226$ Nancy is here and maybe Gene no.

NOTE Confidence: 0.860820174217224

00:01:53.230 --> 00:01:56.430 Many new new John even better than me.

NOTE Confidence: 0.860820174217224

00:01:56.430 --> 00:01:58.985 I knew him from from the clinical

NOTE Confidence: 0.860820174217224

 $00:01:58.985 \longrightarrow 00:02:00.830$ trials and from meetings.

NOTE Confidence: 0.860820174217224

 $00:02:00.830 \longrightarrow 00:02:02.830$ He was really the consummate

NOTE Confidence: 0.860820174217224

 $00:02:02.830 \longrightarrow 00:02:03.630$ medical oncologist.

NOTE Confidence: 0.860820174217224

00:02:03.630 --> 00:02:06.430 Academically, he was just the most caring,

NOTE Confidence: 0.860820174217224

 $00:02:06.430 \longrightarrow 00:02:08.030$ position and active laboratory

NOTE Confidence: 0.860820174217224

 $00{:}02{:}08.030 \dashrightarrow 00{:}02{:}10.030$ working to develop new treatments.

NOTE Confidence: 0.860820174217224

 $00{:}02{:}10.030 \dashrightarrow 00{:}02{:}12.055$ Very insightful and really the

NOTE Confidence: 0.860820174217224

 $00:02:12.055 \longrightarrow 00:02:14.600$ top program that we have today.

 $00:02:14.600 \longrightarrow 00:02:17.950$ Is in large part due to John in his effort.

NOTE Confidence: 0.860820174217224

 $00:02:17.950 \longrightarrow 00:02:19.966$ He had some efforts to his family

NOTE Confidence: 0.860820174217224

 $00:02:19.966 \longrightarrow 00:02:22.828$ with in Las Vegas and Yell had some

NOTE Confidence: 0.860820174217224

 $00:02:22.828 \longrightarrow 00:02:24.368$ interactions there for awhile,

NOTE Confidence: 0.860820174217224

 $00:02:24.370 \longrightarrow 00:02:26.519$ but really he just was someone that

NOTE Confidence: 0.860820174217224

 $00:02:26.519 \longrightarrow 00:02:29.275$ we all knew and loved and one of

NOTE Confidence: 0.860820174217224

 $00{:}02{:}29.275 \dashrightarrow 00{:}02{:}31.040$ the reasons we've invited today's

NOTE Confidence: 0.860820174217224

00:02:31.108 --> 00:02:33.828 speaker is not only is she an amazing

NOTE Confidence: 0.860820174217224

 $00{:}02{:}33.828 \dashrightarrow 00{:}02{:}35.566$ researcher and clinician lung cancer,

NOTE Confidence: 0.860820174217224

 $00:02:35.566 \longrightarrow 00:02:38.412$ but she knew him as well through the

NOTE Confidence: 0.860820174217224

 $00{:}02{:}38.412 \dashrightarrow 00{:}02{:}40.596$ cogs where John was a major person.

NOTE Confidence: 0.860820174217224

 $00:02:40.600 \longrightarrow 00:02:43.296$ So really very glad to have this lecture.

NOTE Confidence: 0.860820174217224

 $00:02:43.300 \longrightarrow 00:02:45.610$ I gave this lecture 11 years ago

NOTE Confidence: 0.860820174217224

 $00:02:45.610 \longrightarrow 00:02:48.075$ so it's really a pleasure to

NOTE Confidence: 0.860820174217224

 $00:02:48.075 \longrightarrow 00:02:49.479$ introduce today's speaker.

NOTE Confidence: 0.860820174217224

 $00{:}02{:}49.480 \dashrightarrow 00{:}02{:}51.331$ We can go to the next slide, please.

00:02:51.331 --> 00:02:53.736 So in thinking about this,

NOTE Confidence: 0.860820174217224

 $00:02:53.740 \longrightarrow 00:02:56.141$ you know this is the second time

NOTE Confidence: 0.860820174217224

00:02:56.141 --> 00:02:57.873 we've scheduled this with Heather

NOTE Confidence: 0.860820174217224

 $00:02:57.873 \longrightarrow 00:03:00.409$ and we just decided we would do this

NOTE Confidence: 0.860820174217224

 $00:03:00.478 \longrightarrow 00:03:02.935$ virtually because we were so lucky to.

NOTE Confidence: 0.860820174217224

 $00:03:02.940 \longrightarrow 00:03:05.668$ Heather agreed to come and speak with us.

NOTE Confidence: 0.860820174217224

 $00:03:05.670 \longrightarrow 00:03:07.536$ Doctor Heather Wakelee is a professor

NOTE Confidence: 0.860820174217224

 $00{:}03{:}07.536 \dashrightarrow 00{:}03{:}09.893$ of medicine and the interim chief in

NOTE Confidence: 0.860820174217224

00:03:09.893 --> 00:03:11.993 the Division of Oncology at Stanford

NOTE Confidence: 0.860820174217224

 $00{:}03{:}11.993 \dashrightarrow 00{:}03{:}13.799$ University and the President elect

NOTE Confidence: 0.860820174217224

 $00:03:13.799 \longrightarrow 00:03:15.199$ of the International Association

NOTE Confidence: 0.860820174217224

00:03:15.199 --> 00:03:17.408 for the Study of Lung Cancer.

NOTE Confidence: 0.860820174217224

 $00{:}03{:}17.408 \dashrightarrow 00{:}03{:}19.670$ PS When she received her medical

NOTE Confidence: 0.860820174217224

 $00:03:19.747 \longrightarrow 00:03:22.472$ degree from Johns Hopkins University

NOTE Confidence: 0.860820174217224

 $00{:}03{:}22.472 \dashrightarrow 00{:}03{:}24.652$ and completed offer postgraduate

00:03:24.652 --> 00:03:26.623 training at Stanford University

NOTE Confidence: 0.860820174217224

 $00:03:26.623 \longrightarrow 00:03:29.323$ before joining the faculty in 2003.

NOTE Confidence: 0.860820174217224

 $00:03:29.330 \longrightarrow 00:03:30.970$ Heather specializes in the

NOTE Confidence: 0.860820174217224

 $00:03:30.970 \longrightarrow 00:03:32.610$ treatment of lung cancer,

NOTE Confidence: 0.860820174217224

00:03:32.610 --> 00:03:33.864 thymoma and mesothelioma,

NOTE Confidence: 0.860820174217224

 $00:03:33.864 \longrightarrow 00:03:35.954$ and had developed research programs

NOTE Confidence: 0.860820174217224

 $00:03:35.954 \longrightarrow 00:03:38.830$ related to lung cancer in SOMA across

NOTE Confidence: 0.860820174217224

 $00:03:38.830 \longrightarrow 00:03:40.805$ multiple areas including clinical trials,

NOTE Confidence: 0.860820174217224

00:03:40.810 --> 00:03:42.638 translation, working, population Sciences.

NOTE Confidence: 0.860820174217224

00:03:42.638 --> 00:03:44.466 He's the principal investigator.

NOTE Confidence: 0.860820174217224

 $00{:}03{:}44.470 \dashrightarrow 00{:}03{:}45.934$ Numerous clinical trials that

NOTE Confidence: 0.860820174217224

 $00:03:45.934 \longrightarrow 00:03:48.130$ have changed the pattern of care,

NOTE Confidence: 0.860820174217224

 $00:03:48.130 \longrightarrow 00:03:50.517$ and as a faculty director of the

NOTE Confidence: 0.860820174217224

 $00:03:50.517 \longrightarrow 00:03:52.390$ Stanford Cancer Clinical Trials Office

NOTE Confidence: 0.860820174217224

 $00:03:52.390 \longrightarrow 00:03:54.355$ and lead investigator for Ikago,

NOTE Confidence: 0.860820174217224

 $00:03:54.360 \longrightarrow 00:03:57.168$ Akron Clinical Trials Group at Stanford.

 $00:03:57.170 \longrightarrow 00:03:59.305$ He also is active in many organizations,

NOTE Confidence: 0.860820174217224

 $00{:}03{:}59.310 \dashrightarrow 00{:}04{:}00.835$ but actually I've shared many

NOTE Confidence: 0.860820174217224

 $00:04:00.835 \longrightarrow 00:04:01.750$ patients with her.

NOTE Confidence: 0.860820174217224

 $00:04:01.750 \longrightarrow 00:04:03.405$ She's also the most compassionate

NOTE Confidence: 0.860820174217224

 $00:04:03.405 \longrightarrow 00:04:05.060$ physician and the person that

NOTE Confidence: 0.842521250247955

 $00:04:05.119 \longrightarrow 00:04:06.464$ I would call whenever I

NOTE Confidence: 0.842521250247955

 $00:04:06.464 \longrightarrow 00:04:07.540$ have a difficult case,

NOTE Confidence: 0.842521250247955

 $00:04:07.540 \longrightarrow 00:04:09.370$ especially in the go far field.

NOTE Confidence: 0.842521250247955

00:04:09.370 --> 00:04:11.505 In fact, I did that last night,

NOTE Confidence: 0.842521250247955

 $00{:}04{:}11.510 \dashrightarrow 00{:}04{:}13.931$ so if you go to the next slide we

NOTE Confidence: 0.842521250247955

 $00{:}04{:}13.931 \dashrightarrow 00{:}04{:}15.936$ normally at this point I would

NOTE Confidence: 0.842521250247955

00:04:15.936 --> 00:04:17.601 be handing Heather her plaque,

NOTE Confidence: 0.842521250247955

00:04:17.610 --> 00:04:19.440 but due to the amazing efficiency

NOTE Confidence: 0.842521250247955

00:04:19.440 --> 00:04:20.660 of Yale Cancer Center,

NOTE Confidence: 0.842521250247955

 $00:04:20.660 \longrightarrow 00:04:23.128$ Heather hasn't show us.

 $00:04:23.130 \longrightarrow 00:04:25.050$ And so it already arrived.

NOTE Confidence: 0.830419361591339

00:04:25.560 --> 00:04:27.989 Can you see it? So here we

NOTE Confidence: 0.830419361591339

 $00:04:27.990 \longrightarrow 00:04:30.230$ go and then last night we actually

NOTE Confidence: 0.830419361591339

 $00:04:30.230 \longrightarrow 00:04:32.149$ had a little get together.

NOTE Confidence: 0.830419361591339

 $00:04:32.150 \longrightarrow 00:04:34.320$ Some of us on zoom and you

NOTE Confidence: 0.830419361591339

 $00:04:34.320 \longrightarrow 00:04:36.319$ can see Pam Kuntz there.

NOTE Confidence: 0.830419361591339

 $00:04:36.320 \longrightarrow 00:04:38.518$ Who we recruited out of Heathers Group

NOTE Confidence: 0.830419361591339

 $00:04:38.518 \longrightarrow 00:04:41.194$ and we did a little zoom happy hour

NOTE Confidence: 0.830419361591339

 $00:04:41.194 \longrightarrow 00:04:43.324$ and without drinks but just talked

NOTE Confidence: 0.830419361591339

 $00:04:43.324 \longrightarrow 00:04:45.688$ about John talked about lung cancer.

NOTE Confidence: 0.830419361591339

 $00{:}04{:}45.690 \dashrightarrow 00{:}04{:}47.420$ So Heather it's wonderful to

NOTE Confidence: 0.830419361591339

 $00:04:47.420 \longrightarrow 00:04:48.804$ have you here today.

NOTE Confidence: 0.830419361591339

 $00:04:48.810 \longrightarrow 00:04:50.230$ Thank you to them.

NOTE Confidence: 0.830419361591339

00:04:50.230 --> 00:04:52.360 You're in family for your support

NOTE Confidence: 0.830419361591339

 $00:04:52.434 \longrightarrow 00:04:54.534$ and again and we Miss John so

NOTE Confidence: 0.830419361591339

 $00{:}04{:}54.534 \dashrightarrow 00{:}04{:}56.810$ much but we're looking forward to.

 $00:04:56.810 \longrightarrow 00:04:58.635$ His memory to a wonderful

NOTE Confidence: 0.830419361591339

00:04:58.635 --> 00:05:00.095 lecture from Heather Wakelee,

NOTE Confidence: 0.830419361591339

 $00:05:00.100 \longrightarrow 00:05:02.296$ 15 years of advances in EGFR

NOTE Confidence: 0.830419361591339

00:05:02.296 --> 00:05:03.765 mutant lung cancer, Heather,

NOTE Confidence: 0.830419361591339

 $00:05:03.765 \longrightarrow 00:05:04.860$ welcome to Yell.

NOTE Confidence: 0.858495354652405

00:05:06.130 --> 00:05:07.810 Thank you, thanks right for

NOTE Confidence: 0.858495354652405

 $00:05:07.810 \longrightarrow 00:05:09.490$ that very kind introduction for

NOTE Confidence: 0.858495354652405

 $00:05:09.548 \longrightarrow 00:05:11.198$ inviting me to be here today.

NOTE Confidence: 0.858495354652405

 $00:05:11.200 \longrightarrow 00:05:13.293$ I really wish I could be there

NOTE Confidence: 0.858495354652405

 $00:05:13.293 \longrightarrow 00:05:15.573$ in person to get to visit with

NOTE Confidence: 0.858495354652405

 $00:05:15.573 \longrightarrow 00:05:17.535$ all of you toward the campus,

NOTE Confidence: 0.858495354652405

 $00:05:17.540 \longrightarrow 00:05:20.076$ which I haven't seen in a few years.

NOTE Confidence: 0.858495354652405

 $00{:}05{:}20.080 \dashrightarrow 00{:}05{:}22.753$ I know there's been a lot of lot of

NOTE Confidence: 0.858495354652405

 $00{:}05{:}22.753 \dashrightarrow 00{:}05{:}24.701$ development an I really would have

NOTE Confidence: 0.858495354652405

 $00:05:24.701 \longrightarrow 00:05:27.049$ valued getting to meet with the family.

00:05:27.050 --> 00:05:29.269 As Roy mentioned, I did know John,

NOTE Confidence: 0.858495354652405

 $00{:}05{:}29.270 \dashrightarrow 00{:}05{:}31.489$ I was pretty early in my career,

NOTE Confidence: 0.858495354652405

00:05:31.490 --> 00:05:33.485 but he was very welcoming when I

NOTE Confidence: 0.858495354652405

 $00:05:33.485 \longrightarrow 00:05:35.928$ showed up at the Kage Akron meanings.

NOTE Confidence: 0.858495354652405

00:05:35.930 --> 00:05:37.242 I remember having several

NOTE Confidence: 0.858495354652405

 $00:05:37.242 \longrightarrow 00:05:38.226$ conversations with him.

NOTE Confidence: 0.858495354652405

 $00{:}05{:}38.230 \dashrightarrow 00{:}05{:}40.378$ And I very vividly remember when

NOTE Confidence: 0.858495354652405

00:05:40.378 --> 00:05:43.089 I heard of his death and how

NOTE Confidence: 0.858495354652405

 $00{:}05{:}43.089 \to 00{:}05{:}45.089$ devastating that was for me.

NOTE Confidence: 0.858495354652405

 $00:05:45.090 \longrightarrow 00:05:46.990$ For whole lung cancer community.

NOTE Confidence: 0.858495354652405

 $00:05:46.990 \longrightarrow 00:05:49.489$ And so when Roy invited me to

NOTE Confidence: 0.858495354652405

 $00:05:49.489 \longrightarrow 00:05:52.239$ give this talk and I was thinking

NOTE Confidence: 0.858495354652405

 $00:05:52.239 \longrightarrow 00:05:54.224$ about what could I cover,

NOTE Confidence: 0.858495354652405

 $00:05:54.230 \longrightarrow 00:05:56.694$ thought about the fact that we lost

NOTE Confidence: 0.858495354652405

00:05:56.694 --> 00:05:59.208 John 15 years ago and thought about

NOTE Confidence: 0.858495354652405

 $00{:}05{:}59.208 \dashrightarrow 00{:}06{:}01.806$ how much the world of lung cancer

 $00:06:01.806 \longrightarrow 00:06:04.518$ therapy has changed during that time.

NOTE Confidence: 0.858495354652405

 $00:06:04.520 \longrightarrow 00:06:06.872$ Ann wanted to use the story

NOTE Confidence: 0.858495354652405

 $00{:}06{:}06.872 \dashrightarrow 00{:}06{:}08.760$ with EGFR lung cancer is.

NOTE Confidence: 0.858495354652405

 $00:06:08.760 \longrightarrow 00:06:10.808$ As a real example of one of the

NOTE Confidence: 0.858495354652405

 $00:06:10.808 \longrightarrow 00:06:12.471$ most dramatic changes that have

NOTE Confidence: 0.858495354652405

00:06:12.471 --> 00:06:14.286 happened during that time period.

NOTE Confidence: 0.858495354652405

 $00:06:14.290 \longrightarrow 00:06:16.537$ But I also wanted to weave in

NOTE Confidence: 0.858495354652405

 $00{:}06{:}16.537 \dashrightarrow 00{:}06{:}18.967$ a different story and that was

NOTE Confidence: 0.858495354652405

00:06:18.967 --> 00:06:20.365 about patient communication,

NOTE Confidence: 0.858495354652405

00:06:20.370 --> 00:06:22.986 and so I'm also going to be talking

NOTE Confidence: 0.858495354652405

00:06:22.986 --> 00:06:25.714 about that and framing it a little

NOTE Confidence: 0.858495354652405

 $00:06:25.714 \longrightarrow 00:06:27.709$ bit around another physician who

NOTE Confidence: 0.858495354652405

 $00{:}06{:}27.780 \to 00{:}06{:}30.174$ unfortunately lost to cancer and who

NOTE Confidence: 0.858495354652405

 $00:06:30.174 \longrightarrow 00:06:32.868$ has a connection to yell as well.

NOTE Confidence: 0.858495354652405

 $00:06:32.868 \longrightarrow 00:06:35.064$ And that's Paul Community and his

 $00:06:35.064 \longrightarrow 00:06:37.467$ breath when breath becomes air book,

NOTE Confidence: 0.858495354652405

00:06:37.470 --> 00:06:39.750 I was I was his physician,

NOTE Confidence: 0.858495354652405

 $00:06:39.750 \longrightarrow 00:06:42.756$ and in that book he talked a lot about

NOTE Confidence: 0.858495354652405

 $00:06:42.756 \longrightarrow 00:06:45.477$ his philosophy about living as a.

NOTE Confidence: 0.858495354652405

00:06:45.480 --> 00:06:46.744 Well, first of all,

NOTE Confidence: 0.858495354652405

00:06:46.744 --> 00:06:48.640 is being a care provider physician

NOTE Confidence: 0.858495354652405

 $00:06:48.708 \longrightarrow 00:06:50.613$ for patients facing cancer and

NOTE Confidence: 0.858495354652405

00:06:50.613 --> 00:06:51.756 other serious issues.

NOTE Confidence: 0.858495354652405

 $00:06:51.760 \longrightarrow 00:06:54.464$ But then as a patient and how he

NOTE Confidence: 0.858495354652405

 $00:06:54.464 \longrightarrow 00:06:56.607$ thought about living as a patient

NOTE Confidence: 0.858495354652405

 $00{:}06{:}56.607 \dashrightarrow 00{:}06{:}59.106$ an he really captured a lot of

NOTE Confidence: 0.858495354652405

 $00:06:59.106 \longrightarrow 00:07:01.188$ the ways I think about things.

NOTE Confidence: 0.858495354652405

 $00:07:01.190 \longrightarrow 00:07:03.535$ And as we talked together and so

NOTE Confidence: 0.858495354652405

 $00:07:03.535 \longrightarrow 00:07:06.069$ I wanted to share that as well.

NOTE Confidence: 0.858495354652405

 $00:07:06.070 \longrightarrow 00:07:09.910$ So I'm going to try to do over the next hour.

NOTE Confidence: 0.858495354652405

 $00:07:09.910 \longrightarrow 00:07:13.222$ I'm going to go ahead and share my

 $00:07:13.222 \longrightarrow 00:07:18.230$ screen here. And give you the talk.

NOTE Confidence: 0.858495354652405 00:07:18.230 --> 00:07:19.930 OK, and. NOTE Confidence: 0.858495354652405

00:07:19.930 --> 00:07:22.480 View slideshow OK,

NOTE Confidence: 0.858495354652405

 $00:07:22.480 \longrightarrow 00:07:24.300$ so hopefully you're seeing this

NOTE Confidence: 0.858495354652405

 $00{:}07{:}24.300 \dashrightarrow 00{:}07{:}27.203$ now show I titled this 15 years of

NOTE Confidence: 0.858495354652405

 $00:07:27.203 \longrightarrow 00:07:29.225$ advances in EGFR mutant non small

NOTE Confidence: 0.858495354652405

 $00:07:29.225 \longrightarrow 00:07:31.657$ cell lung cancer lung cancer and then

NOTE Confidence: 0.858495354652405

00:07:31.657 --> 00:07:34.132 with the focus on patient communication,

NOTE Confidence: 0.858495354652405

 $00:07:34.132 \longrightarrow 00:07:36.376$ these are my disclosures.

NOTE Confidence: 0.858495354652405

 $00:07:36.380 \longrightarrow 00:07:38.366$ And then as I thought about

NOTE Confidence: 0.858495354652405

 $00:07:38.366 \longrightarrow 00:07:39.359$ a learning objectives,

NOTE Confidence: 0.858495354652405

 $00:07:39.360 \longrightarrow 00:07:41.696$ it was to really talk about how that

NOTE Confidence: 0.858495354652405

 $00{:}07{:}41.696 \dashrightarrow 00{:}07{:}43.659$ EGFR Field has evolved overtime,

NOTE Confidence: 0.858495354652405

 $00{:}07{:}43.660 \dashrightarrow 00{:}07{:}44.984$ specifically the EGFR Tkis,

NOTE Confidence: 0.858495354652405

 $00:07:44.984 \longrightarrow 00:07:46.970$ and then a couple of things.

 $00:07:46.970 \longrightarrow 00:07:49.301$ We're looking at adding to Egypt art

NOTE Confidence: 0.858495354652405

 $00:07:49.301 \longrightarrow 00:07:51.939$ therapy and then talk about communication.

NOTE Confidence: 0.858495354652405

 $00:07:51.940 \longrightarrow 00:07:54.532$ So when we think about advances

NOTE Confidence: 0.858495354652405

 $00:07:54.532 \longrightarrow 00:07:56.261$ in the treatment, EGFR,

NOTE Confidence: 0.858495354652405

00:07:56.261 --> 00:07:58.847 TKI eyes alone, veg F combinations,

NOTE Confidence: 0.858495354652405

 $00:07:58.850 \longrightarrow 00:08:00.578$ each diff are combinations,

NOTE Confidence: 0.858495354652405

 $00:08:00.578 \longrightarrow 00:08:02.050$ chemotherapy, immune therapy,

NOTE Confidence: 0.858495354652405

 $00:08:02.050 \longrightarrow 00:08:04.130$ and then the communication.

NOTE Confidence: 0.858495354652405

 $00{:}08{:}04.130 --> 00{:}08{:}06.570$ So back in 2000 and I was just

NOTE Confidence: 0.858495354652405

 $00:08:06.570 \longrightarrow 00:08:08.778$ starting my time as a oncologist.

NOTE Confidence: 0.858495354652405

 $00:08:08.780 \longrightarrow 00:08:11.610$ This was the state of the art and what this

NOTE Confidence: 0.886035919189453

00:08:11.682 --> 00:08:14.265 is showing is survival curve from a

NOTE Confidence: 0.886035919189453

 $00:08:14.265 \longrightarrow 00:08:16.100$ trial where patients were treated

NOTE Confidence: 0.886035919189453

 $00:08:16.100 \longrightarrow 00:08:18.732$ with only things we knew to use which

NOTE Confidence: 0.886035919189453

 $00:08:18.740 \longrightarrow 00:08:20.116$ was just chemotherapy combinations

NOTE Confidence: 0.886035919189453

00:08:20.116 --> 00:08:22.548 and you can see that we hadn't

 $00:08:22.548 \longrightarrow 00:08:24.378$ made a whole lot of progress.

NOTE Confidence: 0.886035919189453

 $00:08:24.380 \longrightarrow 00:08:25.708$ We were helping people,

NOTE Confidence: 0.886035919189453

00:08:25.708 --> 00:08:27.700 but not for nearly long enough,

NOTE Confidence: 0.886035919189453

00:08:27.700 --> 00:08:29.608 and without really being able to

NOTE Confidence: 0.886035919189453

00:08:29.608 --> 00:08:31.286 distinguish what was the best

NOTE Confidence: 0.886035919189453

00:08:31.286 --> 00:08:33.006 treatment for any given patient.

NOTE Confidence: 0.886035919189453

 $00:08:33.010 \longrightarrow 00:08:35.476$ Because all of these different chemotherapy

NOTE Confidence: 0.886035919189453

 $00{:}08{:}35.476 \dashrightarrow 00{:}08{:}37.460$ combinations looked about the same.

NOTE Confidence: 0.886035919189453

 $00{:}08{:}37.460 \dashrightarrow 00{:}08{:}39.428$ But many people realize that there

NOTE Confidence: 0.886035919189453

 $00:08:39.428 \longrightarrow 00:08:41.536$ were things about lung cancer that

NOTE Confidence: 0.886035919189453

 $00:08:41.536 \longrightarrow 00:08:43.738$ were different than some normal tissue,

NOTE Confidence: 0.886035919189453

 $00:08:43.740 \longrightarrow 00:08:45.988$ and one of them was a high expression

NOTE Confidence: 0.886035919189453

 $00{:}08{:}45.988 \dashrightarrow 00{:}08{:}48.364$ of this EGFR or epidermal growth

NOTE Confidence: 0.886035919189453

00:08:48.364 --> 00:08:49.675 factor receptor protein,

NOTE Confidence: 0.886035919189453

 $00:08:49.680 \longrightarrow 00:08:51.666$ and what this cartoon is showing

 $00:08:51.666 \longrightarrow 00:08:53.787$ is that the epidermal growth factor

NOTE Confidence: 0.886035919189453

 $00{:}08{:}53.787 \dashrightarrow 00{:}08{:}56.304$ receptor is a protein. It's a receptor.

NOTE Confidence: 0.886035919189453

 $00:08:56.304 \longrightarrow 00:08:58.740$ It's on the cell surface and it's

NOTE Confidence: 0.886035919189453

 $00:08:58.819 \longrightarrow 00:09:01.888$ sitting out there with the GF binding to it.

NOTE Confidence: 0.886035919189453

 $00:09:01.890 \longrightarrow 00:09:04.682$ And then it sets off a whole bunch

NOTE Confidence: 0.886035919189453 00:09:04.682 --> 00:09:05.380 of signaling, NOTE Confidence: 0.886035919189453

 $00:09:05.380 \longrightarrow 00:09:08.180$ which does a lot within the cell regarding.

NOTE Confidence: 0.886035919189453

 $00{:}09{:}08.180 \dashrightarrow 00{:}09{:}10.016$ Growth of cells. Survival of cell.

NOTE Confidence: 0.886035919189453

 $00:09:10.020 \longrightarrow 00:09:12.132$ So it was thought that each day Fr

NOTE Confidence: 0.886035919189453

00:09:12.132 --> 00:09:14.607 was likely pretty important in lung cancer,

NOTE Confidence: 0.886035919189453

 $00{:}09{:}14.610 \dashrightarrow 00{:}09{:}16.902$ and you'll notice one of these

NOTE Confidence: 0.886035919189453

00:09:16.902 --> 00:09:18.800 reviews was written by Roy.

NOTE Confidence: 0.886035919189453

 $00:09:18.800 \longrightarrow 00:09:20.875$ And so people develop different

NOTE Confidence: 0.886035919189453

 $00:09:20.875 \longrightarrow 00:09:22.535$ ways to target EGFR,

NOTE Confidence: 0.886035919189453

 $00:09:22.540 \longrightarrow 00:09:24.620$ the most common being these

NOTE Confidence: 0.886035919189453

 $00{:}09{:}24.620 \dashrightarrow 00{:}09{:}25.868$ tyrosine kinase inhibitors.

 $00:09:25.870 \longrightarrow 00:09:27.950$ So this is the tyrosine

NOTE Confidence: 0.886035919189453

 $00:09:27.950 \longrightarrow 00:09:29.198$ kinase receptor ligands,

NOTE Confidence: 0.886035919189453

00:09:29.200 --> 00:09:31.755 binds and then you get signaling down

NOTE Confidence: 0.886035919189453

 $00:09:31.755 \longrightarrow 00:09:34.609$ here and the tyrosine kinase domain.

NOTE Confidence: 0.886035919189453

 $00:09:34.610 \longrightarrow 00:09:37.184$ So these medications that are pill

NOTE Confidence: 0.886035919189453

 $00:09:37.184 \longrightarrow 00:09:39.960$ drugs oral can block that tyrosine

NOTE Confidence: 0.886035919189453

 $00:09:39.960 \longrightarrow 00:09:42.906$ kinase activity than the first of

NOTE Confidence: 0.886035919189453

 $00{:}09{:}42.906 \dashrightarrow 00{:}09{:}45.138$ these developed was to fit Nip.

NOTE Confidence: 0.886035919189453

 $00:09:45.140 \longrightarrow 00:09:47.023$ And I'm going to talk about each

NOTE Confidence: 0.886035919189453

 $00:09:47.023 \longrightarrow 00:09:48.638$ of these drugs a little bit,

NOTE Confidence: 0.886035919189453

 $00:09:48.640 \longrightarrow 00:09:49.980$ but there are other strategies

NOTE Confidence: 0.886035919189453

 $00:09:49.980 \longrightarrow 00:09:51.052$ to hit tyrosine kinases.

NOTE Confidence: 0.886035919189453

 $00{:}09{:}51.060 \dashrightarrow 00{:}09{:}53.132$ One is to block where the leg and

NOTE Confidence: 0.886035919189453

00:09:53.132 --> 00:09:54.764 would bind and will talk about

NOTE Confidence: 0.886035919189453

 $00:09:54.764 \longrightarrow 00:09:56.710$ a couple of drugs that do that,

 $00:09:56.710 \longrightarrow 00:09:58.702$ and then another is to just take all

NOTE Confidence: 0.886035919189453

 $00{:}09{:}58.702 \dashrightarrow 00{:}10{:}00.648$ the Ligue and what would be binding

NOTE Confidence: 0.886035919189453

 $00:10:00.648 \longrightarrow 00:10:02.629$ and turning things on and remove it.

NOTE Confidence: 0.886035919189453

 $00:10:02.630 \longrightarrow 00:10:03.970$ And that's a different approach

NOTE Confidence: 0.886035919189453

 $00:10:03.970 \longrightarrow 00:10:05.042$ that used as well.

NOTE Confidence: 0.886035919189453

 $00:10:05.050 \longrightarrow 00:10:06.592$ So just wanted to give this

NOTE Confidence: 0.886035919189453

 $00:10:06.592 \longrightarrow 00:10:08.508$ cartoon to set people in mind about

NOTE Confidence: 0.886035919189453

 $00:10:08.508 \longrightarrow 00:10:09.888$ thinking how the drugs work.

NOTE Confidence: 0.886035919189453

 $00:10:09.890 \longrightarrow 00:10:11.698$ So the first drug to fit him and

NOTE Confidence: 0.886035919189453

00:10:11.698 --> 00:10:13.429 you see Doctor Herbst of course

NOTE Confidence: 0.886035919189453

 $00{:}10{:}13.429 \dashrightarrow 00{:}10{:}15.259$ was involved in that was given

NOTE Confidence: 0.886035919189453

 $00:10:15.320 \longrightarrow 00:10:16.960$ to patients with lung cancer.

NOTE Confidence: 0.886035919189453

00:10:16.960 --> 00:10:19.137 And it was noted that some people,

NOTE Confidence: 0.886035919189453

00:10:19.140 --> 00:10:21.000 maybe 10\%, had these dramatic responses,

NOTE Confidence: 0.886035919189453

 $00:10:21.000 \longrightarrow 00:10:22.196$ which was really exciting.

NOTE Confidence: 0.886035919189453

 $00:10:22.196 \longrightarrow 00:10:24.740$ But we didn't know how to select them.

 $00:10:24.740 \longrightarrow 00:10:27.164$ And this led to a larger trial with

NOTE Confidence: 0.886035919189453

 $00:10:27.164 \longrightarrow 00:10:29.397$ a similar drug called or lot nib,

NOTE Confidence: 0.886035919189453

 $00:10:29.400 \longrightarrow 00:10:31.619$ where patients receive this drug in the

NOTE Confidence: 0.886035919189453

 $00:10:31.619 \longrightarrow 00:10:34.069$ second or third line or they got placebo.

NOTE Confidence: 0.886035919189453

 $00:10:34.070 \longrightarrow 00:10:35.930$ And when you looked at it,

NOTE Confidence: 0.886035919189453

00:10:35.930 --> 00:10:37.490 response rates were about 10%.

NOTE Confidence: 0.886035919189453

00:10:37.490 --> 00:10:38.465 Survival wasn't great,

NOTE Confidence: 0.886035919189453

 $00{:}10{:}38.465 \dashrightarrow 00{:}10{:}41.083$ but it was better than place bo and so

NOTE Confidence: 0.886035919189453

00:10:41.083 --> 00:10:43.078 this was really exciting at the time.

NOTE Confidence: 0.886035919189453

 $00:10:43.080 \dashrightarrow 00:10:45.486$ To see that two month improvement.

NOTE Confidence: 0.886035919189453

 $00:10:45.490 \longrightarrow 00:10:47.075$ But what was really interesting

NOTE Confidence: 0.886035919189453

 $00{:}10{:}47.075 \dashrightarrow 00{:}10{:}49.076$ is people notice that it wasn't

NOTE Confidence: 0.886035919189453

 $00{:}10{:}49.076 \dashrightarrow 00{:}10{:}50.866$ just random who was responding.

NOTE Confidence: 0.886035919189453

 $00:10:50.870 \longrightarrow 00:10:52.628$ It tended to be women attended

NOTE Confidence: 0.886035919189453

 $00:10:52.628 \longrightarrow 00:10:54.269$ the people with the adenocarcinoma

 $00:10:54.269 \longrightarrow 00:10:55.909$ type of lung cancer,

NOTE Confidence: 0.886035919189453

 $00:10:55.910 \longrightarrow 00:10:58.255$ and especially people who had lung cancer,

NOTE Confidence: 0.868898510932922

 $00:10:58.260 \longrightarrow 00:10:59.532$ who had never smoked,

NOTE Confidence: 0.868898510932922

 $00:10:59.532 \longrightarrow 00:11:01.862$ and also notice that a lot of

NOTE Confidence: 0.868898510932922

00:11:01.862 --> 00:11:03.627 them were of Asian ethnicity,

NOTE Confidence: 0.868898510932922

 $00{:}11{:}03.630 \dashrightarrow 00{:}11{:}05.772$ and so this led to at the

NOTE Confidence: 0.868898510932922

 $00:11:05.772 \longrightarrow 00:11:08.000$ study known as the ipass trial,

NOTE Confidence: 0.868898510932922

00:11:08.000 --> 00:11:09.680 which was conducted in Asia.

NOTE Confidence: 0.868898510932922

 $00:11:09.680 \longrightarrow 00:11:11.871$ It was done in people who had

NOTE Confidence: 0.868898510932922

00:11:11.871 --> 00:11:13.380 very limited smoking history,

NOTE Confidence: 0.868898510932922

 $00:11:13.380 \longrightarrow 00:11:14.748$ but who had developed

NOTE Confidence: 0.868898510932922

00:11:14.748 --> 00:11:15.774 adenocarcinoma lung cancer.

NOTE Confidence: 0.868898510932922

00:11:15.780 --> 00:11:18.258 And this ended up being mostly women,

NOTE Confidence: 0.868898510932922

 $00:11:18.260 \longrightarrow 00:11:20.996$ and they were randomized to either get the

NOTE Confidence: 0.868898510932922

00:11:20.996 --> 00:11:23.920 EGFR pill drug you fit nib or chemotherapy,

NOTE Confidence: 0.868898510932922

 $00{:}11{:}23.920 \dashrightarrow 00{:}11{:}26.461$ and I showed you that before the

 $00:11:26.461 \longrightarrow 00:11:28.879$ chemotherapy drugs are all about the same.

NOTE Confidence: 0.868898510932922

 $00:11:28.880 \longrightarrow 00:11:31.240$ When we look at them.

NOTE Confidence: 0.868898510932922

 $00:11:31.240 \longrightarrow 00:11:32.805$ And what was interesting is

NOTE Confidence: 0.868898510932922

 $00:11:32.805 \longrightarrow 00:11:34.370$ when the results came out,

NOTE Confidence: 0.868898510932922

00:11:34.370 --> 00:11:36.248 it was very confusing at first,

NOTE Confidence: 0.868898510932922

 $00:11:36.250 \longrightarrow 00:11:38.128$ because when you look at the

NOTE Confidence: 0.868898510932922

00:11:38.128 --> 00:11:39.067 progression free survival,

NOTE Confidence: 0.868898510932922

 $00:11:39.070 \longrightarrow 00:11:40.816$ which is the time either that

NOTE Confidence: 0.868898510932922

 $00:11:40.816 \longrightarrow 00:11:42.614$ someone is no longer alive or

NOTE Confidence: 0.868898510932922

 $00:11:42.614 \longrightarrow 00:11:44.039$ that their tumor has started

NOTE Confidence: 0.868898510932922

 $00{:}11{:}44.039 \dashrightarrow 00{:}11{:}45.949$ to grow the progression time,

NOTE Confidence: 0.868898510932922

 $00:11:45.950 \longrightarrow 00:11:47.828$ these curves kind of crossed in.

NOTE Confidence: 0.868898510932922

 $00{:}11{:}47.830 \dashrightarrow 00{:}11{:}49.395$ The chemo was looking better

NOTE Confidence: 0.868898510932922

00:11:49.395 --> 00:11:50.647 until about five months,

NOTE Confidence: 0.868898510932922

00:11:50.650 --> 00:11:52.554 six months and then the gym fit

 $00:11:52.554 \longrightarrow 00:11:55.029$ nib so it wasn't very clear story.

NOTE Confidence: 0.868898510932922

 $00{:}11{:}55.030 \dashrightarrow 00{:}11{:}57.102$ And when we looked at overall survival

NOTE Confidence: 0.868898510932922

00:11:57.102 --> 00:11:59.298 there wasn't any different so it wasn't

NOTE Confidence: 0.868898510932922

 $00:11:59.298 \longrightarrow 00:12:01.610$ clear that this definitive drug was really.

NOTE Confidence: 0.868898510932922

 $00:12:01.610 \longrightarrow 00:12:05.610$ Do anything special but.

NOTE Confidence: 0.868898510932922

00:12:05.610 --> 00:12:06.063 Well,

NOTE Confidence: 0.868898510932922

00:12:06.063 --> 00:12:08.328 that study was being conducted,

NOTE Confidence: 0.868898510932922

00:12:08.330 --> 00:12:11.018 so starting at around 2005 and

NOTE Confidence: 0.868898510932922

 $00{:}12{:}11.018 \dashrightarrow 00{:}12{:}14.219$ again this is the 15 year story.

NOTE Confidence: 0.868898510932922

 $00:12:14.220 \longrightarrow 00:12:15.948$ Scientists at several institutions.

NOTE Confidence: 0.868898510932922

 $00{:}12{:}15.948 \mathrel{--}{>} 00{:}12{:}18.108$ An realize that these patients

NOTE Confidence: 0.868898510932922

00:12:18.108 --> 00:12:20.140 who had these dramatic responses

NOTE Confidence: 0.868898510932922

 $00:12:20.140 \longrightarrow 00:12:22.438$ they did testing on their tumors,

NOTE Confidence: 0.868898510932922

 $00:12:22.440 \longrightarrow 00:12:24.856$ and they found that the tumors had a

NOTE Confidence: 0.868898510932922

00:12:24.856 --> 00:12:27.177 change in the Egypt far protein that

NOTE Confidence: 0.868898510932922

 $00:12:27.177 \longrightarrow 00:12:29.660$ was really related to a mutation that

 $00:12:29.660 \longrightarrow 00:12:31.976$ had happened in the cancer cells.

NOTE Confidence: 0.868898510932922

 $00:12:31.980 \longrightarrow 00:12:33.690$ So in the cancer cells,

NOTE Confidence: 0.868898510932922

 $00{:}12{:}33.690 \dashrightarrow 00{:}12{:}35.395$ EGFR became mutated and became

NOTE Confidence: 0.868898510932922

 $00:12:35.395 \longrightarrow 00:12:37.100$ turned on all the time.

NOTE Confidence: 0.868898510932922

00:12:37.100 --> 00:12:38.099 Constitu Tively active,

NOTE Confidence: 0.868898510932922

 $00:12:38.099 \longrightarrow 00:12:40.097$ and that looked like it was

NOTE Confidence: 0.868898510932922

 $00:12:40.097 \longrightarrow 00:12:41.867$ what was driving the cancer,

NOTE Confidence: 0.868898510932922

 $00{:}12{:}41.870 \dashrightarrow 00{:}12{:}43.495$ 'cause none of those mutations

NOTE Confidence: 0.868898510932922

 $00:12:43.495 \longrightarrow 00:12:45.557$ were found in the tumors are

NOTE Confidence: 0.868898510932922

00:12:45.557 --> 00:12:47.069 patients who didn't respond.

NOTE Confidence: 0.868898510932922

 $00:12:47.070 \longrightarrow 00:12:48.834$ But they were found in almost

NOTE Confidence: 0.868898510932922

00:12:48.834 --> 00:12:50.010 every tumor of patient,

NOTE Confidence: 0.868898510932922

 $00{:}12{:}50.010 \dashrightarrow 00{:}12{:}50.775$ who did respond,

NOTE Confidence: 0.868898510932922

 $00:12:50.775 \longrightarrow 00:12:52.903$ and so that led to sort of the

NOTE Confidence: 0.868898510932922

 $00:12:52.903 \longrightarrow 00:12:54.891$ ah ha moment that changed how we

00:12:54.891 --> 00:12:56.770 think about treating lung cancer,

NOTE Confidence: 0.868898510932922

 $00:12:56.770 \longrightarrow 00:12:59.010$ because when they went back and tested

NOTE Confidence: 0.868898510932922

00:12:59.010 --> 00:13:00.785 the tumor samples from the patients

NOTE Confidence: 0.868898510932922

 $00:13:00.785 \longrightarrow 00:13:02.950$ who had been on the I pass trial,

NOTE Confidence: 0.868898510932922

 $00:13:02.950 \longrightarrow 00:13:04.966$ they were able to come up with a

NOTE Confidence: 0.868898510932922

00:13:04.966 --> 00:13:06.458 very different story that patients

NOTE Confidence: 0.868898510932922

 $00:13:06.458 \longrightarrow 00:13:08.324$ whose tumors had the EGFR mutation

NOTE Confidence: 0.868898510932922

00:13:08.324 --> 00:13:10.296 did much better with your fitness,

NOTE Confidence: 0.868898510932922

 $00{:}13{:}10.300 --> 00{:}13{:}11.182 \ \mathrm{much\ higher\ responses}.$

NOTE Confidence: 0.868898510932922

00:13:11.182 --> 00:13:13.240 It worked for much longer than chemo,

NOTE Confidence: 0.868898510932922

 $00{:}13{:}13.240 \dashrightarrow 00{:}13{:}14.705$ but for people whose tumors

NOTE Confidence: 0.868898510932922

 $00:13:14.705 \longrightarrow 00:13:15.877$ did not have that,

NOTE Confidence: 0.868898510932922

 $00:13:15.880 \longrightarrow 00:13:18.687$ even though they otherwise look the same.

NOTE Confidence: 0.868898510932922

00:13:18.690 --> 00:13:20.170 It didn't work at all,

NOTE Confidence: 0.868898510932922

 $00:13:20.170 \longrightarrow 00:13:22.627$ and that was the first moment for us in

NOTE Confidence: 0.868898510932922

 $00:13:22.627 \longrightarrow 00:13:25.197$ lung cancer to realize we needed to look.

00:13:25.200 --> 00:13:27.006 We couldn't just determine based on

NOTE Confidence: 0.868898510932922

 $00{:}13{:}27.006 \dashrightarrow 00{:}13{:}29.117$ what the patient looked like or what

NOTE Confidence: 0.868898510932922

 $00:13:29.117 \longrightarrow 00:13:31.112$ their tumor look like under the microscope.

NOTE Confidence: 0.868898510932922

 $00:13:31.120 \longrightarrow 00:13:32.172$ We needed to test,

NOTE Confidence: 0.868898510932922

 $00:13:32.172 \longrightarrow 00:13:34.669$ and if we could find the EGFR mutation,

NOTE Confidence: 0.868898510932922

 $00:13:34.670 \longrightarrow 00:13:36.609$ then we could actually treat the tumors

NOTE Confidence: 0.868898510932922

 $00:13:36.609 \longrightarrow 00:13:38.230$ differently and have better outcomes.

NOTE Confidence: 0.868898510932922

 $00{:}13{:}38.230 \dashrightarrow 00{:}13{:}40.000$ And I'm focusing on EGFR today,

NOTE Confidence: 0.868898510932922

 $00:13:40.000 \longrightarrow 00:13:41.984$ but we now know to think about and

NOTE Confidence: 0.868898510932922

 $00{:}13{:}41.984 \dashrightarrow 00{:}13{:}43.951$ look for mutations in seven different

NOTE Confidence: 0.868898510932922

 $00:13:43.951 \longrightarrow 00:13:46.063$ genes in lung cancer is actually

NOTE Confidence: 0.853055417537689

00:13:46.127 --> 00:13:48.460 closer to 10 now, but we have 7 where

NOTE Confidence: 0.853055417537689

 $00{:}13{:}48.460 \dashrightarrow 00{:}13{:}49.810$ we have FDA approved medications

NOTE Confidence: 0.853055417537689

 $00:13:49.870 \longrightarrow 00:13:51.984$ and we can think about using those

NOTE Confidence: 0.853055417537689

 $00:13:51.984 \longrightarrow 00:13:53.360$ instead of chemotherapy upfront.

 $00:13:53.360 \longrightarrow 00:13:55.848$ So this was a complete revolution in how

NOTE Confidence: 0.853055417537689

00:13:55.848 --> 00:13:58.258 we think about treating lung cancer,

NOTE Confidence: 0.853055417537689

 $00:13:58.260 \longrightarrow 00:14:01.410$ and this study really was what God is there.

NOTE Confidence: 0.853055417537689

 $00:14:01.410 \longrightarrow 00:14:03.186$ So I'm focusing on that quite

NOTE Confidence: 0.853055417537689

 $00:14:03.186 \longrightarrow 00:14:05.610$ a bit in this particular trial,

NOTE Confidence: 0.853055417537689

 $00:14:05.610 \longrightarrow 00:14:07.710$ by selecting the way they did,

NOTE Confidence: 0.853055417537689

 $00:14:07.710 \longrightarrow 00:14:09.948$ about 60% of patients had EGFR

NOTE Confidence: 0.853055417537689

00:14:09.948 --> 00:14:11.821 mutations and those numbers vary

NOTE Confidence: 0.853055417537689

 $00:14:11.821 \longrightarrow 00:14:14.341$ around the world and based on who the

NOTE Confidence: 0.853055417537689

 $00:14:14.341 \longrightarrow 00:14:16.458$ person is who's gotten the cancer,

NOTE Confidence: 0.853055417537689

 $00{:}14{:}16.460 \rightarrow 00{:}14{:}19.610$ but it's not specific dress to women in Asia,

NOTE Confidence: 0.853055417537689

 $00{:}14{:}19.610 \dashrightarrow 00{:}14{:}22.410$ have a lot of my patients here in California,

NOTE Confidence: 0.853055417537689

 $00:14:22.410 \longrightarrow 00:14:23.806$ wouldn't fit that demographic.

NOTE Confidence: 0.853055417537689

00:14:23.806 --> 00:14:26.470 But yet there tumor hasn't EGFR mutation,

NOTE Confidence: 0.853055417537689

 $00:14:26.470 \longrightarrow 00:14:29.678$ and they respond as if anybody else who

NOTE Confidence: 0.853055417537689

 $00{:}14{:}29.678 \dashrightarrow 00{:}14{:}32.986$ has that EGFR mutation in their tumor.

 $00:14:32.990 \longrightarrow 00:14:35.666$ We've now had multiple other studies

NOTE Confidence: 0.853055417537689

 $00{:}14{:}35.666 \dashrightarrow 00{:}14{:}39.231$ that have looked at giving EGFR TK I

NOTE Confidence: 0.853055417537689

 $00{:}14{:}39.231 \dashrightarrow 00{:}14{:}41.769$ drugs versus chemo in patients whose

NOTE Confidence: 0.853055417537689

00:14:41.849 --> 00:14:44.147 tumor have in Egypt are mutation

NOTE Confidence: 0.853055417537689

 $00:14:44.147 \longrightarrow 00:14:47.800$ and you can see that in all of them,

NOTE Confidence: 0.853055417537689

 $00:14:47.800 \longrightarrow 00:14:50.754$ the progression free survival time was much,

NOTE Confidence: 0.853055417537689

00:14:50.760 --> 00:14:53.714 much better with the Egypt far drug,

NOTE Confidence: 0.853055417537689

 $00:14:53.720 \longrightarrow 00:14:55.676$ but it wasn't forever,

NOTE Confidence: 0.853055417537689

 $00:14:55.676 \longrightarrow 00:14:59.047$ and that's because we get resistance and

NOTE Confidence: 0.853055417537689

00:14:59.047 --> 00:15:02.600 this is a slide goes back a little bit now,

NOTE Confidence: 0.853055417537689

 $00:15:02.600 \longrightarrow 00:15:05.030$ but. Was figured out was that.

NOTE Confidence: 0.853055417537689

 $00{:}15{:}05.030 \dashrightarrow 00{:}15{:}07.982$ In order for a cancer to have an EGFR

NOTE Confidence: 0.853055417537689

 $00:15:07.982 \dashrightarrow 00:15:10.944$ mutation that we don't know why that happens,

NOTE Confidence: 0.853055417537689

 $00:15:10.950 \longrightarrow 00:15:13.379$ we're still trying to figure that out.

NOTE Confidence: 0.853055417537689

 $00:15:13.380 \longrightarrow 00:15:15.420$ But when the tumor has

 $00:15:15.420 \longrightarrow 00:15:17.052$ that easier for mutation.

NOTE Confidence: 0.853055417537689

 $00{:}15{:}17.060 \dashrightarrow 00{:}15{:}19.055$ The tumors respond really well

NOTE Confidence: 0.853055417537689

00:15:19.055 --> 00:15:21.480 to an EGFR tyrosine kinase drug,

NOTE Confidence: 0.853055417537689

 $00:15:21.480 \longrightarrow 00:15:23.088$ but overtime resistance develops

NOTE Confidence: 0.853055417537689

00:15:23.088 --> 00:15:24.696 because new mutations happen,

NOTE Confidence: 0.853055417537689

 $00:15:24.700 \longrightarrow 00:15:27.486$ and in the setting of EGFR with

NOTE Confidence: 0.853055417537689

 $00:15:27.486 \longrightarrow 00:15:29.520$ these initially developed EGFR drugs,

NOTE Confidence: 0.853055417537689

 $00:15:29.520 \longrightarrow 00:15:32.736$ most of the time when those stopped working,

NOTE Confidence: 0.853055417537689

 $00:15:32.740 \longrightarrow 00:15:35.146$ it was because another EGFR mutation

NOTE Confidence: 0.853055417537689

 $00:15:35.146 \longrightarrow 00:15:37.560$ developed, and that was called T790M.

NOTE Confidence: 0.853055417537689

00:15:37.560 --> 00:15:40.283 So we spent many years trying to

NOTE Confidence: 0.853055417537689

00:15:40.283 --> 00:15:43.188 figure out a way to treat T790M.

NOTE Confidence: 0.853055417537689

 $00:15:43.190 \longrightarrow 00:15:46.070$ I have a lot of old studies that

NOTE Confidence: 0.853055417537689

 $00:15:46.070 \longrightarrow 00:15:47.650$ didn't go anywhere.

NOTE Confidence: 0.853055417537689

00:15:47.650 --> 00:15:49.445 And many medications we worked

NOTE Confidence: 0.853055417537689

 $00:15:49.445 \longrightarrow 00:15:51.959$ with but then came along this drug.

 $00:15:51.960 \longrightarrow 00:15:53.037$ Oh, summer NAMM.

NOTE Confidence: 0.853055417537689

 $00:15:53.037 \longrightarrow 00:15:55.550$ Which is the third generation EGFR drug.

NOTE Confidence: 0.853055417537689

 $00:15:55.550 \longrightarrow 00:15:58.056$ And it works when T790M is there.

NOTE Confidence: 0.853055417537689

 $00:15:58.060 \longrightarrow 00:16:00.573$ So what these are showing is you've

NOTE Confidence: 0.85305541753768900:16:00.573 --> 00:16:01.650 got the line. NOTE Confidence: 0.853055417537689

00:16:01.650 --> 00:16:03.810 If you've got lines above it,

NOTE Confidence: 0.853055417537689

 $00:16:03.810 \longrightarrow 00:16:05.958$ that means that the tumors grew,

NOTE Confidence: 0.853055417537689

00:16:05.960 --> 00:16:08.198 but everything below the line means

NOTE Confidence: 0.853055417537689

 $00{:}16{:}08.198 \dashrightarrow 00{:}16{:}10.829$ that the tumor shrunk so you can see

NOTE Confidence: 0.853055417537689

 $00{:}16{:}10.829 \dashrightarrow 00{:}16{:}13.365$ that for most of these the tumors were

NOTE Confidence: 0.853055417537689

00:16:13.365 --> 00:16:15.510 shrinking when a patient received

NOTE Confidence: 0.853055417537689

00:16:15.510 --> 00:16:17.746 oh summerton. If after they die.

NOTE Confidence: 0.853055417537689

 $00:16:17.746 \longrightarrow 00:16:19.810$ Already had either the jefit neighbor

NOTE Confidence: 0.853055417537689

 $00:16:19.879 \longrightarrow 00:16:21.769$ alot nip those so if they've been

NOTE Confidence: 0.853055417537689

00:16:21.769 --> 00:16:24.249 on a first generation drug and it

 $00:16:24.249 \longrightarrow 00:16:25.049$ stopped working.

NOTE Confidence: 0.853055417537689

 $00:16:25.050 \longrightarrow 00:16:27.180$ And then they took this third

NOTE Confidence: 0.853055417537689

 $00:16:27.180 \longrightarrow 00:16:28.600$ generation drug osimertinib most

NOTE Confidence: 0.853055417537689

 $00:16:28.666 \longrightarrow 00:16:30.328$ of the time the tumor shrunk,

NOTE Confidence: 0.853055417537689

 $00:16:30.330 \longrightarrow 00:16:32.646$ and that's because we were actually

NOTE Confidence: 0.853055417537689

 $00:16:32.646 \longrightarrow 00:16:34.190$ getting that T790M target.

NOTE Confidence: 0.853055417537689

 $00:16:34.190 \longrightarrow 00:16:36.710$ So to prove that there was a trial

NOTE Confidence: 0.853055417537689

00:16:36.710 --> 00:16:38.840 for patients who had been found

NOTE Confidence: 0.853055417537689

 $00:16:38.840 \longrightarrow 00:16:40.976$ to have Egypt permutation in their

NOTE Confidence: 0.853055417537689

 $00:16:41.044 \longrightarrow 00:16:42.588$ tumor at the beginning,

NOTE Confidence: 0.853055417537689

 $00:16:42.590 \longrightarrow 00:16:44.690$ they started on a first or

NOTE Confidence: 0.853055417537689

00:16:44.690 --> 00:16:46.090 second generation EGFR medicine,

NOTE Confidence: 0.853055417537689

00:16:46.090 --> 00:16:48.190 and then when it stopped working,

NOTE Confidence: 0.853055417537689

00:16:48.190 --> 00:16:50.608 they were on a trial randomized

NOTE Confidence: 0.853055417537689

 $00:16:50.608 \longrightarrow 00:16:52.220$ to either ghetto smart

NOTE Confidence: 0.841263711452484

00:16:52.295 --> 00:16:54.310 nib or to get chemotherapy.

 $00:16:54.310 \longrightarrow 00:16:56.788$ And what this showed us was that

NOTE Confidence: 0.841263711452484

 $00:16:56.788 \longrightarrow 00:16:58.280$ progression free survival was

NOTE Confidence: 0.841263711452484

00:16:58.280 --> 00:17:00.463 clearly better with, uh, oh, summer.

NOTE Confidence: 0.841263711452484

 $00:17:00.463 \longrightarrow 00:17:01.907$ New versus the chemotherapy.

NOTE Confidence: 0.841263711452484

 $00:17:01.910 \longrightarrow 00:17:04.360$ So it became standard that you would

NOTE Confidence: 0.841263711452484

 $00:17:04.360 \longrightarrow 00:17:06.581$ test when someone was diognosed if

NOTE Confidence: 0.841263711452484

00:17:06.581 --> 00:17:08.783 their tumor had the EGFR mutation,

NOTE Confidence: 0.841263711452484

 $00:17:08.790 \longrightarrow 00:17:10.580$ you would start them on

NOTE Confidence: 0.841263711452484

 $00:17:10.580 \longrightarrow 00:17:12.770$ our lot nib or defend him,

NOTE Confidence: 0.841263711452484

00:17:12.770 --> 00:17:14.580 and when that stopped working,

NOTE Confidence: 0.841263711452484

 $00:17:14.580 \longrightarrow 00:17:17.114$ if the T790M mutation was there then,

NOTE Confidence: 0.841263711452484

00:17:17.120 --> 00:17:19.430 oh, Smyrna was a better choice

NOTE Confidence: 0.841263711452484

 $00{:}17{:}19.430 \dashrightarrow 00{:}17{:}22.060$ than going with the chemotherapy.

NOTE Confidence: 0.841263711452484

00:17:22.060 --> 00:17:24.400 But finding the T790M wasn't perfect,

NOTE Confidence: 0.841263711452484

 $00:17:24.400 \longrightarrow 00:17:26.350$ so there were several different

 $00:17:26.350 \longrightarrow 00:17:27.910$ ways to do that.

NOTE Confidence: 0.841263711452484

00:17:27.910 --> 00:17:31.078 You could either look with tissue.

NOTE Confidence: 0.841263711452484

 $00:17:31.080 \longrightarrow 00:17:33.054$ Or in the blood with plasma testing.

NOTE Confidence: 0.841263711452484

 $00:17:33.060 \longrightarrow 00:17:35.034$ And we also did some urine testing,

NOTE Confidence: 0.841263711452484

 $00:17:35.040 \longrightarrow 00:17:37.264$ but we don't do that so much anymore

NOTE Confidence: 0.841263711452484

 $00:17:37.264 \longrightarrow 00:17:39.523$ and what this is showing is that most

NOTE Confidence: 0.841263711452484

 $00:17:39.523 \longrightarrow 00:17:41.819$ of the time you could find that T790M

NOTE Confidence: 0.841263711452484

 $00:17:41.819 \longrightarrow 00:17:43.907$ in the tissue and in the plasma,

NOTE Confidence: 0.841263711452484

 $00{:}17{:}43.907 \dashrightarrow 00{:}17{:}45.792$ but there were sometimes you

NOTE Confidence: 0.841263711452484

 $00:17:45.792 \longrightarrow 00:17:48.338$ could only find it in that issue.

NOTE Confidence: 0.841263711452484

 $00{:}17{:}48.340 \dashrightarrow 00{:}17{:}49.905$ And that's because it wasn't

NOTE Confidence: 0.841263711452484

 $00:17:49.905 \longrightarrow 00:17:52.150$ being shed out into the DNA wasn't

NOTE Confidence: 0.841263711452484

 $00:17:52.150 \longrightarrow 00:17:53.695$ going out into the blood.

NOTE Confidence: 0.841263711452484

00:17:53.700 --> 00:17:55.877 Or you only found in the plasma,

NOTE Confidence: 0.841263711452484

 $00:17:55.880 \longrightarrow 00:17:57.124$ and that's perhaps because

NOTE Confidence: 0.841263711452484

 $00:17:57.124 \longrightarrow 00:17:58.679$ when you did the biopsy,

00:17:58.680 --> 00:18:01.160 you didn't get enough of the cancer cells,

NOTE Confidence: 0.841263711452484

 $00:18:01.160 \longrightarrow 00:18:02.835$ or there's some heterogeneity meeting

NOTE Confidence: 0.841263711452484

 $00:18:02.835 \longrightarrow 00:18:05.423$ that not all of the cancer cells change

NOTE Confidence: 0.841263711452484

 $00:18:05.423 \longrightarrow 00:18:07.700$ at the same time in the same way,

NOTE Confidence: 0.841263711452484

 $00:18:07.700 \longrightarrow 00:18:09.716$ so you can have areas that are

NOTE Confidence: 0.841263711452484

 $00:18:09.716 \longrightarrow 00:18:11.429$ resistant because of this mutation.

NOTE Confidence: 0.841263711452484

 $00:18:11.430 \longrightarrow 00:18:12.980$ Other areas that are not,

NOTE Confidence: 0.841263711452484

 $00{:}18{:}12.980 --> 00{:}18{:}15.143$ and so I'm not going to go

NOTE Confidence: 0.841263711452484

 $00:18:15.143 \longrightarrow 00:18:17.028$ into that today in one hour.

NOTE Confidence: 0.841263711452484

00:18:17.030 --> 00:18:18.258 You can't cover everything,

NOTE Confidence: 0.841263711452484

 $00:18:18.258 \longrightarrow 00:18:20.484$ but I wanted to at least mention

NOTE Confidence: 0.841263711452484

 $00:18:20.484 \longrightarrow 00:18:22.079$ this because it is important

NOTE Confidence: 0.841263711452484

00:18:22.079 --> 00:18:23.950 to know that when you have.

NOTE Confidence: 0.841263711452484 00:18:23.950 --> 00:18:24.241 Cancer. NOTE Confidence: 0.841263711452484

 $00:18:24.241 \longrightarrow 00:18:25.696$ It's evolving overtime an the

00:18:25.696 --> 00:18:27.599 way that we find out more about

NOTE Confidence: 0.841263711452484

00:18:27.599 --> 00:18:29.181 it as we need to sample it.

NOTE Confidence: 0.841263711452484

 $00:18:29.190 \longrightarrow 00:18:31.017$ An historically that was always with tissue.

NOTE Confidence: 0.841263711452484

 $00:18:31.020 \longrightarrow 00:18:33.036$ But now we can look at it with

NOTE Confidence: 0.841263711452484

 $00:18:33.036 \longrightarrow 00:18:34.429$ these plasma tests as well,

NOTE Confidence: 0.841263711452484

00:18:34.430 --> 00:18:36.536 but that's all I'm going to

NOTE Confidence: 0.841263711452484

 $00:18:36.536 \longrightarrow 00:18:37.940$ say on that topic.

NOTE Confidence: 0.841263711452484

 $00:18:37.940 \longrightarrow 00:18:38.247$ OK,

NOTE Confidence: 0.841263711452484

 $00:18:38.247 \longrightarrow 00:18:40.703$ so we know to test we know to

NOTE Confidence: 0.841263711452484

 $00:18:40.703 \longrightarrow 00:18:43.110$ give a first generation drugs.

NOTE Confidence: 0.841263711452484

 $00:18:43.110 \longrightarrow 00:18:44.422$ We know T790M develops.

NOTE Confidence: 0.841263711452484

 $00:18:44.422 \longrightarrow 00:18:47.530$ So then you use the third generation drug.

NOTE Confidence: 0.841263711452484

00:18:47.530 --> 00:18:49.380 But of course people wondered.

NOTE Confidence: 0.841263711452484

 $00:18:49.380 \longrightarrow 00:18:51.498$ Why don't we just give this

NOTE Confidence: 0.841263711452484

 $00:18:51.498 \longrightarrow 00:18:53.364$ third generation drug 1st and

NOTE Confidence: 0.841263711452484

 $00:18:53.364 \longrightarrow 00:18:54.908$ then T790M won't develop?

 $00:18:54.910 \longrightarrow 00:18:57.178$ So that was this flora trial

NOTE Confidence: 0.841263711452484

 $00:18:57.178 \longrightarrow 00:18:59.209$ where patients who are newly

NOTE Confidence: 0.841263711452484

 $00:18:59.209 \longrightarrow 00:19:01.837$ diagnosed and found to have EGFR

NOTE Confidence: 0.841263711452484

00:19:01.837 --> 00:19:04.025 mutation either goto summer nib

NOTE Confidence: 0.841263711452484

 $00:19:04.025 \dashrightarrow 00:19:06.419$ or got a first generation drugs.

NOTE Confidence: 0.841263711452484

 $00:19:06.420 \longrightarrow 00:19:08.020$ And that study clearly showed

NOTE Confidence: 0.841263711452484

 $00:19:08.020 \longrightarrow 00:19:10.089$ that to Smyrna was better for

NOTE Confidence: 0.841263711452484

 $00:19:10.089 \longrightarrow 00:19:11.717$ that progression free survival,

NOTE Confidence: 0.841263711452484

 $00:19:11.720 \longrightarrow 00:19:13.705$ meaning that it took longer

NOTE Confidence: 0.841263711452484

 $00:19:13.705 \longrightarrow 00:19:16.120$ before the cancer started to grow.

NOTE Confidence: 0.84126371145248400:19:16.120 --> 00:19:16.493 However, NOTE Confidence: 0.841263711452484

00:19:16.493 --> 00:19:19.104 it also led to overall survival improvement,

NOTE Confidence: 0.841263711452484

 $00{:}19{:}19.110 \dashrightarrow 00{:}19{:}20.875$ and that's because even though

NOTE Confidence: 0.841263711452484

 $00:19:20.875 \longrightarrow 00:19:22.640$ theoretically if you started on

NOTE Confidence: 0.841263711452484

 $00:19:22.703 \longrightarrow 00:19:24.663$ a first generation drugs and

 $00:19:24.663 \longrightarrow 00:19:25.839$ then developed resistance,

NOTE Confidence: 0.841263711452484

00:19:25.840 --> 00:19:28.360 you could go on to Summerton

NOTE Confidence: 0.841263711452484

 $00:19:28.360 \longrightarrow 00:19:30.760$ if if it was T790M.

NOTE Confidence: 0.841263711452484

 $00:19:30.760 \longrightarrow 00:19:32.680$ There are other things that cause

NOTE Confidence: 0.841263711452484

 $00:19:32.680 \longrightarrow 00:19:34.721$ resistance an so delaying that time

NOTE Confidence: 0.841263711452484

00:19:34.721 --> 00:19:36.466 to resistance with the osimertinib,

NOTE Confidence: 0.841263711452484

 $00:19:36.470 \longrightarrow 00:19:38.808$ even when it's not resistance due to

NOTE Confidence: 0.841263711452484

00:19:38.808 --> 00:19:40.500 T790M overall survival was better,

NOTE Confidence: 0.841263711452484

 $00:19:40.500 \longrightarrow 00:19:42.558$ and so this has become the

NOTE Confidence: 0.841263711452484

 $00:19:42.558 \longrightarrow 00:19:44.200$ standard approach in the US.

NOTE Confidence: 0.841263711452484

 $00{:}19{:}44.200 \dashrightarrow 00{:}19{:}45.880$ But unfortunately as you see,

NOTE Confidence: 0.834039330482483

 $00:19:45.880 \longrightarrow 00:19:47.896$ these curves eventually do come down.

NOTE Confidence: 0.834039330482483

00:19:47.900 --> 00:19:49.380 We're not curing people,

NOTE Confidence: 0.834039330482483

 $00:19:49.380 \longrightarrow 00:19:51.600$ and therefore we still need to

NOTE Confidence: 0.834039330482483

 $00:19:51.674 \longrightarrow 00:19:53.828$ think about better things to do.

NOTE Confidence: 0.834039330482483

 $00:19:53.830 \longrightarrow 00:19:55.405$ So we kind of summarize

 $00:19:55.405 \longrightarrow 00:19:56.980$ this first part about EGFR.

NOTE Confidence: 0.834039330482483

 $00:19:56.980 \longrightarrow 00:19:59.815$ We need to test to find the Jaffar mutation,

NOTE Confidence: 0.834039330482483

 $00:19:59.820 \longrightarrow 00:20:01.080$ and usually that's done

NOTE Confidence: 0.834039330482483

 $00:20:01.080 \longrightarrow 00:20:02.340$ with the tissue testing.

NOTE Confidence: 0.834039330482483

 $00:20:02.340 \longrightarrow 00:20:04.342$ But we can also sometimes find it

NOTE Confidence: 0.834039330482483

 $00:20:04.342 \longrightarrow 00:20:06.428$ in the plasma with blood testing.

NOTE Confidence: 0.834039330482483

00:20:06.430 --> 00:20:08.320 If you find the Jaffar mutation,

NOTE Confidence: 0.834039330482483

 $00:20:08.320 \longrightarrow 00:20:09.900$ we start on an EGFR.

NOTE Confidence: 0.834039330482483

 $00:20:09.900 \longrightarrow 00:20:11.168$ TKI osimertinib is what

NOTE Confidence: 0.834039330482483

 $00:20:11.168 \longrightarrow 00:20:13.360$ we tend to use in the US,

NOTE Confidence: 0.834039330482483

 $00:20:13.360 \longrightarrow 00:20:15.358$ but there are multiple other EGFR

NOTE Confidence: 0.834039330482483

 $00:20:15.358 \longrightarrow 00:20:17.032$ tyrosine kinase drugs which are

NOTE Confidence: 0.834039330482483

 $00:20:17.032 \dashrightarrow 00:20:18.718$ approved in the US and globally,

NOTE Confidence: 0.834039330482483

 $00{:}20{:}18.720 \dashrightarrow 00{:}20{:}22.600$ and so those are still used as well.

NOTE Confidence: 0.834039330482483

00:20:22.600 --> 00:20:23.964 And as I mentioned,

 $00:20:23.964 \longrightarrow 00:20:25.669$ we're still thwarted by resistance.

NOTE Confidence: 0.834039330482483

00:20:25.670 --> 00:20:26.702 So with Osimertinib,

NOTE Confidence: 0.834039330482483

 $00:20:26.702 \longrightarrow 00:20:29.110$ it's not such a simple story like

NOTE Confidence: 0.834039330482483

 $00:20:29.175 \longrightarrow 00:20:31.464$ it is with the 1st generation drugs,

NOTE Confidence: 0.834039330482483

 $00:20:31.470 \longrightarrow 00:20:33.510$ where we get one resistance pathway,

NOTE Confidence: 0.834039330482483

 $00{:}20{:}33.510 \dashrightarrow 00{:}20{:}35.424$ which is that T790M resistance as

NOTE Confidence: 0.834039330482483

 $00:20:35.424 \longrightarrow 00:20:37.122$ the predominant one when there's

NOTE Confidence: 0.834039330482483

 $00{:}20{:}37.122 \dashrightarrow 00{:}20{:}39.270$ osimertinib resistance are a lot of

NOTE Confidence: 0.834039330482483

 $00:20:39.270 \longrightarrow 00:20:41.009$ different things that can happen.

NOTE Confidence: 0.834039330482483

00:20:41.010 --> 00:20:43.738 You can get other EGFR mutations and C.

NOTE Confidence: 0.834039330482483

00:20:43.740 --> 00:20:47.490 797 and T 797 S is the most common of those,

NOTE Confidence: 0.834039330482483

 $00{:}20{:}47.490 \to 00{:}20{:}50.202$ but there are a lot of other pathways

NOTE Confidence: 0.834039330482483

 $00:20:50.202 \longrightarrow 00:20:52.650$ that can go be abnormal as well.

NOTE Confidence: 0.834039330482483

00:20:52.650 --> 00:20:54.320 And so it's quite complex,

NOTE Confidence: 0.834039330482483

 $00:20:54.320 \longrightarrow 00:20:56.750$ and there's a lot of research

NOTE Confidence: 0.834039330482483

 $00:20:56.750 \longrightarrow 00:20:57.965$ being done there.

00:20:57.970 --> 00:21:00.245 And this is just a graphic showing

NOTE Confidence: 0.834039330482483

 $00:21:00.245 \longrightarrow 00:21:02.110$ some of those resistances,

NOTE Confidence: 0.834039330482483

 $00:21:02.110 \longrightarrow 00:21:04.357$ so I talked about the T790M here

NOTE Confidence: 0.834039330482483

00:21:04.357 --> 00:21:06.620 with first and second generation.

NOTE Confidence: 0.834039330482483

 $00:21:06.620 \longrightarrow 00:21:08.530$ But there's also overexpression of

NOTE Confidence: 0.834039330482483

00:21:08.530 --> 00:21:11.226 met and mutations in other genes and

NOTE Confidence: 0.834039330482483

00:21:11.226 --> 00:21:13.386 also a transformation to small cell,

NOTE Confidence: 0.834039330482483

 $00{:}21{:}13.390 \to 00{:}21{:}15.270$ so it's not always straightforward.

NOTE Confidence: 0.834039330482483

00:21:15.270 --> 00:21:17.508 And then with Osama RNIB there

NOTE Confidence: 0.834039330482483

 $00:21:17.508 \longrightarrow 00:21:20.170$ can be changes in what the EGFR.

NOTE Confidence: 0.834039330482483

00:21:20.170 --> 00:21:22.006 Just kind of percentage of expression

NOTE Confidence: 0.834039330482483

 $00:21:22.006 \longrightarrow 00:21:24.354$ of the protein is it's never gone

NOTE Confidence: 0.834039330482483

 $00{:}21{:}24.354 \dashrightarrow 00{:}21{:}26.129$ that Egypt permutations never gone,

NOTE Confidence: 0.834039330482483

 $00:21:26.130 \longrightarrow 00:21:28.447$ but the way it's expressed is different.

NOTE Confidence: 0.834039330482483

 $00:21:28.450 \longrightarrow 00:21:30.767$ But you also can have met overexpression

00:21:30.767 --> 00:21:32.992 is fairly common, or changes in BRAF.

NOTE Confidence: 0.834039330482483

 $00:21:32.992 \longrightarrow 00:21:34.840$ We just were talking about a

NOTE Confidence: 0.834039330482483

 $00:21:34.905 \longrightarrow 00:21:36.060$ case this morning.

NOTE Confidence: 0.834039330482483

00:21:36.060 --> 00:21:37.348 Stanford, with another patient

NOTE Confidence: 0.834039330482483

00:21:37.348 --> 00:21:39.700 who had been on no smart Niman,

NOTE Confidence: 0.834039330482483

00:21:39.700 --> 00:21:42.017 then developed in be reputation and many,

NOTE Confidence: 0.834039330482483

 $00:21:42.020 \longrightarrow 00:21:42.648$ many others.

NOTE Confidence: 0.834039330482483

00:21:42.648 --> 00:21:44.846 So again, it's quite complex as we

NOTE Confidence: 0.834039330482483

00:21:44.846 --> 00:21:46.976 talk about bypass tracks in Egypt,

NOTE Confidence: 0.834039330482483

 $00:21:46.980 \longrightarrow 00:21:48.072$ our target alterations.

NOTE Confidence: 0.834039330482483

 $00:21:48.072 \longrightarrow 00:21:50.256$ So then what do you do?

NOTE Confidence: 0.834039330482483

 $00:21:50.260 \longrightarrow 00:21:51.148$ You gotta think.

NOTE Confidence: 0.834039330482483

00:21:51.148 --> 00:21:52.934 Outside of that, here's the target.

NOTE Confidence: 0.834039330482483

 $00:21:52.934 \longrightarrow 00:21:53.828$ Hit the target.

NOTE Confidence: 0.834039330482483

 $00:21:53.830 \longrightarrow 00:21:55.606$ We've got to be broader that.

NOTE Confidence: 0.834039330482483

00:21:55.610 --> 00:21:56.794 Here's a target hit.

 $00:21:56.794 \longrightarrow 00:21:57.978$ The target is great,

NOTE Confidence: 0.834039330482483

 $00:21:57.980 \longrightarrow 00:21:59.465$ and it's helped people tremendously

NOTE Confidence: 0.834039330482483

 $00:21:59.465 \longrightarrow 00:22:01.250$ for an extended period of time.

NOTE Confidence: 0.834039330482483

 $00:22:01.250 \longrightarrow 00:22:03.626$ But we want to extend it even further.

NOTE Confidence: 0.834039330482483

 $00:22:03.630 \longrightarrow 00:22:05.110$ So what can we do?

NOTE Confidence: 0.834039330482483

 $00:22:05.110 \longrightarrow 00:22:06.595$ We try something else so

NOTE Confidence: 0.834039330482483

 $00:22:06.595 \longrightarrow 00:22:07.783$ we have anti angiogenesis.

NOTE Confidence: 0.834039330482483

 $00:22:07.790 \longrightarrow 00:22:08.970$ Is one of those,

NOTE Confidence: 0.834039330482483

 $00:22:08.970 \longrightarrow 00:22:10.740$ and that's getting at the fact

NOTE Confidence: 0.834039330482483

00:22:10.804 --> 00:22:12.574 that in order for some tumor

NOTE Confidence: 0.834039330482483

00:22:12.574 --> 00:22:14.320 cells to grow into a tumor,

NOTE Confidence: 0.834039330482483

 $00:22:14.320 \longrightarrow 00:22:16.984$ they need to have blood vessels to feed them.

NOTE Confidence: 0.834039330482483

 $00{:}22{:}16.990 \dashrightarrow 00{:}22{:}18.480$ And so this is angiogenesis.

NOTE Confidence: 0.834039330482483

 $00:22:18.480 \longrightarrow 00:22:20.010$ And that's where we get.

NOTE Confidence: 0.834039330482483

00:22:20.010 --> 00:22:23.420 Drugs such as Bevis is a map and many others,

 $00:22:23.420 \longrightarrow 00:22:25.688$ and there's been a ton of trials

NOTE Confidence: 0.834039330482483

00:22:25.688 --> 00:22:27.848 noticing that in the EGFR setting,

NOTE Confidence: 0.834039330482483

 $00:22:27.850 \longrightarrow 00:22:29.896$ veg F is also very important,

NOTE Confidence: 0.834039330482483

 $00:22:29.900 \longrightarrow 00:22:32.650$ and so a lot of trials looking at in a

NOTE Confidence: 0.834039330482483

00:22:32.727 --> 00:22:35.695 patient with Egypt or mutant lung cancer,

NOTE Confidence: 0.800156116485596

00:22:35.700 --> 00:22:37.740 you can give the EGFR medicine,

NOTE Confidence: 0.800156116485596

 $00:22:37.740 \longrightarrow 00:22:39.788$ but you can also give it with anti

NOTE Confidence: 0.800156116485596

 $00:22:39.788 \longrightarrow 00:22:42.078$ veg F therapy and that theoretically

NOTE Confidence: 0.800156116485596

00:22:42.078 --> 00:22:44.213 should make a big difference.

NOTE Confidence: 0.800156116485596

 $00:22:44.220 \longrightarrow 00:22:47.190$ So one of the first trials was the beta

NOTE Confidence: 0.800156116485596

 $00:22:47.190 \longrightarrow 00:22:50.377$ trial led by Doctor Herbst an in this study.

NOTE Confidence: 0.800156116485596

00:22:50.380 --> 00:22:52.702 It wasn't selected for patients with

NOTE Confidence: 0.800156116485596

 $00{:}22{:}52.702 \dashrightarrow 00{:}22{:}54.952$ EGFR mutation because this was done

NOTE Confidence: 0.800156116485596

 $00:22:54.952 \longrightarrow 00:22:57.185$ before we knew about the EGFR mutations

NOTE Confidence: 0.800156116485596

00:22:57.185 --> 00:22:59.875 and we were just giving our lot nib to

NOTE Confidence: 0.800156116485596

00:22:59.875 --> 00:23:01.796 people after they already had chemo.

00:23:01.796 --> 00:23:04.160 But this showed that even without that,

NOTE Confidence: 0.800156116485596

 $00:23:04.160 \longrightarrow 00:23:06.584$ giving the bevacizumab added to the

NOTE Confidence: 0.800156116485596

00:23:06.584 --> 00:23:08.200 Erlotinib improved progression free

NOTE Confidence: 0.800156116485596

00:23:08.260 --> 00:23:10.409 survival Anne in the small group of

NOTE Confidence: 0.800156116485596

 $00:23:10.409 \longrightarrow 00:23:12.529$ patients who would went back and found

NOTE Confidence: 0.800156116485596

 $00:23:12.529 \longrightarrow 00:23:14.591$ that they actually had an EGFR mutation.

NOTE Confidence: 0.800156116485596

00:23:14.591 --> 00:23:17.048 There was a signal that there might

NOTE Confidence: 0.800156116485596

 $00:23:17.048 \longrightarrow 00:23:19.678$ even be an overall survival benefit by

NOTE Confidence: 0.800156116485596

 $00{:}23{:}19.678 \dashrightarrow 00{:}23{:}22.598$ adding the beva system up to her latinum.

NOTE Confidence: 0.800156116485596

 $00:23:22.600 \longrightarrow 00:23:24.766$ So there's been multiple other trials.

NOTE Confidence: 0.800156116485596

 $00:23:24.770 \longrightarrow 00:23:28.019$ This was a phase two study done in Japan,

NOTE Confidence: 0.800156116485596

 $00:23:28.020 \longrightarrow 00:23:30.600$ where patients with EGFR mutant lung

NOTE Confidence: 0.800156116485596

 $00{:}23{:}30.600 \dashrightarrow 00{:}23{:}32.962$ cancer received either or lot nib

NOTE Confidence: 0.800156116485596

 $00:23:32.962 \longrightarrow 00:23:35.234$ the pill everyday or a lot in it.

NOTE Confidence: 0.800156116485596

 $00:23:35.240 \longrightarrow 00:23:37.238$ Plus the breakfast is a Babin

00:23:37.238 --> 00:23:39.210 beverages and Maps and antibody.

NOTE Confidence: 0.800156116485596

00:23:39.210 --> 00:23:41.370 It's given Ivy every three weeks,

NOTE Confidence: 0.800156116485596

 $00:23:41.370 \longrightarrow 00:23:43.416$ and this study showed a marked

NOTE Confidence: 0.800156116485596

 $00:23:43.416 \longrightarrow 00:23:45.340$ improvement in progression free survival.

NOTE Confidence: 0.800156116485596

 $00:23:45.340 \longrightarrow 00:23:47.145$ This actually led to approval

NOTE Confidence: 0.800156116485596

00:23:47.145 --> 00:23:48.950 of this combination in Europe,

NOTE Confidence: 0.800156116485596

 $00:23:48.950 \longrightarrow 00:23:50.870$ but not in the US.

NOTE Confidence: 0.800156116485596

00:23:50.870 --> 00:23:52.770 With longer term follow-up, though,

NOTE Confidence: 0.800156116485596

 $00:23:52.770 \longrightarrow 00:23:55.325$ it turned out that Despite that huge

NOTE Confidence: 0.800156116485596

00:23:55.325 --> 00:23:56.950 progression free survival benefit,

NOTE Confidence: 0.800156116485596

 $00:23:56.950 \longrightarrow 00:23:59.230$ there was no overall survival benefit,

NOTE Confidence: 0.800156116485596

 $00:23:59.230 \longrightarrow 00:24:02.198$ and so it's a little bit unclear what

NOTE Confidence: 0.800156116485596

 $00:24:02.198 \longrightarrow 00:24:04.690$ we're doing with this combination.

NOTE Confidence: 0.800156116485596

00:24:04.690 --> 00:24:07.035 Perhaps it's helpful to prolong the time

NOTE Confidence: 0.800156116485596

 $00:24:07.035 \longrightarrow 00:24:09.507$ that there are lot name stops working.

NOTE Confidence: 0.800156116485596

 $00:24:09.510 \longrightarrow 00:24:11.561$ It turns out that if you're on

00:24:11.561 --> 00:24:13.629 our lot nib and bevacizumab,

NOTE Confidence: 0.800156116485596

 $00:24:13.630 \longrightarrow 00:24:15.006$ the most common resistance

NOTE Confidence: 0.800156116485596

 $00:24:15.006 \longrightarrow 00:24:16.726$ pathway still is that T790M.

NOTE Confidence: 0.800156116485596

00:24:16.730 --> 00:24:18.794 So if you're on this combination

NOTE Confidence: 0.800156116485596

 $00:24:18.794 \longrightarrow 00:24:20.170$ and then resistance develops,

NOTE Confidence: 0.800156116485596

 $00:24:20.170 \longrightarrow 00:24:22.914$ you can go on to get the oh,

NOTE Confidence: 0.800156116485596

00:24:22.920 --> 00:24:24.942 Smyrna, but there are other things

NOTE Confidence: 0.800156116485596

 $00:24:24.942 \longrightarrow 00:24:26.696$ that happen besides T790M and

NOTE Confidence: 0.800156116485596

00:24:26.696 --> 00:24:28.426 then no overall survival benefit,

NOTE Confidence: 0.800156116485596

 $00:24:28.430 \longrightarrow 00:24:29.914$ so little bit unclear.

NOTE Confidence: 0.800156116485596

00:24:29.914 --> 00:24:32.600 There was a randomized phase three trial,

NOTE Confidence: 0.800156116485596

00:24:32.600 --> 00:24:33.676 also done in Japan,

NOTE Confidence: 0.800156116485596

 $00{:}24{:}33.676 {\:\dashrightarrow\:} 00{:}24{:}35.736$ which had very similar results in Nice

NOTE Confidence: 0.800156116485596

00:24:35.736 --> 00:24:37.316 progression free survival benefit,

NOTE Confidence: 0.800156116485596

 $00:24:37.320 \longrightarrow 00:24:38.900$ but no overall survival benefit.

 $00:24:38.900 \longrightarrow 00:24:40.148$ And this is news.

NOTE Confidence: 0.800156116485596

 $00{:}24{:}40.148 \operatorname{--}{>} 00{:}24{:}42.412$ This just was presented at our ASCO

NOTE Confidence: 0.800156116485596

 $00:24:42.412 \longrightarrow 00:24:44.964$ meeting in June of 2020 show we were

NOTE Confidence: 0.800156116485596

 $00:24:44.964 \longrightarrow 00:24:47.445$ waiting on this and now we kind of

NOTE Confidence: 0.800156116485596

00:24:47.445 --> 00:24:48.994 seeing this similar story emerging,

NOTE Confidence: 0.800156116485596

 $00:24:48.994 \longrightarrow 00:24:51.810$ but it's more complex than that in the US.

NOTE Confidence: 0.800156116485596

 $00:24:51.810 \longrightarrow 00:24:54.015$ We also have this or lot inhibin

NOTE Confidence: 0.800156116485596

 $00:24:54.015 \longrightarrow 00:24:54.960$ Rammus serum app,

NOTE Confidence: 0.800156116485596

 $00:24:54.960 \longrightarrow 00:24:57.725$ so I showed at the very beginning.

NOTE Confidence: 0.800156116485596

 $00:24:57.730 \longrightarrow 00:24:59.470$ The different ways to target

NOTE Confidence: 0.800156116485596

 $00{:}24{:}59.470 \dashrightarrow 00{:}25{:}00.514$ these tyrosine kinase.

NOTE Confidence: 0.800156116485596

 $00:25:00.520 \longrightarrow 00:25:02.710$ So the first where the tyrosine

NOTE Confidence: 0.800156116485596

 $00{:}25{:}02.710 \dashrightarrow 00{:}25{:}04.879$ kinase inhibitors which work in the

NOTE Confidence: 0.800156116485596

 $00:25:04.879 \longrightarrow 00:25:06.799$ cell at the tyrosine kinase domain.

NOTE Confidence: 0.800156116485596

 $00:25:06.800 \longrightarrow 00:25:09.243$ But then I also showed two different

NOTE Confidence: 0.800156116485596

 $00:25:09.243 \longrightarrow 00:25:10.290$ strategies for antibodies,

 $00:25:10.290 \longrightarrow 00:25:12.414$ one the bevacizumab type or its

NOTE Confidence: 0.800156116485596

 $00:25:12.414 \longrightarrow 00:25:14.858$ binding to the leg and so sucking

NOTE Confidence: 0.800156116485596

 $00{:}25{:}14.858 \dashrightarrow 00{:}25{:}17.527$ all the leg and away the other binds

NOTE Confidence: 0.800156116485596

 $00:25:17.527 \longrightarrow 00:25:20.068$ to the receptor and sort of blocks.

NOTE Confidence: 0.800156116485596

 $00:25:20.070 \longrightarrow 00:25:22.164$ The way the receptor works oramus

NOTE Confidence: 0.800156116485596

00:25:22.164 --> 00:25:24.598 serum AB is blocking veg F receptor.

NOTE Confidence: 0.800156116485596

00:25:24.600 --> 00:25:26.924 So you've got our Latin if the

NOTE Confidence: 0.800156116485596

 $00{:}25{:}26.924 \rightarrow 00{:}25{:}28.520$ tyrosine kinase inhibitor blocking.

NOTE Confidence: 0.800156116485596

 $00:25:28.520 \longrightarrow 00:25:29.796$ The tyrosine kinase activity

NOTE Confidence: 0.800156116485596

00:25:29.796 --> 00:25:31.710 and then veg F Knotty Jeff

NOTE Confidence: 0.832051396369934

 $00{:}25{:}31.768 {\:\dashrightarrow\:} 00{:}25{:}34.029$ but Jeff related is also being blocked,

NOTE Confidence: 0.832051396369934

 $00:25:34.030 \longrightarrow 00:25:36.530$ so you get a blockage that way and you see

NOTE Confidence: 0.832051396369934

 $00{:}25{:}36.590 \dashrightarrow 00{:}25{:}39.206$ this nice progression free survival benefit.

NOTE Confidence: 0.832051396369934

 $00:25:39.210 \longrightarrow 00:25:40.830$ But we haven't seen the

NOTE Confidence: 0.832051396369934

 $00:25:40.830 \longrightarrow 00:25:42.126$ overall survival data yet.

 $00:25:42.130 \longrightarrow 00:25:43.680$ This combination is approved as

NOTE Confidence: 0.832051396369934

 $00{:}25{:}43.680 \dashrightarrow 00{:}25{:}45.690$ an option in the United States,

NOTE Confidence: 0.832051396369934

 $00:25:45.690 \longrightarrow 00:25:48.105$ but I think that we're waiting to

NOTE Confidence: 0.832051396369934

 $00:25:48.105 \longrightarrow 00:25:50.025$ see what that overall survival

NOTE Confidence: 0.832051396369934

 $00:25:50.025 \longrightarrow 00:25:52.449$ data is going to look like.

NOTE Confidence: 0.832051396369934

00:25:52.450 --> 00:25:54.767 Now I've talked about her Latin appear,

NOTE Confidence: 0.832051396369934

 $00:25:54.770 \longrightarrow 00:25:56.420$ but you're thinking the beginning.

NOTE Confidence: 0.832051396369934

 $00:25:56.420 \longrightarrow 00:25:58.406$ You were telling me that Osimertinib

NOTE Confidence: 0.832051396369934

 $00:25:58.406 \longrightarrow 00:25:59.730$ is preferred over these.

NOTE Confidence: 0.832051396369934 00:25:59.730 --> 00:26:00.428 So what? NOTE Confidence: 0.832051396369934

 $00:26:00.428 \longrightarrow 00:26:02.522$ Where is the oh summer nimbuses

NOTE Confidence: 0.832051396369934

 $00:26:02.522 \longrightarrow 00:26:04.700$ amount data will hear some of it.

NOTE Confidence: 0.832051396369934

 $00:26:04.700 \longrightarrow 00:26:07.012$ So this was data that Hanyu and the

NOTE Confidence: 0.832051396369934

00:26:07.012 --> 00:26:09.328 Memorial Sloan Kettering Group put together.

NOTE Confidence: 0.832051396369934

 $00:26:09.330 \longrightarrow 00:26:10.582$ This is not randomized.

NOTE Confidence: 0.832051396369934

 $00:26:10.582 \longrightarrow 00:26:12.147$ This is just patients getting

00:26:12.147 --> 00:26:13.969 oh smart nib and bevacizumab,

NOTE Confidence: 0.832051396369934

 $00:26:13.970 \longrightarrow 00:26:15.750$ and the toxicity was manageable

NOTE Confidence: 0.832051396369934

 $00:26:15.750 \longrightarrow 00:26:17.849$ and the response rate was very

NOTE Confidence: 0.832051396369934

 $00:26:17.849 \longrightarrow 00:26:20.260$ high at 80% and the median time to

NOTE Confidence: 0.832051396369934

 $00:26:20.260 \longrightarrow 00:26:21.920$ win the drugs combination stopped

NOTE Confidence: 0.832051396369934

 $00:26:21.980 \longrightarrow 00:26:23.640$ working was about 19 months.

NOTE Confidence: 0.832051396369934

 $00:26:23.640 \longrightarrow 00:26:24.768$ So that's interesting.

NOTE Confidence: 0.832051396369934

00:26:24.768 --> 00:26:25.896 It's not overwhelming,

NOTE Confidence: 0.832051396369934

 $00:26:25.900 \longrightarrow 00:26:26.746$ but interesting.

NOTE Confidence: 0.832051396369934

00:26:26.746 --> 00:26:29.284 However, there is this very confusing

NOTE Confidence: 0.832051396369934

00:26:29.284 --> 00:26:31.539 presentation that we saw just last month,

NOTE Confidence: 0.832051396369934

 $00:26:31.540 \longrightarrow 00:26:33.802$ two months ago now in September

NOTE Confidence: 0.832051396369934

 $00:26:33.802 \longrightarrow 00:26:36.049$ at the ESMO meeting the 2020,

NOTE Confidence: 0.832051396369934

 $00:26:36.050 \longrightarrow 00:26:37.930$ which was a randomized phase,

NOTE Confidence: 0.832051396369934

00:26:37.930 --> 00:26:40.938 two of oh Smyrna plus or minus bevacizumab,

 $00:26:40.940 \longrightarrow 00:26:42.444$ again done in Japan.

NOTE Confidence: 0.832051396369934

00:26:42.444 --> 00:26:44.700 And when you look at this,

NOTE Confidence: 0.832051396369934

 $00:26:44.700 \longrightarrow 00:26:46.950$ the blue line is actually the

NOTE Confidence: 0.832051396369934

 $00:26:46.950 \longrightarrow 00:26:47.700$ osimertinib alone.

NOTE Confidence: 0.832051396369934

 $00:26:47.700 \longrightarrow 00:26:49.580$ So in this particular trial,

NOTE Confidence: 0.832051396369934

 $00:26:49.580 \longrightarrow 00:26:51.260$ the combination of oh Summerton

NOTE Confidence: 0.832051396369934

 $00{:}26{:}51.260 \dashrightarrow 00{:}26{:}53.462$ Bevis is a map looked worse

NOTE Confidence: 0.832051396369934

 $00:26:53.462 \longrightarrow 00:26:55.286$ for progression free survival.

NOTE Confidence: 0.832051396369934

 $00:26:55.290 \longrightarrow 00:26:57.000$ Then just the ocean alone.

NOTE Confidence: 0.832051396369934

 $00:26:57.000 \longrightarrow 00:26:59.568$ So that's a little bit confusing.

NOTE Confidence: 0.832051396369934

 $00{:}26{:}59.570 \dashrightarrow 00{:}27{:}01.290$ The ultimate number alone progression

NOTE Confidence: 0.832051396369934

 $00:27:01.290 \longrightarrow 00:27:03.590$ free survival was lower than were well.

NOTE Confidence: 0.832051396369934

00:27:03.590 --> 00:27:06.270 I should say this was done second line,

NOTE Confidence: 0.832051396369934

 $00:27:06.270 \longrightarrow 00:27:08.142$ so this was for patients who

NOTE Confidence: 0.832051396369934

00:27:08.142 --> 00:27:10.628 had or lot nip or just fitting.

NOTE Confidence: 0.832051396369934

00:27:10.630 --> 00:27:12.305 It already had developed T790M

00:27:12.305 --> 00:27:13.980 and then went on Osimertinib.

NOTE Confidence: 0.832051396369934

 $00:27:13.980 \longrightarrow 00:27:16.660$ So it's different than the first line data.

NOTE Confidence: 0.832051396369934

00:27:16.660 --> 00:27:19.549 But in this trial with no smart net plus

NOTE Confidence: 0.832051396369934

 $00:27:19.549 \longrightarrow 00:27:22.706$ or minus bevacizumab in the second line.

NOTE Confidence: 0.832051396369934

 $00:27:22.710 \longrightarrow 00:27:24.365$ The awesome art never alone

NOTE Confidence: 0.832051396369934

 $00:27:24.365 \longrightarrow 00:27:25.358$ outperformed the combination.

NOTE Confidence: 0.832051396369934

00:27:25.360 --> 00:27:28.008 Now this is a randomized phase two trial,

NOTE Confidence: 0.832051396369934

 $00:27:28.010 \longrightarrow 00:27:29.990$ so we have to be careful,

NOTE Confidence: 0.832051396369934

 $00:27:29.990 \longrightarrow 00:27:31.314$ sometimes randomized phase two

NOTE Confidence: 0.832051396369934

 $00:27:31.314 \longrightarrow 00:27:33.300$ studies don't give us the truth.

NOTE Confidence: 0.832051396369934

 $00:27:33.300 \longrightarrow 00:27:35.298$ This study did show that the

NOTE Confidence: 0.832051396369934

 $00{:}27{:}35.298 \dashrightarrow 00{:}27{:}37.011$ response to the combination was

NOTE Confidence: 0.832051396369934

00:27:37.011 --> 00:27:38.596 higher than the single drug,

NOTE Confidence: 0.832051396369934

 $00:27:38.600 \longrightarrow 00:27:40.255$ but just not that progression

NOTE Confidence: 0.832051396369934

 $00:27:40.255 \longrightarrow 00:27:41.248$ free survival benefit.

 $00:27:41.250 \longrightarrow 00:27:43.890$ So we're kind of confused by this result.

NOTE Confidence: 0.832051396369934

 $00:27:43.890 \longrightarrow 00:27:46.123$ We are moving forward with the next

NOTE Confidence: 0.832051396369934

00:27:46.123 --> 00:27:47.870 line of exploring this though,

NOTE Confidence: 0.832051396369934

 $00:27:47.870 \longrightarrow 00:27:49.904$ so this is a randomized phase

NOTE Confidence: 0.832051396369934

 $00:27:49.904 \longrightarrow 00:27:51.636$ study randomised three study in

NOTE Confidence: 0.832051396369934

 $00:27:51.636 \longrightarrow 00:27:52.896$ the Kaga Chrome network.

NOTE Confidence: 0.832051396369934

 $00:27:52.900 \longrightarrow 00:27:55.175$ Led by Helena, you and bless Hamas.

NOTE Confidence: 0.832051396369934

 $00:27:55.180 \longrightarrow 00:27:57.130$ And this is for patients who

NOTE Confidence: 0.832051396369934

 $00:27:57.130 \longrightarrow 00:27:58.430$ have just been diagnosed.

NOTE Confidence: 0.832051396369934

 $00:27:58.430 \longrightarrow 00:28:01.030$ So the one I just showed you with

NOTE Confidence: 0.832051396369934

 $00:28:01.030 \longrightarrow 00:28:03.104$ second line. This is first line.

NOTE Confidence: 0.832051396369934

00:28:03.104 --> 00:28:04.884 Patients just diagnosed with lung

NOTE Confidence: 0.832051396369934

 $00{:}28{:}04.884 \dashrightarrow 00{:}28{:}07.089$ cancer that has an EGFR mutation and

NOTE Confidence: 0.832051396369934

 $00:28:07.089 \longrightarrow 00:28:09.266$ they're going to get oh smart nib

NOTE Confidence: 0.832051396369934

00:28:09.266 --> 00:28:11.429 plus or minus bevacizumab in the study.

NOTE Confidence: 0.832051396369934

 $00:28:11.430 \longrightarrow 00:28:12.730$ Just opened to enrollment,

 $00:28:12.730 \longrightarrow 00:28:14.680$ so we have no data yet,

NOTE Confidence: 0.832051396369934

00:28:14.680 --> 00:28:16.300 but we're excited to participate,

NOTE Confidence: 0.832051396369934

00:28:16.300 --> 00:28:16.950 though again,

NOTE Confidence: 0.832051396369934

 $00:28:16.950 \longrightarrow 00:28:19.550$ as we talk about this option with patients,

NOTE Confidence: 0.832051396369934

00:28:19.550 --> 00:28:21.362 it will be important to talk

NOTE Confidence: 0.832051396369934

 $00:28:21.362 \longrightarrow 00:28:22.570$ about the data that

NOTE Confidence: 0.822217047214508

 $00:28:22.641 \longrightarrow 00:28:23.838$ we've already seen.

NOTE Confidence: 0.822217047214508

 $00:28:23.840 \longrightarrow 00:28:25.886$ With the other combinations of EGFR,

NOTE Confidence: 0.822217047214508

00:28:25.890 --> 00:28:28.626 TKI plus or minus a veg F drug,

NOTE Confidence: 0.822217047214508

 $00:28:28.630 \longrightarrow 00:28:30.340$ so we're still kind of

NOTE Confidence: 0.822217047214508

 $00:28:30.340 \longrightarrow 00:28:31.708$ trying to understand this,

NOTE Confidence: 0.822217047214508

 $00:28:31.710 \longrightarrow 00:28:34.538$ and there's also a no smart nib

NOTE Confidence: 0.822217047214508

 $00{:}28{:}34.538 \to 00{:}28{:}36.440$ romanisti remap trial ongoing.

NOTE Confidence: 0.822217047214508

 $00:28:36.440 \longrightarrow 00:28:40.016$ OK, so leaving that and now on too,

NOTE Confidence: 0.822217047214508

00:28:40.020 --> 00:28:43.884 so hopefully you're following EGFR TK alone.

 $00:28:43.890 \longrightarrow 00:28:46.200$ EGFR TK I plus or minus.

NOTE Confidence: 0.822217047214508

 $00{:}28{:}46.200 \dashrightarrow 00{:}28{:}47.740$ These anti angiogenesis drugs.

NOTE Confidence: 0.822217047214508

00:28:47.740 --> 00:28:50.050 And now we're looking at Egypt,

NOTE Confidence: 0.822217047214508

00:28:50.050 --> 00:28:52.360 far EGFR combinations and this is

NOTE Confidence: 0.822217047214508

 $00:28:52.360 \longrightarrow 00:28:54.574$ where we're looking at, an EGFR,

NOTE Confidence: 0.822217047214508

00:28:54.574 --> 00:28:56.933 TKI pill, plus an antibody drug that's

NOTE Confidence: 0.822217047214508

 $00:28:56.933 \longrightarrow 00:28:59.290$ also hitting that EGFR approaching.

NOTE Confidence: 0.822217047214508

00:28:59.290 --> 00:29:01.978 And if you think again about that first

NOTE Confidence: 0.822217047214508

00:29:01.978 --> 00:29:05.000 slide I showed you or early on where

NOTE Confidence: 0.822217047214508

 $00:29:05.000 \longrightarrow 00:29:07.380$ you've got the different strategies.

NOTE Confidence: 0.822217047214508

00:29:07.380 --> 00:29:09.690 If you've got a receptor tyrosine

NOTE Confidence: 0.822217047214508

00:29:09.690 --> 00:29:11.230 kinase inhibitor tyrosine kinase,

NOTE Confidence: 0.822217047214508

 $00:29:11.230 \longrightarrow 00:29:14.374$ if you block at the tyrosine kinase domain.

NOTE Confidence: 0.822217047214508

 $00:29:14.380 \longrightarrow 00:29:16.596$ But you also block at that receptor point.

NOTE Confidence: 0.822217047214508

00:29:16.600 --> 00:29:18.808 You're hitting the same protein in two ways,

NOTE Confidence: 0.822217047214508

 $00:29:18.810 \longrightarrow 00:29:20.545$ and so that's the theory

 $00:29:20.545 \longrightarrow 00:29:22.280$ behind why this would work.

NOTE Confidence: 0.822217047214508

 $00:29:22.280 \longrightarrow 00:29:23.534$ And long ago.

NOTE Confidence: 0.822217047214508

 $00:29:23.534 \longrightarrow 00:29:25.624$ Before we had to simmer

NOTE Confidence: 0.822217047214508

 $00:29:25.624 \longrightarrow 00:29:28.359$ nib so going back in time.

NOTE Confidence: 0.822217047214508

 $00:29:28.360 \longrightarrow 00:29:31.726$ We had a combination of an EGFR pill drug

NOTE Confidence: 0.822217047214508

00:29:31.726 --> 00:29:34.198 afatinib plus in Egypt are antibody,

NOTE Confidence: 0.822217047214508 00:29:34.200 --> 00:29:35.092 so tux, NOTE Confidence: 0.822217047214508

 $00:29:35.092 \longrightarrow 00:29:38.214$ omab and this combination was the first

NOTE Confidence: 0.822217047214508

00:29:38.214 --> 00:29:40.472 combination and the first thing we

NOTE Confidence: 0.822217047214508

 $00{:}29{:}40.472 \dashrightarrow 00{:}29{:}43.536$ ever saw that worked in the setting of

NOTE Confidence: 0.822217047214508

 $00:29:43.536 \longrightarrow 00:29:46.441$ having been on a first generation drugs.

NOTE Confidence: 0.822217047214508

00:29:46.450 --> 00:29:47.294 Developing T790M,

NOTE Confidence: 0.822217047214508

 $00:29:47.294 \longrightarrow 00:29:49.826$ so before we add anything else,

NOTE Confidence: 0.822217047214508

 $00:29:49.830 \longrightarrow 00:29:52.225$ this combination worked and so

NOTE Confidence: 0.822217047214508

 $00:29:52.225 \longrightarrow 00:29:54.620$ this was the original publication.

00:29:54.620 --> 00:29:55.168 And again,

NOTE Confidence: 0.822217047214508

00:29:55.168 --> 00:29:56.538 this is a waterfall plot,

NOTE Confidence: 0.822217047214508

 $00:29:56.540 \longrightarrow 00:29:58.458$ so you see a few patients had

NOTE Confidence: 0.822217047214508

 $00:29:58.458 \longrightarrow 00:29:59.280$ there's tumors grow,

NOTE Confidence: 0.822217047214508

 $00:29:59.280 \longrightarrow 00:30:00.918$ but a lot of patients had

NOTE Confidence: 0.822217047214508

00:30:00.918 --> 00:30:01.737 their tumors shrink,

NOTE Confidence: 0.822217047214508

 $00:30:01.740 \longrightarrow 00:30:03.252$ and so this was an option

NOTE Confidence: 0.822217047214508

 $00:30:03.252 \longrightarrow 00:30:04.760$ that we had for awhile.

NOTE Confidence: 0.822217047214508

 $00:30:04.760 \dashrightarrow 00:30:07.500$ But now we don't use it too much because of,

NOTE Confidence: 0.822217047214508

 $00{:}30{:}08.184 \dashrightarrow 00{:}30{:}10.578$ but we're coming back to study it.

NOTE Confidence: 0.822217047214508

 $00:30:10.580 \longrightarrow 00:30:12.480$ And so there are combinations.

NOTE Confidence: 0.822217047214508

00:30:12.480 --> 00:30:13.845 Nauvo summer nib.

NOTE Confidence: 0.822217047214508

 $00:30:13.845 \longrightarrow 00:30:15.665$ Plus and different Egypt

NOTE Confidence: 0.822217047214508

00:30:15.665 --> 00:30:17.030 for antibody necitum umab.

NOTE Confidence: 0.822217047214508

 $00:30:17.030 \longrightarrow 00:30:19.774$ So we are part of this California

 $00:30:19.774 \longrightarrow 00:30:20.950$ Cancer Consortium study.

NOTE Confidence: 0.822217047214508

00:30:20.950 --> 00:30:22.830 Jonathan Rhys has talked about

NOTE Confidence: 0.822217047214508

 $00:30:22.830 \longrightarrow 00:30:25.185$ some of the preliminary data at

NOTE Confidence: 0.822217047214508

00:30:25.185 --> 00:30:27.597 ASCO last year and more ongoing

NOTE Confidence: 0.822217047214508

 $00:30:27.597 \longrightarrow 00:30:29.190$ show encouraging information here,

NOTE Confidence: 0.822217047214508

 $00:30:29.190 \longrightarrow 00:30:31.899$ and this is being looked at for

NOTE Confidence: 0.822217047214508

00:30:31.899 --> 00:30:34.754 patients who have already been on oh

NOTE Confidence: 0.822217047214508

 $00:30:34.754 \dashrightarrow 00:30:38.200$ smart nib and it stopped working or early on.

NOTE Confidence: 0.822217047214508

 $00:30:38.200 \dashrightarrow 00:30:40.250$ There also combinations with a fatinib

NOTE Confidence: 0.822217047214508

 $00{:}30{:}40.250 \dashrightarrow 00{:}30{:}42.708$ and necitum umab and were part of

NOTE Confidence: 0.822217047214508

 $00:30:42.708 \longrightarrow 00:30:44.446$ that consortium which is, uh.

NOTE Confidence: 0.822217047214508

 $00:30:44.446 \longrightarrow 00:30:44.812$ Vanderbilt,

NOTE Confidence: 0.822217047214508

 $00{:}30{:}44.812 \dashrightarrow 00{:}30{:}48.190$ so a lot of work still looking at this,

NOTE Confidence: 0.822217047214508

00:30:48.190 --> 00:30:50.344 but some toxicity issues so we'll

NOTE Confidence: 0.822217047214508

 $00:30:50.344 \longrightarrow 00:30:52.559$ see how that pans out there.

00:30:52.560 --> 00:30:52.924 Also,

NOTE Confidence: 0.822217047214508

 $00{:}30{:}52.924 \dashrightarrow 00{:}30{:}54.744$ newer drugs being developed at

NOTE Confidence: 0.822217047214508

 $00:30:54.744 \longrightarrow 00:30:57.157$ build on this so one is this

NOTE Confidence: 0.822217047214508

00:30:57.157 --> 00:30:58.927 aim of antonyms drug which is

NOTE Confidence: 0.822217047214508

 $00:30:58.927 \longrightarrow 00:31:01.289$ a combination of EGFR antibody.

NOTE Confidence: 0.822217047214508

00:31:01.290 --> 00:31:02.602 It's also hitting met,

NOTE Confidence: 0.822217047214508

 $00:31:02.602 \longrightarrow 00:31:05.300$ so if you were paying close attention,

NOTE Confidence: 0.822217047214508

 $00:31:05.300 \longrightarrow 00:31:07.484$ you'll notice I mentioned met as

NOTE Confidence: 0.822217047214508

 $00:31:07.484 \longrightarrow 00:31:08.940$ a potential resistance mechanism,

NOTE Confidence: 0.822217047214508

 $00:31:08.940 \longrightarrow 00:31:11.118$ and so there's data to support

NOTE Confidence: 0.822217047214508

 $00{:}31{:}11.118 \dashrightarrow 00{:}31{:}12.207$ looking at met.

NOTE Confidence: 0.822217047214508

 $00:31:12.210 \longrightarrow 00:31:14.548$ There have been a lot of trials

NOTE Confidence: 0.822217047214508

 $00:31:14.548 \longrightarrow 00:31:16.599$ looking at specific met inhibitors.

NOTE Confidence: 0.822217047214508

00:31:16.600 --> 00:31:18.526 In the Egypt Far world again,

NOTE Confidence: 0.822217047214508

00:31:18.530 --> 00:31:20.140 I only have an hour,

NOTE Confidence: 0.822217047214508

 $00:31:20.140 \longrightarrow 00:31:22.716$ so I couldn't go into all of that.

00:31:22.720 --> 00:31:24.967 They haven't really gone too far yet.

NOTE Confidence: 0.822217047214508

00:31:24.970 --> 00:31:26.580 We haven't changed our treatment,

NOTE Confidence: 0.822217047214508

 $00{:}31{:}26.580 \dashrightarrow 00{:}31{:}28.506$ but we haven't forgotten about that.

NOTE Confidence: 0.817905068397522

 $00:31:28.510 \longrightarrow 00:31:30.120$ So we're looking at Egypt.

NOTE Confidence: 0.817905068397522

 $00:31:30.120 \longrightarrow 00:31:31.101$ Are met antibody,

NOTE Confidence: 0.817905068397522

 $00:31:31.101 \longrightarrow 00:31:33.063$ hear them if antonym in combination

NOTE Confidence: 0.817905068397522

 $00:31:33.063 \longrightarrow 00:31:34.754$ with the different third generation

NOTE Confidence: 0.817905068397522

 $00:31:34.754 \longrightarrow 00:31:37.068$ drug lizard nip and to this was

NOTE Confidence: 0.817905068397522

 $00:31:37.068 \longrightarrow 00:31:39.140$ dated that was just presented at Asmo,

NOTE Confidence: 0.817905068397522

 $00:31:39.140 \longrightarrow 00:31:40.133$ which was encouraging,

NOTE Confidence: 0.817905068397522

 $00:31:40.133 \longrightarrow 00:31:42.450$ showing that for patients who had already

NOTE Confidence: 0.817905068397522

 $00:31:42.510 \longrightarrow 00:31:43.985$ had osimertinib in this preliminary

NOTE Confidence: 0.817905068397522

 $00{:}31{:}43.985 \dashrightarrow 00{:}31{:}46.176$ data about a third of those patients

NOTE Confidence: 0.817905068397522

 $00{:}31{:}46.176 \dashrightarrow 00{:}31{:}48.156$ did respond to this new combination.

NOTE Confidence: 0.817905068397522

 $00:31:48.160 \longrightarrow 00:31:50.505$ So we'll see how that pans out.

00:31:50.510 --> 00:31:51.858 How long it works,

NOTE Confidence: 0.817905068397522

 $00:31:51.858 \longrightarrow 00:31:53.880$ what the toxicity is show there's

NOTE Confidence: 0.817905068397522

 $00:31:53.950 \longrightarrow 00:31:56.180$ there is movement in excitement

NOTE Confidence: 0.817905068397522

 $00:31:56.180 \longrightarrow 00:31:57.518$ about new combinations.

NOTE Confidence: 0.817905068397522

 $00:31:57.520 \longrightarrow 00:31:59.585$ There are also newer EGFR

NOTE Confidence: 0.817905068397522

 $00:31:59.585 \longrightarrow 00:32:02.520$ pathway or this is the new drug.

NOTE Confidence: 0.817905068397522

 $00:32:02.520 \longrightarrow 00:32:04.866$ Use U Three 1402 or projection

NOTE Confidence: 0.817905068397522

 $00:32:04.866 \longrightarrow 00:32:07.312$ map directed hand and this drug

NOTE Confidence: 0.817905068397522

 $00:32:07.312 \longrightarrow 00:32:09.706$ is looking at her three and

NOTE Confidence: 0.817905068397522

00:32:09.706 --> 00:32:11.939 you're confused now because you

NOTE Confidence: 0.817905068397522

 $00:32:11.939 \dashrightarrow 00:32:14.199$ think you're talking about EGFR.

NOTE Confidence: 0.817905068397522

00:32:14.200 --> 00:32:16.696 Why, you talking bout her three?

NOTE Confidence: 0.817905068397522

 $00:32:16.700 \longrightarrow 00:32:19.619$ Well, EGFR is also called her one,

NOTE Confidence: 0.817905068397522

 $00:32:19.620 \longrightarrow 00:32:22.588$ so her one an heard too and her

NOTE Confidence: 0.817905068397522

 $00:32:22.588 \longrightarrow 00:32:26.025$ three are all in the same family and

NOTE Confidence: 0.817905068397522

 $00:32:26.025 \longrightarrow 00:32:28.849$ they do interact with each other.

 $00:32:28.850 \longrightarrow 00:32:31.040$ Show this antibody drug conjugate

NOTE Confidence: 0.817905068397522

 $00{:}32{:}31.040 \dashrightarrow 00{:}32{:}33.676$ that's focused on her three there's

NOTE Confidence: 0.817905068397522

 $00{:}32{:}33.676 \dashrightarrow 00{:}32{:}36.287$ hope and some data now showing that

NOTE Confidence: 0.817905068397522

 $00:32:36.287 \longrightarrow 00:32:39.085$ it could be active in the setting

NOTE Confidence: 0.817905068397522

 $00:32:39.085 \longrightarrow 00:32:41.060$ where there's Egypt far drive.

NOTE Confidence: 0.817905068397522

00:32:41.060 --> 00:32:43.909 Sorry, each of our driven lung cancer,

NOTE Confidence: 0.817905068397522

 $00:32:43.910 \longrightarrow 00:32:46.458$ so this is again some very preliminary

NOTE Confidence: 0.817905068397522

 $00:32:46.458 \dashrightarrow 00:32:49.128$ data where they were looking at patients

NOTE Confidence: 0.817905068397522

 $00{:}32{:}49.128 \dashrightarrow 00{:}32{:}51.930$ who had EGFR mutant lung cancer had

NOTE Confidence: 0.817905068397522

 $00:32:51.930 \longrightarrow 00:32:53.790$ previously received osimertinib or

NOTE Confidence: 0.817905068397522

 $00:32:53.790 \longrightarrow 00:32:57.341$ had developed resistance to or lot in a

NOTE Confidence: 0.817905068397522

 $00:32:57.341 \longrightarrow 00:33:00.190$ bridge of fitness and didn't have T790M.

NOTE Confidence: 0.817905068397522

 $00{:}33{:}00.190 \dashrightarrow 00{:}33{:}03.853$ And again on this waterfall plot you can see.

NOTE Confidence: 0.817905068397522

 $00:33:03.860 \longrightarrow 00:33:05.918$ That there have definitely been some

NOTE Confidence: 0.817905068397522

 $00:33:05.918 \longrightarrow 00:33:08.180$ responses and the details here all about.

00:33:08.180 --> 00:33:10.166 Well, what was the resistance mechanism?

NOTE Confidence: 0.817905068397522

 $00:33:10.170 \longrightarrow 00:33:12.487$ What was the underlying you do fermentation?

NOTE Confidence: 0.817905068397522

 $00:33:12.490 \longrightarrow 00:33:13.714$ What resistance was seen?

NOTE Confidence: 0.817905068397522

 $00:33:13.714 \longrightarrow 00:33:16.808$ And so there's a lot to parse through here,

NOTE Confidence: 0.817905068397522

 $00:33:16.810 \longrightarrow 00:33:18.470$ which I won't go into.

NOTE Confidence: 0.817905068397522

 $00:33:18.470 \longrightarrow 00:33:20.850$ But encouraging new data.

NOTE Confidence: 0.817905068397522

00:33:20.850 --> 00:33:21.196 OK.

NOTE Confidence: 0.817905068397522

00:33:21.196 --> 00:33:24.310 So now we talked about Egypt Party K eyes

NOTE Confidence: 0.817905068397522

 $00{:}33{:}24.393 \dashrightarrow 00{:}33{:}27.315$ alone veg F novelly Jeff combinations.

NOTE Confidence: 0.817905068397522

00:33:27.320 --> 00:33:29.784 And now I'm going to switch to chemo

NOTE Confidence: 0.817905068397522

 $00{:}33{:}29.784 \dashrightarrow 00{:}33{:}31.952$ and talk about Egypt, Kaizen, chemo.

NOTE Confidence: 0.817905068397522

 $00:33:31.952 \longrightarrow 00:33:33.938$ So again, bear with me here.

NOTE Confidence: 0.817905068397522

00:33:33.940 --> 00:33:36.910 This is a it's a lot to go through,

NOTE Confidence: 0.817905068397522

 $00:33:36.910 \longrightarrow 00:33:39.101$ but there's been so much we've been

NOTE Confidence: 0.817905068397522

00:33:39.101 --> 00:33:40.889 exploring and trying to understand.

NOTE Confidence: 0.817905068397522

 $00:33:40.890 \longrightarrow 00:33:43.590$ How do we do a better job of treating

00:33:43.590 --> 00:33:45.847 patients with EGFR mutant lung cancer?

NOTE Confidence: 0.817905068397522

 $00:33:45.850 \longrightarrow 00:33:47.510$ So what about combining chemotherapy?

NOTE Confidence: 0.817905068397522

 $00:33:47.510 \longrightarrow 00:33:49.160$ Need you 40K as well?

NOTE Confidence: 0.817905068397522

00:33:49.160 --> 00:33:51.808 When we first learned about the EGFR Tkis,

NOTE Confidence: 0.817905068397522

 $00:33:51.810 \longrightarrow 00:33:53.062$ there were big trials.

NOTE Confidence: 0.817905068397522

 $00:33:53.062 \longrightarrow 00:33:54.940$ Combining them with chemo misses the

NOTE Confidence: 0.817905068397522

00:33:54.996 --> 00:33:56.958 tribute and talent studies and again,

NOTE Confidence: 0.817905068397522

 $00:33:56.960 \longrightarrow 00:33:59.496$ Doctor Herbs played a big role in these.

NOTE Confidence: 0.817905068397522

 $00:33:59.500 \longrightarrow 00:34:01.250$ And these studies were completely,

NOTE Confidence: 0.817905068397522

00:34:01.250 --> 00:34:01.938 absolutely negative,

NOTE Confidence: 0.817905068397522

 $00:34:01.938 \longrightarrow 00:34:04.346$ and this was done before we knew

NOTE Confidence: 0.817905068397522

 $00{:}34{:}04.346 \dashrightarrow 00{:}34{:}06.150$ about EGFR mutant lung cancer.

NOTE Confidence: 0.817905068397522

 $00{:}34{:}06.150 \dashrightarrow 00{:}34{:}08.250$ We were treating every body the same,

NOTE Confidence: 0.817905068397522

 $00:34:08.250 \longrightarrow 00:34:09.534$ but totally negative studies.

NOTE Confidence: 0.817905068397522

 $00:34:09.534 \longrightarrow 00:34:12.282$ And so this let us to not really

 $00:34:12.282 \longrightarrow 00:34:14.550$ think about this approach for awhile.

NOTE Confidence: 0.817905068397522

 $00:34:14.550 \longrightarrow 00:34:16.300$ There was some work looking

NOTE Confidence: 0.817905068397522

 $00:34:16.300 \longrightarrow 00:34:17.700$ at this intercalation idea,

NOTE Confidence: 0.817905068397522

 $00:34:17.700 \longrightarrow 00:34:20.150$ where you would use the Jaffar TK.

NOTE Confidence: 0.817905068397522

00:34:20.150 --> 00:34:22.950 I stop it, get the cell signaling again,

NOTE Confidence: 0.817905068397522

 $00:34:22.950 \longrightarrow 00:34:23.958$ give the chemo,

NOTE Confidence: 0.817905068397522

 $00:34:23.958 \longrightarrow 00:34:25.974$ restart the EGFR TKI and this

NOTE Confidence: 0.817905068397522

00:34:25.974 --> 00:34:27.500 actually was encourage ING.

NOTE Confidence: 0.817905068397522

 $00:34:27.500 \longrightarrow 00:34:29.648$ It showed an overall survival benefit.

NOTE Confidence: 0.817905068397522

00:34:29.650 --> 00:34:31.185 But people don't really want

NOTE Confidence: 0.817905068397522

 $00:34:31.185 \longrightarrow 00:34:32.720$ to be on chemo if

NOTE Confidence: 0.870585560798645

 $00:34:32.795 \longrightarrow 00:34:34.804$ they can just take a pill and

NOTE Confidence: 0.870585560798645

 $00:34:34.804 \longrightarrow 00:34:37.008$ said this was not widely adopted.

NOTE Confidence: 0.870585560798645

 $00:34:37.010 \longrightarrow 00:34:38.954$ People looked at whether when you

NOTE Confidence: 0.870585560798645

 $00:34:38.954 \longrightarrow 00:34:41.064$ were on a first generation EGFR

NOTE Confidence: 0.870585560798645

00:34:41.064 --> 00:34:43.290 drug and it stopped working and

 $00:34:43.290 \longrightarrow 00:34:45.489$ you were going to go to chemo.

NOTE Confidence: 0.870585560798645

 $00:34:45.490 \longrightarrow 00:34:47.180$ This was before Osama Rname.

NOTE Confidence: 0.870585560798645

00:34:47.180 --> 00:34:49.546 Whether you could continue the EGFR drug,

NOTE Confidence: 0.870585560798645

 $00:34:49.550 \longrightarrow 00:34:51.590$ or if you should stop it.

NOTE Confidence: 0.870585560798645

 $00:34:51.590 \longrightarrow 00:34:53.534$ And this was the impressed trial

NOTE Confidence: 0.870585560798645

 $00:34:53.534 \longrightarrow 00:34:55.320$ and that didn't work either.

NOTE Confidence: 0.870585560798645

 $00:34:55.320 \longrightarrow 00:34:57.686$ There was no benefit an if anything.

NOTE Confidence: 0.870585560798645

 $00:34:57.690 \longrightarrow 00:34:59.370$ Survival favored stopping that Jeff

NOTE Confidence: 0.870585560798645

 $00{:}34{:}59.370 \dashrightarrow 00{:}35{:}01.759$ it nib and just going with chemo.

NOTE Confidence: 0.870585560798645

 $00:35:01.760 \longrightarrow 00:35:03.344$ So we had these.

NOTE Confidence: 0.870585560798645

 $00:35:03.344 \dashrightarrow 00:35:05.324$ Big randomized trials combinations negative.

NOTE Confidence: 0.870585560798645

 $00:35:05.330 \longrightarrow 00:35:06.990$ We had this trial negative.

NOTE Confidence: 0.870585560798645

 $00{:}35{:}06.990 \dashrightarrow 00{:}35{:}08.985$ If you combine the EGFR TK I

NOTE Confidence: 0.870585560798645

 $00:35:08.985 \longrightarrow 00:35:11.167$ in the chemo together so not a

NOTE Confidence: 0.870585560798645

 $00:35:11.167 \longrightarrow 00:35:13.502$ lot of enthusiasm but a lot of

00:35:13.502 --> 00:35:15.417 questions still because the idea

NOTE Confidence: 0.870585560798645

00:35:15.417 --> 00:35:18.000 of stopping in EGFR TKI for tumor

NOTE Confidence: 0.870585560798645

 $00:35:18.000 \longrightarrow 00:35:20.100$ that was dependent on it doesn't

NOTE Confidence: 0.870585560798645

 $00:35:20.100 \longrightarrow 00:35:22.585$ make a lot of sense to many people,

NOTE Confidence: 0.870585560798645

 $00:35:22.590 \longrightarrow 00:35:25.656$ and so lots of debate on this.

NOTE Confidence: 0.870585560798645

00:35:25.660 --> 00:35:27.935 And then we had two trials that

NOTE Confidence: 0.870585560798645

 $00:35:27.935 \longrightarrow 00:35:29.969$ came out which changed again,

NOTE Confidence: 0.870585560798645

 $00:35:29.970 \longrightarrow 00:35:32.476$ the way that we think about this.

NOTE Confidence: 0.870585560798645

00:35:32.480 --> 00:35:35.344 Should this was a trial done in Japan,

NOTE Confidence: 0.870585560798645

 $00:35:35.350 \longrightarrow 00:35:35.787$ chemotherapy.

NOTE Confidence: 0.870585560798645

 $00:35:35.787 \longrightarrow 00:35:38.846$ Within Egypt Artique I so this was

NOTE Confidence: 0.870585560798645

 $00:35:38.846 \longrightarrow 00:35:41.430$ in patients who are newly diagnosed

NOTE Confidence: 0.870585560798645

00:35:41.430 --> 00:35:43.878 EGFR mutant lung cancer and they

NOTE Confidence: 0.870585560798645

 $00:35:43.955 \longrightarrow 00:35:46.579$ either got the EGFR drugs if it new.

NOTE Confidence: 0.870585560798645

 $00:35:46.580 \longrightarrow 00:35:49.136$ Or they got to fit in it plus chemo.

NOTE Confidence: 0.870585560798645

00:35:49.140 --> 00:35:51.114 If they got your fitness plus chemo,

 $00:35:51.120 \longrightarrow 00:35:52.445$ they stayed on maintenance and

NOTE Confidence: 0.870585560798645

 $00:35:52.445 \longrightarrow 00:35:54.230$ at the time it stopped working.

NOTE Confidence: 0.870585560798645

 $00:35:54.230 \longrightarrow 00:35:55.928$ They went on to other therapy.

NOTE Confidence: 0.870585560798645

 $00:35:55.930 \longrightarrow 00:35:57.628$ If they got ripped fitness alone

NOTE Confidence: 0.870585560798645

 $00:35:57.628 \longrightarrow 00:35:58.760$ when it stopped working,

NOTE Confidence: 0.870585560798645

 $00:35:58.760 \longrightarrow 00:36:00.734$ they stopped it and went to chemo.

NOTE Confidence: 0.870585560798645

 $00:36:00.740 \longrightarrow 00:36:02.160$ So it's sort of a.

NOTE Confidence: 0.870585560798645

 $00{:}36{:}02.160 \dashrightarrow 00{:}36{:}03.292$ You get everything together.

NOTE Confidence: 0.870585560798645

 $00:36:03.292 \longrightarrow 00:36:04.707$ The chemo plus that huge.

NOTE Confidence: 0.870585560798645

00:36:04.710 --> 00:36:06.966 If our drug or you get it sequentially,

NOTE Confidence: 0.870585560798645

 $00:36:06.970 \longrightarrow 00:36:08.610$ EGFR and then chemo.

NOTE Confidence: 0.870585560798645

00:36:08.610 --> 00:36:10.178 Again, no summerton period.

NOTE Confidence: 0.870585560798645

 $00{:}36{:}10.178 \dashrightarrow 00{:}36{:}12.108$ This was designed before there

NOTE Confidence: 0.870585560798645

 $00:36:12.108 \longrightarrow 00:36:13.619$ was oh so Barnum.

NOTE Confidence: 0.870585560798645

 $00:36:13.620 \longrightarrow 00:36:16.668$ An when you think about this was the.

00:36:16.670 --> 00:36:18.358 First progression free survival.

NOTE Confidence: 0.870585560798645

 $00{:}36{:}18.358 \dashrightarrow 00{:}36{:}20.890$ So this was for patients who

NOTE Confidence: 0.870585560798645

 $00:36:20.968 \longrightarrow 00:36:22.768$ had chemo plus to fit nib.

NOTE Confidence: 0.870585560798645

 $00:36:22.770 \longrightarrow 00:36:25.122$ Versus Jeff it Nip and the chemo plus

NOTE Confidence: 0.870585560798645

 $00:36:25.122 \longrightarrow 00:36:27.647$ your fitness group stayed on the drug longer.

NOTE Confidence: 0.870585560798645

 $00:36:27.650 \longrightarrow 00:36:29.610$ It was they were still on it

NOTE Confidence: 0.870585560798645

 $00:36:29.610 \longrightarrow 00:36:31.010$ for like 21 months.

NOTE Confidence: 0.870585560798645

 $00:36:31.010 \longrightarrow 00:36:32.840$ The group only get into fitting.

NOTE Confidence: 0.870585560798645

 $00:36:32.840 \longrightarrow 00:36:34.670$ It only worked about 11 months.

NOTE Confidence: 0.87058556079864500:36:34.670 --> 00:36:35.118 You like?

NOTE Confidence: 0.870585560798645

 $00{:}36{:}35.118 \dashrightarrow 00{:}36{:}36.910$ OK right but what you really want to

NOTE Confidence: 0.870585560798645

 $00:36:36.966 \longrightarrow 00:36:38.796$ know is everybody everything together

NOTE Confidence: 0.870585560798645

 $00:36:38.796 \longrightarrow 00:36:40.626$ concurrent versus sequential and when

NOTE Confidence: 0.870585560798645

 $00:36:40.678 \longrightarrow 00:36:42.288$ you looked at everybody together,

NOTE Confidence: 0.870585560798645

00:36:42.290 --> 00:36:43.283 concurrent versus sequential,

NOTE Confidence: 0.870585560798645

 $00:36:43.283 \longrightarrow 00:36:45.600$ the sequential group cut up when they

 $00:36:45.651 \longrightarrow 00:36:47.478$ went from there drifting it to chemo.

NOTE Confidence: 0.870585560798645

 $00:36:47.480 \longrightarrow 00:36:49.629$ It all ended up being the same

NOTE Confidence: 0.870585560798645

00:36:49.629 --> 00:36:51.582 and you think OK, negative study.

NOTE Confidence: 0.870585560798645

00:36:51.582 --> 00:36:53.086 Except overall survival actually

NOTE Confidence: 0.870585560798645

 $00:36:53.086 \longrightarrow 00:36:55.427$ showed a benefit to having gotten

NOTE Confidence: 0.870585560798645

 $00:36:55.427 \longrightarrow 00:36:56.975$ everything at the beginning,

NOTE Confidence: 0.870585560798645

 $00:36:56.980 \longrightarrow 00:36:59.844$ so this has this a little bit confused.

NOTE Confidence: 0.870585560798645

 $00:36:59.850 \longrightarrow 00:37:02.742$ Why would getting everything at the

NOTE Confidence: 0.870585560798645

 $00:37:02.742 \longrightarrow 00:37:06.118$ beginning be better than the A and then B?

NOTE Confidence: 0.870585560798645

 $00:37:06.120 \longrightarrow 00:37:07.180$ If the progression free

NOTE Confidence: 0.870585560798645

 $00:37:07.180 \longrightarrow 00:37:08.240$ Survival's were the same,

NOTE Confidence: 0.870585560798645

 $00:37:08.240 \longrightarrow 00:37:10.360$ so we're still trying to figure this out,

NOTE Confidence: 0.870585560798645

 $00{:}37{:}10.360 \dashrightarrow 00{:}37{:}11.965$ but there was another study

NOTE Confidence: 0.870585560798645

 $00:37:11.965 \longrightarrow 00:37:13.570$ that was done in India.

NOTE Confidence: 0.870585560798645

 $00:37:13.570 \longrightarrow 00:37:14.290$ Similar design.

 $00:37:14.290 \longrightarrow 00:37:16.090$ That also showed that there

NOTE Confidence: 0.870585560798645

 $00{:}37{:}16.090 \dashrightarrow 00{:}37{:}17.962$ was a progression free and

NOTE Confidence: 0.870585560798645

 $00:37:17.962 \longrightarrow 00:37:19.418$ an overall survival benefit.

NOTE Confidence: 0.870585560798645

 $00:37:19.420 \longrightarrow 00:37:21.082$ This is a little bit different

NOTE Confidence: 0.870585560798645

 $00:37:21.082 \longrightarrow 00:37:22.190$ 'cause not everybody who

NOTE Confidence: 0.838739812374115

00:37:22.241 --> 00:37:23.666 started on Jefit and I've

NOTE Confidence: 0.838739812374115

 $00:37:23.666 \longrightarrow 00:37:24.806$ got the chemo afterwards,

NOTE Confidence: 0.838739812374115

00:37:24.810 --> 00:37:26.520 but it kind of supports that,

NOTE Confidence: 0.838739812374115

 $00:37:26.520 \longrightarrow 00:37:27.940$ and so they're ongoing trials.

NOTE Confidence: 0.838739812374115

 $00:37:27.940 \longrightarrow 00:37:29.452$ Now looking at this question and

NOTE Confidence: 0.838739812374115

00:37:29.452 --> 00:37:31.060 including a study with Osimertinib,

NOTE Confidence: 0.838739812374115

 $00:37:31.060 \longrightarrow 00:37:33.797$ so we'll see how this pans out.

NOTE Confidence: 0.838739812374115

00:37:33.800 --> 00:37:35.676 And now I'm going to shift gears

NOTE Confidence: 0.838739812374115

 $00:37:35.676 \longrightarrow 00:37:38.075$ again and go to immune therapy because

NOTE Confidence: 0.838739812374115

 $00:37:38.075 \longrightarrow 00:37:39.985$ that's the other big revolution

NOTE Confidence: 0.838739812374115

00:37:39.985 --> 00:37:42.217 that's happened in the last 15 years,

 $00:37:42.220 \longrightarrow 00:37:43.840$ but it hasn't changed each

NOTE Confidence: 0.838739812374115

 $00:37:43.840 \longrightarrow 00:37:45.460$ of our lung cancer much.

NOTE Confidence: 0.838739812374115

 $00:37:45.460 \longrightarrow 00:37:47.497$ When we look at the first trials

NOTE Confidence: 0.838739812374115

 $00:37:47.497 \longrightarrow 00:37:49.350$ that got immune therapies with,

NOTE Confidence: 0.838739812374115

 $00{:}37{:}49.350 \dashrightarrow 00{:}37{:}51.974$ these are the PD One PD L1 checkpoint

NOTE Confidence: 0.838739812374115

 $00:37:51.974 \longrightarrow 00:37:53.559$ inhibitors approved in lung cancer.

NOTE Confidence: 0.838739812374115

 $00:37:53.560 \longrightarrow 00:37:55.834$ They were given second line to

NOTE Confidence: 0.838739812374115

 $00{:}37{:}55.834 \dashrightarrow 00{:}37{:}58.140$ people who had already had chemo.

NOTE Confidence: 0.838739812374115

 $00:37:58.140 \dashrightarrow 00:37:59.940$ Some of whom were Egypt Farhadi,

NOTE Confidence: 0.838739812374115

 $00:37:59.940 \longrightarrow 00:38:02.292$ Jaffar lung cancer, and it already

NOTE Confidence: 0.838739812374115

 $00:38:02.292 \longrightarrow 00:38:04.420$ had Egypt farland treatments as well.

NOTE Confidence: 0.838739812374115

 $00:38:04.420 \longrightarrow 00:38:06.170$ They were the only subgroup that didn't

NOTE Confidence: 0.838739812374115

 $00{:}38{:}06.170 \dashrightarrow 00{:}38{:}08.178$ do well with the checkpoint inhibitor,

NOTE Confidence: 0.838739812374115

 $00:38:08.180 \longrightarrow 00:38:10.166$ so in these older trials and

NOTE Confidence: 0.838739812374115

 $00:38:10.166 \longrightarrow 00:38:12.179$ I'm looking here at this box.

00:38:12.180 --> 00:38:13.922 Of docetaxel,

NOTE Confidence: 0.838739812374115

 $00{:}38{:}13.922 \dashrightarrow 00{:}38{:}17.406$ chemo versus checkpoint inhibitors.

NOTE Confidence: 0.838739812374115

 $00:38:17.410 \longrightarrow 00:38:18.710$ Those trials showed overall

NOTE Confidence: 0.838739812374115

00:38:18.710 --> 00:38:20.010 survival benefits to everybody.

NOTE Confidence: 0.838739812374115

00:38:20.010 --> 00:38:21.310 With the checkpoint inhibitors,

NOTE Confidence: 0.838739812374115

 $00:38:21.310 \longrightarrow 00:38:23.260$ except patients who had EGFR mutations.

NOTE Confidence: 0.838739812374115

 $00:38:23.260 \longrightarrow 00:38:25.465$ So when we got to the bigger

NOTE Confidence: 0.838739812374115

 $00:38:25.465 \longrightarrow 00:38:26.840$ trials in first line,

NOTE Confidence: 0.838739812374115

 $00:38:26.840 \longrightarrow 00:38:28.136$ patients with EGFR mutations

NOTE Confidence: 0.838739812374115

00:38:28.136 --> 00:38:29.756 weren't included on those trials,

NOTE Confidence: 0.838739812374115

 $00{:}38{:}29.760 \dashrightarrow 00{:}38{:}31.710$ so we have very little data.

NOTE Confidence: 0.838739812374115

 $00:38:31.710 \longrightarrow 00:38:33.660$ This one study that came out

NOTE Confidence: 0.838739812374115

 $00:38:33.660 \longrightarrow 00:38:34.960$ of the Atlantic trial,

NOTE Confidence: 0.838739812374115

 $00:38:34.960 \longrightarrow 00:38:36.910$ which was a phase two study,

NOTE Confidence: 0.838739812374115

00:38:36.910 --> 00:38:39.638 showed that if you had a Jaffar mutant

NOTE Confidence: 0.838739812374115

00:38:39.638 --> 00:38:42.436 lung cancer and your PD L1 level was high,

 $00:38:42.440 \longrightarrow 00:38:43.736$ sometimes the checkpoint inhibitors

NOTE Confidence: 0.838739812374115

 $00:38:43.736 \longrightarrow 00:38:45.685$ worked about 12% of the time.

NOTE Confidence: 0.838739812374115

00:38:45.685 --> 00:38:47.310 But if you had low,

NOTE Confidence: 0.838739812374115

00:38:47.310 --> 00:38:49.338 any different mutation, it never worked.

NOTE Confidence: 0.838739812374115

 $00:38:49.340 \longrightarrow 00:38:50.920$ People have done TK.

NOTE Confidence: 0.838739812374115

 $00:38:50.920 \longrightarrow 00:38:52.500$ I plus immunotherapy combinations.

NOTE Confidence: 0.838739812374115

 $00:38:52.500 \longrightarrow 00:38:54.480$ And they've been totally negative.

NOTE Confidence: 0.838739812374115

 $00{:}38{:}54.480 \dashrightarrow 00{:}38{:}58.248$ Alot of Texas city. No clear benefit so.

NOTE Confidence: 0.838739812374115

 $00:38:58.250 \longrightarrow 00:39:00.374$ We don't get too enthusiastic about

NOTE Confidence: 0.838739812374115

00:39:00.374 --> 00:39:01.790 checkpoint inhibitors in EGFR,

NOTE Confidence: 0.838739812374115

 $00:39:01.790 \longrightarrow 00:39:03.560$ but we think about it.

NOTE Confidence: 0.838739812374115

 $00:39:03.560 \longrightarrow 00:39:04.760$ There was one trial.

NOTE Confidence: 0.838739812374115

 $00:39:04.760 \dashrightarrow 00:39:07.011$ This empower 150 study that was chemo

NOTE Confidence: 0.838739812374115

 $00:39:07.011 \longrightarrow 00:39:08.971$ plus bevacizumab plus immune therapy

NOTE Confidence: 0.838739812374115

 $00:39:08.971 \longrightarrow 00:39:11.316$ that allowed for patients with EGFR

00:39:11.316 --> 00:39:13.458 mutations and that study did show

NOTE Confidence: 0.838739812374115

00:39:13.458 --> 00:39:15.604 potential survival benefit in the subset,

NOTE Confidence: 0.838739812374115

 $00:39:15.604 \longrightarrow 00:39:18.588$ but we have to be really careful because

NOTE Confidence: 0.838739812374115

 $00:39:18.588 \longrightarrow 00:39:22.210$ this was a small subset of a bigger study.

NOTE Confidence: 0.838739812374115

 $00:39:22.210 \longrightarrow 00:39:23.745$ It wasn't really powered to

NOTE Confidence: 0.838739812374115

 $00:39:23.745 \longrightarrow 00:39:24.973$ answer the specific question,

NOTE Confidence: 0.838739812374115

 $00:39:24.980 \longrightarrow 00:39:26.520$ and so their ongoing trials.

NOTE Confidence: 0.838739812374115

 $00:39:26.520 \longrightarrow 00:39:28.368$ Now looking at this so we

NOTE Confidence: 0.838739812374115

 $00{:}39{:}28.368 \dashrightarrow 00{:}39{:}29.600$ still have questions around.

NOTE Confidence: 0.838739812374115

 $00:39:29.600 \longrightarrow 00:39:31.693$ What do we do with immune therapy

NOTE Confidence: 0.838739812374115

 $00{:}39{:}31.693 \dashrightarrow 00{:}39{:}33.909$ in the setting of EGFR lung cancer?

NOTE Confidence: 0.838739812374115

 $00:39:33.910 \longrightarrow 00:39:35.758$ So to put all that together,

NOTE Confidence: 0.838739812374115

 $00:39:35.760 \longrightarrow 00:39:37.415$ each of Arcgis improved response

NOTE Confidence: 0.838739812374115

 $00:39:37.415 \longrightarrow 00:39:39.070$ rate and progression free survival

NOTE Confidence: 0.838739812374115

 $00:39:39.127 \longrightarrow 00:39:40.907$ compared to first line chemotherapy.

NOTE Confidence: 0.838739812374115

 $00:39:40.910 \longrightarrow 00:39:42.240$ But you have to test.

 $00:39:42.240 \longrightarrow 00:39:43.824$ You have to know the Jaffar

NOTE Confidence: 0.838739812374115

 $00:39:43.824 \longrightarrow 00:39:44.616$ mutation is there.

NOTE Confidence: 0.838739812374115

 $00:39:44.620 \longrightarrow 00:39:46.210$ Oh, summer at numbers are preferred.

NOTE Confidence: 0.838739812374115

 $00:39:46.210 \longrightarrow 00:39:48.866$ First line treatment from the floor of trial.

NOTE Confidence: 0.838739812374115

 $00:39:48.870 \longrightarrow 00:39:50.860$ Veg F inhibition improves response

NOTE Confidence: 0.838739812374115

 $00:39:50.860 \longrightarrow 00:39:52.452$ and progression free survival

NOTE Confidence: 0.838739812374115

 $00:39:52.452 \longrightarrow 00:39:54.120$ when added DJ fourty keys.

NOTE Confidence: 0.838739812374115

00:39:54.120 --> 00:39:56.724 But we don't know if it

NOTE Confidence: 0.838739812374115

 $00:39:56.724 \longrightarrow 00:39:58.026$ improves overall survival.

NOTE Confidence: 0.838739812374115

 $00:39:58.030 \longrightarrow 00:39:59.955$ Chemotherapy may improve overall survival

NOTE Confidence: 0.838739812374115

00:39:59.955 --> 00:40:02.220 in combination with EGFR TK eyes,

NOTE Confidence: 0.838739812374115

 $00:40:02.220 \longrightarrow 00:40:04.485$ but we're still trying to

NOTE Confidence: 0.838739812374115

 $00{:}40{:}04.485 \dashrightarrow 00{:}40{:}05.844$ answer that definitively.

NOTE Confidence: 0.838739812374115

 $00{:}40{:}05.850 \to 00{:}40{:}07.548$ And immune the rapy activity is very,

NOTE Confidence: 0.838739812374115

 $00:40:07.550 \longrightarrow 00:40:08.910$ very limited and there's a

 $00:40:08.910 \longrightarrow 00:40:10.660$ lot we need to explore there,

NOTE Confidence: 0.838739812374115

 $00:40:10.660 \longrightarrow 00:40:12.358$ and we're still dealing with the

NOTE Confidence: 0.838739812374115

 $00:40:12.358 \longrightarrow 00:40:13.490$ fact that resistance develops.

NOTE Confidence: 0.861340403556824

 $00:40:13.490 \longrightarrow 00:40:14.910$ So we've made tremendous progress.

NOTE Confidence: 0.861340403556824

 $00:40:14.910 \longrightarrow 00:40:18.015$ But we still have a long way to go.

NOTE Confidence: 0.861340403556824

 $00:40:18.020 \longrightarrow 00:40:19.544$ And that's where communication

NOTE Confidence: 0.861340403556824

 $00:40:19.544 \longrightarrow 00:40:21.449$ is so important because some

NOTE Confidence: 0.861340403556824

00:40:21.449 --> 00:40:22.948 patients do remarkably well.

NOTE Confidence: 0.861340403556824

00:40:22.950 --> 00:40:24.840 I was meeting with one

NOTE Confidence: 0.861340403556824

 $00:40:24.840 \longrightarrow 00:40:26.352$ of my patients yesterday.

NOTE Confidence: 0.861340403556824

 $00:40:26.360 \longrightarrow 00:40:28.250$ I've been caring for for

NOTE Confidence: 0.861340403556824

00:40:28.250 --> 00:40:29.762 eight years with metastatic,

NOTE Confidence: 0.861340403556824

00:40:29.770 --> 00:40:30.794 EGFR, mutant lung cancer.

NOTE Confidence: 0.861340403556824

 $00:40:30.794 \longrightarrow 00:40:32.074$ We're having to think about

NOTE Confidence: 0.861340403556824

 $00:40:32.074 \longrightarrow 00:40:33.408$ her next line of treatment,

NOTE Confidence: 0.861340403556824

 $00:40:33.410 \longrightarrow 00:40:34.887$ but she still on her last name,

 $00:40:34.890 \longrightarrow 00:40:36.660$ but it's been eight years.

NOTE Confidence: 0.861340403556824

00:40:36.660 --> 00:40:37.868 Some patients do remarkably

NOTE Confidence: 0.861340403556824

 $00:40:37.868 \longrightarrow 00:40:39.076$ poorly on another patient.

NOTE Confidence: 0.861340403556824

00:40:39.080 --> 00:40:41.504 I saw, I think it was last Thursday.

NOTE Confidence: 0.861340403556824

00:40:41.510 --> 00:40:43.926 He's never had a good response to anything,

NOTE Confidence: 0.861340403556824

 $00:40:43.930 \longrightarrow 00:40:45.925$ even though he has a clear EGFR

NOTE Confidence: 0.861340403556824

 $00:40:45.925 \longrightarrow 00:40:47.570$ mutation in his lung cancer.

NOTE Confidence: 0.861340403556824

 $00:40:47.570 \longrightarrow 00:40:49.388$ And so why are they different?

NOTE Confidence: 0.861340403556824

 $00:40:49.390 \longrightarrow 00:40:51.758$ I can't tell, just based on the age

NOTE Confidence: 0.861340403556824

 $00{:}40{:}51.758 \dashrightarrow 00{:}40{:}53.630$ of permutations that they have there.

NOTE Confidence: 0.861340403556824

 $00:40:53.630 \longrightarrow 00:40:54.838$ Other factors that we're

NOTE Confidence: 0.861340403556824

 $00:40:54.838 \longrightarrow 00:40:55.744$ trying to understand,

NOTE Confidence: 0.861340403556824

 $00{:}40{:}55.750 \dashrightarrow 00{:}40{:}57.906$ but it's important than that when we

NOTE Confidence: 0.861340403556824

00:40:57.906 --> 00:41:00.288 talk about how someone is going to do,

NOTE Confidence: 0.861340403556824

00:41:00.290 --> 00:41:02.108 we're never too confident in that,

00:41:02.110 --> 00:41:03.630 because we really don't know,

NOTE Confidence: 0.861340403556824

 $00{:}41{:}03.630 \dashrightarrow 00{:}41{:}05.496$ so we can often predict outcomes

NOTE Confidence: 0.861340403556824

 $00:41:05.496 \longrightarrow 00:41:06.740$ that we can't always.

NOTE Confidence: 0.861340403556824

 $00:41:06.740 \longrightarrow 00:41:09.228$ And we have to strive to keep the

NOTE Confidence: 0.861340403556824

00:41:09.228 --> 00:41:12.079 trust of the patient and their family,

NOTE Confidence: 0.861340403556824

 $00:41:12.080 \longrightarrow 00:41:14.565$ to help them relive given this uncertainty.

NOTE Confidence: 0.861340403556824

00:41:14.570 --> 00:41:15.990 And so I mentioned,

NOTE Confidence: 0.861340403556824

00:41:15.990 --> 00:41:19.199 I'm going to leave this in a little bit.

NOTE Confidence: 0.861340403556824

 $00:41:19.200 \longrightarrow 00:41:21.162$ So this was when breath becomes

NOTE Confidence: 0.861340403556824

00:41:21.162 --> 00:41:23.120 air written by Paul Kleanthi,

NOTE Confidence: 0.861340403556824

00:41:23.120 --> 00:41:23.972 Yale medical student,

NOTE Confidence: 0.861340403556824

 $00:41:23.972 \longrightarrow 00:41:25.960$ who then went on to do his

NOTE Confidence: 0.861340403556824

00:41:26.025 --> 00:41:28.097 neurosurgical training at Stanford,

NOTE Confidence: 0.861340403556824

 $00:41:28.100 \longrightarrow 00:41:29.520$ and he unfortunately died

NOTE Confidence: 0.861340403556824

 $00:41:29.520 \longrightarrow 00:41:31.295$ of EGFR mutant lung cancer.

NOTE Confidence: 0.861340403556824

00:41:31.300 --> 00:41:34.148 But while he was living with his disease,

 $00:41:34.150 \longrightarrow 00:41:35.930$ he wrote this amazing book.

NOTE Confidence: 0.861340403556824

 $00:41:35.930 \longrightarrow 00:41:38.378$ One of the quotes that I.

NOTE Confidence: 0.861340403556824

 $00:41:38.380 \longrightarrow 00:41:40.977$ Lee love from this is that the

NOTE Confidence: 0.861340403556824

00:41:40.977 --> 00:41:43.128 reason that doctors don't give

NOTE Confidence: 0.861340403556824

00:41:43.128 --> 00:41:45.056 patients specific prognosis is

NOTE Confidence: 0.861340403556824

 $00:41:45.056 \longrightarrow 00:41:47.580$ not merely because they cannot.

NOTE Confidence: 0.861340403556824

 $00:41:47.580 \longrightarrow 00:41:49.902$ What patients seek is not scientific

NOTE Confidence: 0.861340403556824

00:41:49.902 --> 00:41:51.450 knowledge that doctors hide

NOTE Confidence: 0.861340403556824

 $00:41:51.515 \longrightarrow 00:41:53.270$ but existential authenticity.

NOTE Confidence: 0.861340403556824

00:41:53.270 --> 00:41:56.014 Each person must find on his or

NOTE Confidence: 0.861340403556824

 $00:41:56.014 \longrightarrow 00:41:58.600$ her own getting too deeply into

NOTE Confidence: 0.861340403556824

 $00:41:58.600 \longrightarrow 00:42:00.810$ statistics is like trying to

NOTE Confidence: 0.861340403556824

 $00{:}42{:}00.810 \dashrightarrow 00{:}42{:}03.778$ quench a thirst with salty water.

NOTE Confidence: 0.861340403556824

 $00{:}42{:}03.780 \dashrightarrow 00{:}42{:}06.582$ The angst of facing mortality has

NOTE Confidence: 0.861340403556824

 $00:42:06.582 \longrightarrow 00:42:09.100$ no remedy in probability and so.

 $00:42:09.100 \longrightarrow 00:42:10.540$ That really captures the essence,

NOTE Confidence: 0.861340403556824

 $00:42:10.540 \longrightarrow 00:42:12.260$ I think of what we tried to do

NOTE Confidence: 0.861340403556824

00:42:12.260 --> 00:42:14.000 in this communication setting,

NOTE Confidence: 0.861340403556824

 $00:42:14.000 \longrightarrow 00:42:15.815$ but written obviously far more

NOTE Confidence: 0.861340403556824

00:42:15.815 --> 00:42:17.630 eloquently than I would ever

NOTE Confidence: 0.861340403556824

 $00:42:17.696 \longrightarrow 00:42:18.986$ have been able to do.

NOTE Confidence: 0.861340403556824

 $00:42:18.990 \longrightarrow 00:42:21.986$ And obviously we miss Paul very much.

NOTE Confidence: 0.861340403556824

 $00:42:21.990 \longrightarrow 00:42:23.957$ So when we talk about communication tips,

NOTE Confidence: 0.861340403556824

 $00:42:23.960 \longrightarrow 00:42:25.808$ these are a few of the things

NOTE Confidence: 0.861340403556824

00:42:25.808 --> 00:42:27.630 that I like to think about.

NOTE Confidence: 0.861340403556824

 $00{:}42{:}27.630 \dashrightarrow 00{:}42{:}29.597$ One of them is about the numbers.

NOTE Confidence: 0.861340403556824

00:42:29.600 --> 00:42:30.728 Just as Paul said,

NOTE Confidence: 0.861340403556824

00:42:30.728 --> 00:42:32.138 but numbers to get remembered.

NOTE Confidence: 0.861340403556824

00:42:32.140 --> 00:42:33.988 If you have an hour long conversation

NOTE Confidence: 0.861340403556824

 $00:42:33.988 \longrightarrow 00:42:36.013$ with the patient and you mentioned a

NOTE Confidence: 0.861340403556824

 $00:42:36.013 \longrightarrow 00:42:37.777$ number when they leave the conversation,

 $00:42:37.780 \longrightarrow 00:42:39.190$ that's what they will remember.

NOTE Confidence: 0.861340403556824

 $00:42:39.190 \longrightarrow 00:42:41.444$ So you have to be very careful

NOTE Confidence: 0.861340403556824

 $00:42:41.444 \longrightarrow 00:42:42.880$ how you use them.

NOTE Confidence: 0.861340403556824

 $00:42:42.880 \longrightarrow 00:42:44.405$ And be very careful how

NOTE Confidence: 0.861340403556824

 $00:42:44.405 \longrightarrow 00:42:45.320$ prognosis is discussed.

NOTE Confidence: 0.861340403556824

 $00:42:45.320 \longrightarrow 00:42:46.235$ Avoid codeine medians,

NOTE Confidence: 0.861340403556824

00:42:46.235 --> 00:42:48.370 because if you quote a median again,

NOTE Confidence: 0.861340403556824

 $00:42:48.370 \longrightarrow 00:42:50.200$ that's the only thing Tillman remembers.

NOTE Confidence: 0.861340403556824

 $00:42:50.200 \longrightarrow 00:42:51.725$ Not everything we know about

NOTE Confidence: 0.861340403556824

 $00:42:51.725 \longrightarrow 00:42:52.945$ what a median means.

NOTE Confidence: 0.861340403556824

00:42:52.950 --> 00:42:56.325 She always give a range and I talk about.

NOTE Confidence: 0.861340403556824

 $00:42:56.330 \longrightarrow 00:42:57.558$ The worst possible outcomes,

NOTE Confidence: 0.861340403556824

 $00{:}42{:}57.558 \to 00{:}42{:}59.750$ but I also talk about the best,

NOTE Confidence: 0.861340403556824

 $00:42:59.750 \longrightarrow 00:43:01.927$ but whenever I talk about the best,

NOTE Confidence: 0.861340403556824

00:43:01.930 --> 00:43:03.580 I always make sure I can

 $00:43:03.580 \longrightarrow 00:43:04.680$ think of a specific

NOTE Confidence: 0.885906279087067

 $00:43:04.752 \longrightarrow 00:43:07.293$ person who was there 'cause otherwise you're

NOTE Confidence: 0.885906279087067

 $00:43:07.293 \longrightarrow 00:43:10.018$ just making a number up out of the air.

NOTE Confidence: 0.885906279087067

 $00:43:10.020 \longrightarrow 00:43:11.555$ The prognosis for any individual

NOTE Confidence: 0.885906279087067

 $00:43:11.555 \longrightarrow 00:43:12.783$ of course evolves overtime,

NOTE Confidence: 0.885906279087067

 $00:43:12.790 \longrightarrow 00:43:14.330$ so help the patient to

NOTE Confidence: 0.885906279087067

00:43:14.330 --> 00:43:15.870 understand that and their family,

NOTE Confidence: 0.885906279087067

 $00:43:15.870 \longrightarrow 00:43:17.795$ and then try to guide the patient

NOTE Confidence: 0.885906279087067

 $00:43:17.795 \longrightarrow 00:43:19.260$ in making life decisions.

NOTE Confidence: 0.885906279087067

 $00:43:19.260 \longrightarrow 00:43:21.395$ They need to make based on probabilities

NOTE Confidence: 0.885906279087067

00:43:21.395 --> 00:43:23.260 of how much time they have,

NOTE Confidence: 0.885906279087067

00:43:23.260 --> 00:43:25.040 knowing that there's that uncertainty

NOTE Confidence: 0.885906279087067

 $00{:}43{:}25.040 \dashrightarrow 00{:}43{:}26.464$ encouraging patients to think

NOTE Confidence: 0.885906279087067

 $00:43:26.464 \longrightarrow 00:43:28.186$ about the things they need to do,

NOTE Confidence: 0.885906279087067

 $00:43:28.190 \longrightarrow 00:43:31.398$ and also about what they want to do.

NOTE Confidence: 0.885906279087067

 $00:43:31.400 \longrightarrow 00:43:33.398$ It's really a terrible thing if

 $00:43:33.398 \longrightarrow 00:43:35.104$ someone assumes that they have

NOTE Confidence: 0.885906279087067

 $00:43:35.104 \longrightarrow 00:43:36.958$ many years left when they don't,

NOTE Confidence: 0.885906279087067

 $00:43:36.960 \longrightarrow 00:43:39.361$ and therefore they don't do the things

NOTE Confidence: 0.885906279087067

 $00:43:39.361 \longrightarrow 00:43:42.015$ they would have done if they had known

NOTE Confidence: 0.885906279087067

 $00:43:42.015 \longrightarrow 00:43:44.149$ that they didn't have alot of time.

NOTE Confidence: 0.885906279087067

00:43:44.150 --> 00:43:45.902 But it's also terrible to take

NOTE Confidence: 0.885906279087067

 $00:43:45.902 \longrightarrow 00:43:47.505$ away hope from someone because

NOTE Confidence: 0.885906279087067

00:43:47.505 --> 00:43:49.385 there always is that uncertainty,

NOTE Confidence: 0.885906279087067

 $00:43:49.390 \longrightarrow 00:43:51.352$ so you try to guide but

NOTE Confidence: 0.885906279087067

 $00:43:51.352 \longrightarrow 00:43:52.660$ leave uncertainty on going.

NOTE Confidence: 0.885906279087067

00:43:52.660 --> 00:43:54.024 Dialogue is obviously critical.

NOTE Confidence: 0.885906279087067

 $00:43:54.024 \longrightarrow 00:43:57.050$ I would bring up the concept of risk and

NOTE Confidence: 0.885906279087067

 $00:43:57.050 \longrightarrow 00:43:59.199$ benefit in regard to the rapy fairly early,

NOTE Confidence: 0.885906279087067

 $00{:}43{:}59.200 \dashrightarrow 00{:}44{:}01.864$ and I frequently talk about that when I'm.

NOTE Confidence: 0.885906279087067

 $00:44:01.870 \longrightarrow 00:44:03.688$ Reviewing things because at some point

 $00:44:03.688 \longrightarrow 00:44:06.387$ we are going to get to that point where

NOTE Confidence: 0.885906279087067

 $00:44:06.387 \longrightarrow 00:44:08.564$ the risk of ongoing therapy is going

NOTE Confidence: 0.885906279087067

 $00:44:08.564 \longrightarrow 00:44:10.284$ to outweigh the potential benefit.

NOTE Confidence: 0.885906279087067

00:44:10.290 --> 00:44:13.410 And if you bring that up for the first time,

NOTE Confidence: 0.885906279087067

 $00:44:13.410 \longrightarrow 00:44:15.594$ then it's a much more difficult conversation.

NOTE Confidence: 0.885906279087067

00:44:15.600 --> 00:44:17.676 It's also really critical as you

NOTE Confidence: 0.885906279087067

 $00:44:17.676 \longrightarrow 00:44:19.650$ engage the patient who is the

NOTE Confidence: 0.885906279087067

 $00:44:19.650 \longrightarrow 00:44:21.210$ key person in all of this.

NOTE Confidence: 0.885906279087067

00:44:21.210 --> 00:44:23.642 You've got to make sure the family also

NOTE Confidence: 0.885906279087067

 $00:44:23.642 \longrightarrow 00:44:26.135$ knows what's going on and is aware of

NOTE Confidence: 0.885906279087067

 $00{:}44{:}26.135 \dashrightarrow 00{:}44{:}27.726$ the information that you're talking

NOTE Confidence: 0.885906279087067

 $00:44:27.726 \longrightarrow 00:44:29.931$ about because the family is going to

NOTE Confidence: 0.885906279087067

 $00:44:29.931 \longrightarrow 00:44:32.150$ live with the memory of the patient.

NOTE Confidence: 0.885906279087067 00:44:32.150 --> 00:44:32.502 Forever. NOTE Confidence: 0.885906279087067

 $00:44:32.502 \longrightarrow 00:44:34.966$ And it's really important that there are

NOTE Confidence: 0.885906279087067

 $00:44:34.966 \longrightarrow 00:44:37.955$ part of those decisions as much as possible,

00:44:37.960 --> 00:44:39.766 not to guide the patient 'cause it

NOTE Confidence: 0.885906279087067

 $00:44:39.766 \longrightarrow 00:44:41.779$ needs to be the patients choices,

NOTE Confidence: 0.885906279087067

00:44:41.780 --> 00:44:44.540 but to understand how the patient was making

NOTE Confidence: 0.885906279087067

 $00:44:44.540 \longrightarrow 00:44:46.776$ those choices and to be a part of that.

NOTE Confidence: 0.885906279087067

00:44:46.780 --> 00:44:48.892 I think that really helps with the piece

NOTE Confidence: 0.885906279087067

00:44:48.892 --> 00:44:51.186 for the family as they continue living,

NOTE Confidence: 0.885906279087067

00:44:51.190 --> 00:44:53.698 having lost an important loved one.

NOTE Confidence: 0.885906279087067

 $00{:}44{:}53.700 \dashrightarrow 00{:}44{:}55.180$ When having a Frank conversation,

NOTE Confidence: 0.885906279087067

 $00{:}44{:}55.180 \dashrightarrow 00{:}44{:}57.420$ pay careful attention to how you're being

NOTE Confidence: 0.885906279087067

 $00{:}44{:}57.420 \dashrightarrow 00{:}44{:}59.600$ heard harder now in the world of zoom.

NOTE Confidence: 0.885906279087067

 $00:44:59.600 \longrightarrow 00:45:01.532$ Much easier when you're in a room

NOTE Confidence: 0.885906279087067

 $00:45:01.532 \longrightarrow 00:45:03.525$ with someone and you can watch the

NOTE Confidence: 0.885906279087067

 $00{:}45{:}03.525 \dashrightarrow 00{:}45{:}05.181$ body language in their faces and

NOTE Confidence: 0.885906279087067

 $00:45:05.242 \longrightarrow 00:45:07.162$ and make sure that everyone is

NOTE Confidence: 0.885906279087067

 $00:45:07.162 \longrightarrow 00:45:08.988$ engaged in an understanding so that

 $00:45:08.988 \longrightarrow 00:45:10.871$ you can back up and rephrase as

NOTE Confidence: 0.885906279087067

00:45:10.871 --> 00:45:12.579 necessary to get everyone together.

NOTE Confidence: 0.885906279087067

 $00{:}45{:}12.580 \dashrightarrow 00{:}45{:}14.284$ Really important to check back in

NOTE Confidence: 0.885906279087067

 $00:45:14.284 \longrightarrow 00:45:16.120$ with the family and the patient.

NOTE Confidence: 0.885906279087067

 $00:45:16.120 \longrightarrow 00:45:18.670$ If you're making a decision.

NOTE Confidence: 0.885906279087067

 $00:45:18.670 \longrightarrow 00:45:19.850$ To have everybody understand

NOTE Confidence: 0.885906279087067

 $00:45:19.850 \longrightarrow 00:45:21.325$ why that decision was made.

NOTE Confidence: 0.885906279087067

 $00{:}45{:}21.330 \dashrightarrow 00{:}45{:}22.810$ While we're choosing a particular

NOTE Confidence: 0.885906279087067

 $00:45:22.810 \longrightarrow 00:45:23.698$ therapy or another,

NOTE Confidence: 0.885906279087067

00:45:23.700 --> 00:45:25.765 because it's not always going to work,

NOTE Confidence: 0.885906279087067

 $00:45:25.770 \longrightarrow 00:45:27.856$ and it's important that everybody knows how

NOTE Confidence: 0.885906279087067

 $00:45:27.856 \longrightarrow 00:45:30.428$ we ended up making that decision together.

NOTE Confidence: 0.885906279087067

 $00:45:30.430 \longrightarrow 00:45:32.224$ If you hear a really strong

NOTE Confidence: 0.885906279087067

00:45:32.224 --> 00:45:34.040 objection to a particular treatment,

NOTE Confidence: 0.885906279087067

 $00:45:34.040 \longrightarrow 00:45:35.992$ it's always good to ask why there are

NOTE Confidence: 0.885906279087067

00:45:35.992 --> 00:45:38.188 a lot of stories that people have

 $00:45:38.188 \longrightarrow 00:45:40.296$ based on their own experiences of

NOTE Confidence: 0.885906279087067

 $00{:}45{:}40.296 {\:{\mbox{--}}}{>}\ 00{:}45{:}41.888$ family members experiences someone

NOTE Confidence: 0.885906279087067

 $00:45:41.888 \longrightarrow 00:45:43.878$ they heard about bad experience.

NOTE Confidence: 0.868247807025909

 $00:45:43.880 \longrightarrow 00:45:46.058$ To understand why what the particular

NOTE Confidence: 0.868247807025909

 $00:45:46.058 \longrightarrow 00:45:48.143$ patients facing is either similar or

NOTE Confidence: 0.868247807025909

 $00:45:48.143 \longrightarrow 00:45:50.159$ different that can help a lot too.

NOTE Confidence: 0.868247807025909

 $00:45:50.160 \longrightarrow 00:45:51.276$ If it all possible,

NOTE Confidence: 0.868247807025909

 $00:45:51.276 \longrightarrow 00:45:52.950$ never answer your phone or pager

NOTE Confidence: 0.868247807025909

 $00{:}45{:}53.003 \dashrightarrow 00{:}45{:}54.719$ when you're in with the patient.

NOTE Confidence: 0.868247807025909

 $00:45:54.720 \longrightarrow 00:45:55.880$ 100% attention really matters,

NOTE Confidence: 0.868247807025909

 $00:45:55.880 \longrightarrow 00:45:57.330$ no matter how crazy are

NOTE Confidence: 0.868247807025909

 $00:45:57.330 \longrightarrow 00:45:58.708$ busy things are in clinic,

NOTE Confidence: 0.868247807025909

 $00{:}45{:}58.710 \dashrightarrow 00{:}46{:}00.990$ and for those of us who are commissioned,

NOTE Confidence: 0.868247807025909

 $00:46:00.990 \longrightarrow 00:46:02.985$ you walk out of the exam room.

NOTE Confidence: 0.868247807025909

00:46:02.990 --> 00:46:04.742 We got five people talking with

 $00:46:04.742 \longrightarrow 00:46:06.231$ you about different things you're

NOTE Confidence: 0.868247807025909

 $00:46:06.231 \longrightarrow 00:46:07.827$ trying to process all of that,

NOTE Confidence: 0.868247807025909

00:46:07.830 --> 00:46:10.395 but when you go back in the exam room,

NOTE Confidence: 0.868247807025909

 $00:46:10.400 \longrightarrow 00:46:12.388$ you just have to be there 100%.

NOTE Confidence: 0.868247807025909

 $00:46:12.390 \longrightarrow 00:46:13.815$ Com you're listening and that

NOTE Confidence: 0.868247807025909

 $00:46:13.815 \longrightarrow 00:46:15.240$ really makes a big difference.

NOTE Confidence: 0.868247807025909

 $00:46:15.240 \longrightarrow 00:46:16.932$ Also, because the really the key

NOTE Confidence: 0.868247807025909

 $00:46:16.932 \longrightarrow 00:46:18.686$ is being sure that the patient

NOTE Confidence: 0.868247807025909

 $00:46:18.686 \longrightarrow 00:46:20.660$ and the family know that you care

NOTE Confidence: 0.868247807025909

 $00:46:20.660 \longrightarrow 00:46:22.389$ about your patient as a person.

NOTE Confidence: 0.868247807025909

 $00{:}46{:}22.390 \dashrightarrow 00{:}46{:}24.094$ And that you are guiding them

NOTE Confidence: 0.868247807025909

 $00:46:24.094 \longrightarrow 00:46:25.945$ to make the best decisions that

NOTE Confidence: 0.868247807025909

 $00:46:25.945 \longrightarrow 00:46:28.234$ can help them live as long as

NOTE Confidence: 0.868247807025909

 $00{:}46{:}28.234 \dashrightarrow 00{:}46{:}30.230$ possible with good quality of life.

NOTE Confidence: 0.868247807025909

 $00:46:30.230 \longrightarrow 00:46:32.491$ And I'm going to and then with

NOTE Confidence: 0.868247807025909

00:46:32.491 --> 00:46:34.440 a quote from Joe Feldman,

00:46:34.440 --> 00:46:35.844 who's amazing patient advocate

NOTE Confidence: 0.868247807025909

 $00:46:35.844 \longrightarrow 00:46:37.248$ within the EGFR world.

NOTE Confidence: 0.868247807025909

 $00:46:37.250 \longrightarrow 00:46:40.058$ And she does a lot with the ice,

NOTE Confidence: 0.868247807025909

 $00:46:40.060 \longrightarrow 00:46:42.166$ else she is a frequent speaker

NOTE Confidence: 0.868247807025909

 $00:46:42.166 \longrightarrow 00:46:43.219$ at our conferences,

NOTE Confidence: 0.868247807025909

00:46:43.220 --> 00:46:44.970 including our North America conference.

NOTE Confidence: 0.868247807025909

 $00:46:44.970 \longrightarrow 00:46:47.076$ And we were just there together

NOTE Confidence: 0.868247807025909

 $00:46:47.076 \longrightarrow 00:46:48.480$ a few weeks ago,

NOTE Confidence: 0.868247807025909

 $00:46:48.480 \longrightarrow 00:46:50.310$ and she talked about treating the

NOTE Confidence: 0.868247807025909

 $00:46:50.310 \longrightarrow 00:46:52.340$ whole patient on an intellectual,

NOTE Confidence: 0.868247807025909

 $00{:}46{:}52.340 \dashrightarrow 00{:}46{:}53.800$ physical, and emotional level.

NOTE Confidence: 0.868247807025909

 $00{:}46{:}53.800 \dashrightarrow 00{:}46{:}56.377$ We need to know that the doctor

NOTE Confidence: 0.868247807025909

 $00{:}46{:}56.377 \dashrightarrow 00{:}46{:}58.603$ caring for us as a patient cares

NOTE Confidence: 0.868247807025909

00:46:58.603 --> 00:47:00.746 about us what we patients value

NOTE Confidence: 0.868247807025909

 $00:47:00.746 \longrightarrow 00:47:02.894$ can't be quantified in a curve.

 $00:47:02.900 \longrightarrow 00:47:04.630$ Hazard ratios and clinical trial

NOTE Confidence: 0.868247807025909

 $00:47:04.630 \longrightarrow 00:47:06.360$ numbers don't translate to real

NOTE Confidence: 0.868247807025909

 $00:47:06.422 \longrightarrow 00:47:08.306$ life or capture the une quantifiable

NOTE Confidence: 0.868247807025909

 $00:47:08.306 \longrightarrow 00:47:10.160$ meaning of a person's life show.

NOTE Confidence: 0.868247807025909

00:47:10.160 --> 00:47:11.480 I thought she really,

NOTE Confidence: 0.868247807025909

 $00:47:11.480 \longrightarrow 00:47:14.120$ really captured summed it up well with that.

NOTE Confidence: 0.868247807025909

 $00:47:14.120 \longrightarrow 00:47:16.460$ OK, so overall summary.

NOTE Confidence: 0.868247807025909

 $00:47:16.460 \longrightarrow 00:47:18.170$ Over the last 15 years,

NOTE Confidence: 0.868247807025909

 $00{:}47{:}18.170 \dashrightarrow 00{:}47{:}20.258$ we know that Egypt artique eyes

NOTE Confidence: 0.868247807025909

 $00{:}47{:}20.258 \rightarrow 00{:}47{:}23.034$ alone or the standard of care in a

NOTE Confidence: 0.868247807025909

 $00{:}47{:}23.034 \to 00{:}47{:}25.002$ patient with EGFR mutant lung cancer,

NOTE Confidence: 0.868247807025909

 $00:47:25.010 \longrightarrow 00:47:28.139$ but you have to find the mutation.

NOTE Confidence: 0.868247807025909

 $00:47:28.140 \longrightarrow 00:47:28.964$ Veggie combinations.

NOTE Confidence: 0.868247807025909

00:47:28.964 --> 00:47:29.376 Encouraging,

NOTE Confidence: 0.868247807025909

 $00:47:29.376 \longrightarrow 00:47:32.260$ but so far we haven't seen that

NOTE Confidence: 0.868247807025909

 $00:47:32.324 \longrightarrow 00:47:34.859$ translate into overall survival benefit.

00:47:34.860 --> 00:47:36.600 EGFR combinations look really promising,

NOTE Confidence: 0.868247807025909

 $00:47:36.600 \longrightarrow 00:47:38.012$ but not standard yet.

NOTE Confidence: 0.868247807025909

 $00:47:38.012 \longrightarrow 00:47:39.071$ Chemotherapy combinations maybe

NOTE Confidence: 0.868247807025909

 $00:47:39.071 \longrightarrow 00:47:40.760$ will move into first line,

NOTE Confidence: 0.868247807025909

 $00:47:40.760 \longrightarrow 00:47:42.836$ seemed to have improved overall survival,

NOTE Confidence: 0.868247807025909

00:47:42.840 --> 00:47:44.922 but we haven't really bought that

NOTE Confidence: 0.868247807025909

 $00:47:44.922 \longrightarrow 00:47:46.328$ concept yet. Immune therapy.

NOTE Confidence: 0.868247807025909

 $00:47:46.328 \longrightarrow 00:47:48.452$ That's a promise for so many

NOTE Confidence: 0.868247807025909

 $00{:}47{:}48.452 \dashrightarrow 00{:}47{:}50.203$ patients living with lung cancer

NOTE Confidence: 0.868247807025909

 $00:47:50.203 \longrightarrow 00:47:51.515$ and other cancers now.

NOTE Confidence: 0.868247807025909

 $00{:}47{:}51.520 \dashrightarrow 00{:}47{:}53.949$ But in the world of EGFR therapy,

NOTE Confidence: 0.868247807025909

 $00:47:53.950 \longrightarrow 00:47:56.379$ there's a lot we don't understand yet.

NOTE Confidence: 0.868247807025909

 $00{:}47{:}56.380 \dashrightarrow 00{:}47{:}58.718$ And that brings us to the fact

NOTE Confidence: 0.868247807025909

 $00:47:58.718 \longrightarrow 00:48:01.102$ that for all of these advances

NOTE Confidence: 0.868247807025909

 $00:48:01.102 \longrightarrow 00:48:03.694$ and wonderful things that we do.

 $00:48:03.700 \longrightarrow 00:48:05.230$ We still aren't caring people.

NOTE Confidence: 0.868247807025909

 $00{:}48{:}05.230 \dashrightarrow 00{:}48{:}08.280$ We still have a lot that we need to do.

NOTE Confidence: 0.868247807025909

 $00{:}48{:}08.280 \dashrightarrow 00{:}48{:}10.110$ I have intentionally not gone into.

NOTE Confidence: 0.868247807025909

 $00:48:10.110 \longrightarrow 00:48:12.238$ What do we do in early stage?

NOTE Confidence: 0.868247807025909

 $00:48:12.240 \longrightarrow 00:48:13.652$ Lung cancer therapy today?

NOTE Confidence: 0.868247807025909

 $00{:}48{:}13.652 \dashrightarrow 00{:}48{:}15.417$ 'cause there's been some tremendous

NOTE Confidence: 0.868247807025909

00:48:15.417 --> 00:48:16.892 advances there or with Theodore

NOTE Confidence: 0.868247807025909

00:48:16.892 --> 00:48:18.500 trial and other ways of looking

NOTE Confidence: 0.868247807025909

 $00{:}48{:}18.555 \dashrightarrow 00{:}48{:}20.253$ at bringing in Egypt or the rapy

NOTE Confidence: 0.868247807025909

 $00:48:20.253 \longrightarrow 00:48:21.696$ earlier to those patients who

NOTE Confidence: 0.868247807025909

 $00:48:21.696 \longrightarrow 00:48:22.920$ can theoretically be cured.

NOTE Confidence: 0.868247807025909 00:48:22.920 --> 00:48:23.263 Today,

NOTE Confidence: 0.868247807025909

00:48:23.263 --> 00:48:25.321 I wanted to really focus on

NOTE Confidence: 0.868247807025909

00:48:25.321 --> 00:48:26.926 metastatic disease and the fact

NOTE Confidence: 0.868247807025909

 $00:48:26.926 \longrightarrow 00:48:28.704$ that with all that we can do,

NOTE Confidence: 0.868247807025909

 $00:48:28.710 \longrightarrow 00:48:29.978$ it still about communication

 $00:48:29.978 \longrightarrow 00:48:31.880$ and really to help our patients

NOTE Confidence: 0.858358144760132

 $00:48:31.941 \longrightarrow 00:48:33.411$ make the right choices for

NOTE Confidence: 0.858358144760132

 $00:48:33.411 \longrightarrow 00:48:34.587$ themselves and to live.

NOTE Confidence: 0.858358144760132

 $00:48:34.590 \longrightarrow 00:48:35.810$ Despite facing the uncertainties

NOTE Confidence: 0.858358144760132

 $00:48:35.810 \longrightarrow 00:48:37.338$ of cancer, so thank you.

NOTE Confidence: 0.700433671474457

00:48:43.290 --> 00:48:45.220 Oh, you're on mute Roy.

NOTE Confidence: 0.819496035575867

 $00:48:45.940 \longrightarrow 00:48:46.963$ Thanks thanks Heather.

NOTE Confidence: 0.819496035575867

 $00{:}48{:}46.963 \dashrightarrow 00{:}48{:}49.630$ That was wonderful and really great to see.

NOTE Confidence: 0.819496035575867

 $00:48:49.630 \longrightarrow 00:48:51.640$ The progress made in 15 years.

NOTE Confidence: 0.819496035575867

00:48:51.640 --> 00:48:54.326 You know a lot of it, you know,

NOTE Confidence: 0.819496035575867

 $00:48:54.326 \longrightarrow 00:48:56.692$ stimulated by the work of John and

NOTE Confidence: 0.819496035575867

 $00{:}48{:}56.692 \dashrightarrow 00{:}48{:}59.133$ others were open for questions now I

NOTE Confidence: 0.819496035575867

 $00{:}48{:}59.133 \dashrightarrow 00{:}49{:}01.688$ guess we do questions in the chat room.

NOTE Confidence: 0.819496035575867

 $00:49:01.690 \longrightarrow 00:49:03.030$ I see Charlie here.

NOTE Confidence: 0.819496035575867

 $00:49:03.030 \longrightarrow 00:49:05.040$ Maybe we'll start with a few.

00:49:05.040 --> 00:49:07.133 Well, while you get while they come

NOTE Confidence: 0.819496035575867

 $00{:}49{:}07.133 \dashrightarrow 00{:}49{:}09.730$ in for the EGFR mutated lung cancer.

NOTE Confidence: 0.819496035575867

00:49:09.730 --> 00:49:13.080 Of course you know you see a lot of it,

NOTE Confidence: 0.819496035575867

 $00:49:13.080 \longrightarrow 00:49:14.885$ especially where you live because

NOTE Confidence: 0.819496035575867

 $00:49:14.885 \longrightarrow 00:49:16.690$ it's so prominent in Asians.

NOTE Confidence: 0.819496035575867

00:49:16.690 --> 00:49:19.106 Do we know yet why has anyone looked

NOTE Confidence: 0.819496035575867

 $00:49:19.106 \longrightarrow 00:49:21.896$ at the genetics of that to understand

NOTE Confidence: 0.819496035575867

 $00:49:21.896 \longrightarrow 00:49:24.476$ no other certain sequences or or

NOTE Confidence: 0.819496035575867

 $00:49:24.476 \longrightarrow 00:49:26.840$ genetics that can explain that? So

NOTE Confidence: 0.810994863510132

 $00:49:26.840 \longrightarrow 00:49:28.348$ it's a great question,

NOTE Confidence: 0.810994863510132

 $00{:}49{:}28.348 \dashrightarrow 00{:}49{:}30.610$ so there are multiple efforts on

NOTE Confidence: 0.810994863510132

 $00{:}49{:}30.688 \to 00{:}49{:}33.229$ going to try to answer that question.

NOTE Confidence: 0.810994863510132

 $00:49:33.230 \longrightarrow 00:49:35.378$ So my last international trip before

NOTE Confidence: 0.810994863510132

 $00:49:35.378 \longrightarrow 00:49:37.370$ travel ended was last January.

NOTE Confidence: 0.810994863510132

 $00:49:37.370 \longrightarrow 00:49:39.250$ I was in Taipei, Taiwan.

NOTE Confidence: 0.810994863510132

00:49:39.250 --> 00:49:42.386 James Gang had invited me to the National

 $00:49:42.386 \longrightarrow 00:49:44.509$ Taiwan University and there are some.

NOTE Confidence: 0.810994863510132

 $00:49:44.510 \longrightarrow 00:49:46.664$ There's amazing work being done there

NOTE Confidence: 0.810994863510132

 $00:49:46.664 \longrightarrow 00:49:48.951$ in in multiple other Asian countries

NOTE Confidence: 0.810994863510132

 $00:49:48.951 \longrightarrow 00:49:51.695$ trying to look at the nuances subtleties.

NOTE Confidence: 0.810994863510132

 $00:49:51.700 \longrightarrow 00:49:54.628$ They've done a lot of screening as well.

NOTE Confidence: 0.810994863510132

 $00:49:54.630 \longrightarrow 00:49:56.790$ That involves screening and non smokers

NOTE Confidence: 0.810994863510132

00:49:56.790 --> 00:49:59.388 trying to figure out you know what

NOTE Confidence: 0.810994863510132

 $00:49:59.388 \longrightarrow 00:50:01.218$ what's happening and they've very

NOTE Confidence: 0.810994863510132

 $00:50:01.218 \longrightarrow 00:50:03.090$ clearly shown as significant increase

NOTE Confidence: 0.810994863510132

 $00:50:03.090 \longrightarrow 00:50:05.610$ in a denocarcinoma of the lung in non

NOTE Confidence: 0.810994863510132

 $00:50:05.610 \longrightarrow 00:50:08.895$ smokers and it's both in men and in women.

NOTE Confidence: 0.810994863510132

 $00:50:08.900 \longrightarrow 00:50:10.886$ I'm actually working with the group

NOTE Confidence: 0.810994863510132

 $00{:}50{:}10.886 \to 00{:}50{:}12.930$ that's from UC Davis University,

NOTE Confidence: 0.810994863510132

00:50:12.930 --> 00:50:14.760 California Davis University of California,

NOTE Confidence: 0.810994863510132

 $00:50:14.760 \longrightarrow 00:50:16.590$ San Francisco and then Stanford.

00:50:16.590 --> 00:50:19.398 We have a narrow one that's finding a

NOTE Confidence: 0.810994863510132

 $00{:}50{:}19.398 \to 00{:}50{:}21.790$ case control study that we're doing.

NOTE Confidence: 0.810994863510132

 $00:50:21.790 \longrightarrow 00:50:23.590$ Looking at Asian never smoking women

NOTE Confidence: 0.810994863510132

 $00:50:23.590 \longrightarrow 00:50:25.200$ who have developed lung cancer,

NOTE Confidence: 0.810994863510132

 $00:50:25.200 \longrightarrow 00:50:27.489$ almost all of whom have EGFR mutants

NOTE Confidence: 0.810994863510132

 $00:50:27.489 \longrightarrow 00:50:29.826$ mutations and then doing a match control

NOTE Confidence: 0.810994863510132

 $00:50:29.826 \longrightarrow 00:50:32.085$ of very similar people who don't have

NOTE Confidence: 0.810994863510132

00:50:32.085 --> 00:50:34.229 lung cancer and trying to see if what

NOTE Confidence: 0.810994863510132

 $00:50:34.229 \longrightarrow 00:50:36.724$ we are able to find there and that of

NOTE Confidence: 0.810994863510132

 $00:50:36.724 \longrightarrow 00:50:38.838$ course involves sample collections as well.

NOTE Confidence: 0.810994863510132

 $00{:}50{:}38.840 --> 00{:}50{:}40.700$ And there are a lot of

NOTE Confidence: 0.810994863510132

 $00:50:40.700 \longrightarrow 00:50:41.630$ similar ongoing efforts,

NOTE Confidence: 0.810994863510132

00:50:41.630 --> 00:50:43.702 but we don't know an for me here

NOTE Confidence: 0.810994863510132

00:50:43.702 --> 00:50:45.349 in Northern California though,

NOTE Confidence: 0.810994863510132

 $00:50:45.350 \longrightarrow 00:50:47.950$ a lot of my patience with you from you and

NOTE Confidence: 0.810994863510132

 $00:50:48.016 \dashrightarrow 00:50:50:620$ lung cancer are never smoking Asian women.

 $00:50:50.620 \longrightarrow 00:50:52.797$ I also have a whole lot of.

NOTE Confidence: 0.810994863510132

 $00:50:52.800 \longrightarrow 00:50:54.230$ People have multiple different ethnicities,

NOTE Confidence: 0.810994863510132

 $00:50:54.230 \longrightarrow 00:50:55.088$ men and women.

NOTE Confidence: 0.810994863510132

 $00:50:55.088 \longrightarrow 00:50:56.804$ Different age ranges who also have

NOTE Confidence: 0.810994863510132

00:50:56.804 --> 00:50:58.790 each of our mutant lung cancer show.

NOTE Confidence: 0.810994863510132

00:50:58.790 --> 00:51:00.210 It's not just that group,

NOTE Confidence: 0.810994863510132

00:51:00.210 --> 00:51:01.938 but there's gotta be something 'cause

NOTE Confidence: 0.810994863510132

 $00{:}51{:}01.938 \dashrightarrow 00{:}51{:}04.283$ it does seem to be increasing and were

NOTE Confidence: 0.810994863510132

 $00:51:04.283 \longrightarrow 00:51:06.200$ still baffled as to what that is.

NOTE Confidence: 0.788463294506073

 $00{:}51{:}06.750 \dashrightarrow 00{:}51{:}08.556$ Great, thanks for the next question.

NOTE Confidence: 0.788463294506073

 $00{:}51{:}08.560 \dashrightarrow 00{:}51{:}10.457$ We have a special guest 15 years

NOTE Confidence: 0.788463294506073

 $00:51:10.457 \longrightarrow 00:51:11.995$ ago a young Scott Guettinger

NOTE Confidence: 0.788463294506073

 $00{:}51{:}11.995 \dashrightarrow 00{:}51{:}14.298$ came to Yale to work with Sean.

NOTE Confidence: 0.788463294506073

 $00{:}51{:}14.300 \dashrightarrow 00{:}51{:}15.810$ Sadly they only work together

NOTE Confidence: 0.788463294506073

 $00:51:15.810 \longrightarrow 00:51:17.320$ for know about a year,

00:51:17.320 --> 00:51:19.126 but if Scott turned out OK,

NOTE Confidence: 0.788463294506073

 $00{:}51{:}19.130 --> 00{:}51{:}20.640$ Jonathan done OK with you.

NOTE Confidence: 0.788463294506073

 $00:51:20.640 \longrightarrow 00:51:23.260$ So Scott, I'm going to move over so I put

NOTE Confidence: 0.788463294506073

 $00:51:23.328 \longrightarrow 00:51:26.080$ my mask on Scotts gonna ask alive question.

NOTE Confidence: 0.788463294506073

 $00:51:26.080 \longrightarrow 00:51:28.510$ These are Dart leader in lung cancer.

NOTE Confidence: 0.788463294506073

 $00:51:28.510 \longrightarrow 00:51:31.200$ To see you. Very talk I'm.

NOTE Confidence: 0.788463294506073

00:51:31.200 --> 00:51:33.855 I don't think you got into it too much,

NOTE Confidence: 0.788463294506073

 $00:51:33.860 \longrightarrow 00:51:35.172$ but I'm just curious.

NOTE Confidence: 0.788463294506073

 $00:51:35.172 \longrightarrow 00:51:37.640$ For your patience to progress at multiple

NOTE Confidence: 0.788463294506073

00:51:37.640 --> 00:51:39.815 sites while getting closer mitted,

NOTE Confidence: 0.788463294506073

 $00:51:39.820 \longrightarrow 00:51:42.588$ do you continue who are not trial candidates?

NOTE Confidence: 0.788463294506073

00:51:42.590 --> 00:51:44.840 Do you continue the those fermented

NOTE Confidence: 0.788463294506073

 $00:51:44.840 \longrightarrow 00:51:47.028$ with the chemotherapy to stop it

NOTE Confidence: 0.788463294506073

 $00:51:47.028 \longrightarrow 00:51:49.212$ along the lines of the impress trial

NOTE Confidence: 0.788463294506073

 $00:51:49.212 \longrightarrow 00:51:51.615$ where we didn't see a benefit from

NOTE Confidence: 0.788463294506073

 $00:51:51.615 \longrightarrow 00:51:53.662$ at least first generation TK eyes?

00:51:53.662 --> 00:51:55.738 How do you handle that situation?

NOTE Confidence: 0.802918910980225

 $00:51:55.740 \longrightarrow 00:51:57.798$ That's a great question, so obviously

NOTE Confidence: 0.802918910980225

 $00:51:57.798 \longrightarrow 00:51:59.540$ it's different for different folks.

NOTE Confidence: 0.802918910980225

 $00:51:59.540 \longrightarrow 00:52:01.969$ So first answer the first part of

NOTE Confidence: 0.802918910980225

 $00{:}52{:}01.969 \dashrightarrow 00{:}52{:}04.099$ question you didn't actually ask which

NOTE Confidence: 0.802918910980225

00:52:04.099 --> 00:52:06.879 is if someone is progressing in a single

NOTE Confidence: 0.802918910980225

 $00:52:06.879 \longrightarrow 00:52:09.255$ or a couple areas, we will continue.

NOTE Confidence: 0.802918910980225

 $00{:}52{:}09.255 \dashrightarrow 00{:}52{:}10.675$ Oh Smyrna bandu radiation.

NOTE Confidence: 0.802918910980225

 $00:52:10.680 \longrightarrow 00:52:12.112$ Or other local ablative

NOTE Confidence: 0.802918910980225

00:52:12.112 --> 00:52:13.186 therapy whenever possible,

NOTE Confidence: 0.802918910980225

 $00:52:13.190 \longrightarrow 00:52:15.110$ and try to extend the time

NOTE Confidence: 0.802918910980225

 $00:52:15.110 \longrightarrow 00:52:17.140$ that they're on that treatment.

NOTE Confidence: 0.802918910980225

 $00:52:17.140 \longrightarrow 00:52:19.796$ If we get to a point where they

NOTE Confidence: 0.802918910980225

 $00{:}52{:}19.796 \dashrightarrow 00{:}52{:}21.890$ are definitely needing to switch

NOTE Confidence: 0.802918910980225

 $00:52:21.890 \longrightarrow 00:52:24.145$ to a different systemic therapy.

00:52:24.150 --> 00:52:25.920 Looking for trials first and foremost,

NOTE Confidence: 0.802918910980225

 $00{:}52{:}25.920 \dashrightarrow 00{:}52{:}27.894$ but if we can't and they're going

NOTE Confidence: 0.802918910980225

 $00:52:27.894 \longrightarrow 00:52:30.610$ to switch to chemo, not on a trial.

NOTE Confidence: 0.802918910980225

 $00:52:30.610 \longrightarrow 00:52:32.770$ It will depend on the patient.

NOTE Confidence: 0.802918910980225

 $00:52:32.770 \longrightarrow 00:52:34.280$ If they have significant amount

NOTE Confidence: 0.802918910980225

 $00.52:34.280 \longrightarrow 00.52:35.186$ of brain disease,

NOTE Confidence: 0.802918910980225

 $00:52:35.190 \longrightarrow 00:52:37.038$ I will continue the OCE myrnam and

NOTE Confidence: 0.802918910980225

 $00:52:37.038 \longrightarrow 00:52:38.830$ start them on the chemotherapy.

NOTE Confidence: 0.802918910980225

00:52:38.830 --> 00:52:40.038 Continuing osimertinib for some

NOTE Confidence: 0.802918910980225

 $00:52:40.038 \longrightarrow 00:52:41.246$ of the other patients.

NOTE Confidence: 0.802918910980225

 $00:52:41.250 \longrightarrow 00:52:43.887$ I will stop it and part of my rationale

NOTE Confidence: 0.802918910980225

 $00:52:43.887 \longrightarrow 00:52:46.097$ behind that is I watch them closely.

NOTE Confidence: 0.802918910980225

00:52:46.100 --> 00:52:48.214 I do believe that flare can happen,

NOTE Confidence: 0.802918910980225

 $00:52:48.220 \longrightarrow 00:52:49.356$ but it's pretty rare.

NOTE Confidence: 0.802918910980225

00:52:49.356 --> 00:52:51.437 So if a patient suddenly gets drastically

NOTE Confidence: 0.802918910980225

00:52:51.437 --> 00:52:53.677 worse when we stop the jiffi drug,

 $00:52:53.680 \longrightarrow 00:52:54.888$ their back on it,

NOTE Confidence: 0.802918910980225

 $00:52:54.888 \longrightarrow 00:52:56.096$ they stay on it.

NOTE Confidence: 0.802918910980225

00:52:56.100 --> 00:52:58.102 I've had that happen more in people

NOTE Confidence: 0.802918910980225

 $00:52:58.102 \longrightarrow 00:53:00.040$ with bone metastases and interesting Lee.

NOTE Confidence: 0.802918910980225

 $00:53:00.040 \longrightarrow 00:53:02.812$ I think if you read when breath becomes air.

NOTE Confidence: 0.802918910980225

 $00:53:02.820 \longrightarrow 00:53:04.708$ Paul was one of the people I continued

NOTE Confidence: 0.802918910980225

00:53:04.708 --> 00:53:06.600 on and he never got to Summerton,

NOTE Confidence: 0.802918910980225

00:53:06.600 --> 00:53:08.322 but I can't because he was his

NOTE Confidence: 0.802918910980225

 $00:53:08.322 \longrightarrow 00:53:09.370$ disease was before that,

NOTE Confidence: 0.802918910980225

00:53:09.370 --> 00:53:11.330 but I continued him on 'cause he was

NOTE Confidence: 0.802918910980225

00:53:11.330 --> 00:53:13.127 someone who as soon as we stopped he

NOTE Confidence: 0.802918910980225

00:53:13.127 --> 00:53:14.891 had bone pain so he was definitely

NOTE Confidence: 0.802918910980225

 $00:53:14.891 \longrightarrow 00:53:16.928$ someone who flared so he never stopped.

NOTE Confidence: 0.802918910980225

 $00:53:16.930 \longrightarrow 00:53:19.464$ But for other patients I will stop.

NOTE Confidence: 0.802918910980225

 $00:53:19.470 \longrightarrow 00:53:21.718$ And part of the rationale is that by

 $00:53:21.718 \longrightarrow 00:53:23.695$ stopping the EGFR TK on the patients

NOTE Confidence: 0.802918910980225

 $00:53:23.695 \longrightarrow 00:53:26.139$ where you can and doing with chemotherapy,

NOTE Confidence: 0.802918910980225

 $00:53:26.140 \longrightarrow 00:53:28.066$ the resistance that develops at that

NOTE Confidence: 0.802918910980225

00:53:28.066 --> 00:53:29.954 point after they've been on chemo

NOTE Confidence: 0.802918910980225

 $00:53:29.954 \longrightarrow 00:53:31.872$ for awhile tends to be much more

NOTE Confidence: 0.802918910980225

 $00:53:31.872 \longrightarrow 00:53:33.410$ responsive to restarting and EGFR.

NOTE Confidence: 0.802918910980225

 $00:53:33.410 \longrightarrow 00:53:35.394$ TK I in the context of a study

NOTE Confidence: 0.802918910980225

 $00:53:35.394 \longrightarrow 00:53:36.740$ or some combination,

NOTE Confidence: 0.802918910980225

00:53:36.740 --> 00:53:38.854 so that's while I'll think about it,

NOTE Confidence: 0.802918910980225

 $00:53:38.860 \longrightarrow 00:53:40.972$ but a lot of my longer term patients

NOTE Confidence: 0.802918910980225

00:53:40.972 --> 00:53:43.217 right now we don't stop because they've

NOTE Confidence: 0.802918910980225

 $00:53:43.217 \longrightarrow 00:53:45.230$ had things where they've had bone,

NOTE Confidence: 0.802918910980225

 $00:53:45.230 \longrightarrow 00:53:46.740$ Mets, or other problems so.

NOTE Confidence: 0.89730766415596

 $00{:}53{:}47.540 {\:{\circ}{\circ}{\circ}}>00{:}53{:}50.816$ Thank you yeah. Great thanks Scott.

NOTE Confidence: 0.89730766415596

 $00:53:50.820 \longrightarrow 00:53:54.726$ We have a few questions in the chat room.

NOTE Confidence: 0.89730766415596

00:53:54.730 --> 00:53:58.762 Um? It's from Sarah Goldberg.

00:53:58.762 --> 00:54:01.570 Hi Sarah, was with us last night.

NOTE Confidence: 0.89730766415596

00:54:01.570 --> 00:54:03.665 You see any promising avenues

NOTE Confidence: 0.89730766415596

00:54:03.665 --> 00:54:05.760 for using immunotherapy in each

NOTE Confidence: 0.89730766415596

 $00:54:05.833 \longrightarrow 00:54:07.933$ year from you in lung cancer

NOTE Confidence: 0.89730766415596

00:54:07.933 --> 00:54:10.840 patients beyond PD one agents? Oh,

NOTE Confidence: 0.829002901911736

 $00:54:10.840 \longrightarrow 00:54:12.433$ great question Sarah.

NOTE Confidence: 0.829002901911736

 $00:54:12.433 \longrightarrow 00:54:17.019$ So I'll again into the first part of that.

NOTE Confidence: 0.829002901911736

00:54:17.020 --> 00:54:18.625 So with the checkpoint inhibitors

NOTE Confidence: 0.829002901911736

 $00{:}54{:}18.625 \longrightarrow 00{:}54{:}20.930$ I've had a couple of patients who

NOTE Confidence: 0.829002901911736

 $00:54:20.930 \longrightarrow 00:54:22.844$ have done very well with them.

NOTE Confidence: 0.829002901911736

 $00:54:22.850 \longrightarrow 00:54:24.590$ They were patients tending to

NOTE Confidence: 0.829002901911736

 $00:54:24.590 \longrightarrow 00:54:27.136$ VL 858 are not deletion 19 and

NOTE Confidence: 0.829002901911736

 $00{:}54{:}27.136 \dashrightarrow 00{:}54{:}28.966$ that's something that in Taiwan.

NOTE Confidence: 0.829002901911736

 $00:54:28.970 \longrightarrow 00:54:30.434$ They've been looking at.

NOTE Confidence: 0.829002901911736

00:54:30.434 --> 00:54:33.290 A lot also tended to have high PD,

 $00:54:33.290 \longrightarrow 00:54:35.738$ L1, and so a couple folks

NOTE Confidence: 0.829002901911736

 $00:54:35.738 \longrightarrow 00:54:38.120$ have done well and we are.

NOTE Confidence: 0.829002901911736

00:54:38.120 --> 00:54:40.340 Participating in a study looking

NOTE Confidence: 0.829002901911736

 $00:54:40.340 \longrightarrow 00:54:42.116$ at chemo with carboplatin.

NOTE Confidence: 0.829002901911736

 $00:54:42.120 \longrightarrow 00:54:43.449$ Pemetrexed said bevacizumab

NOTE Confidence: 0.829002901911736

 $00:54:43.449 \longrightarrow 00:54:45.664$ plus or minus a Tezos.

NOTE Confidence: 0.829002901911736

 $00:54:45.670 \longrightarrow 00:54:48.880$ This is a multi site study.

NOTE Confidence: 0.829002901911736

 $00:54:48.880 \longrightarrow 00:54:51.656$ So I think there might be room there,

NOTE Confidence: 0.829002901911736

00:54:51.660 --> 00:54:53.430 but it's not something I'm doing

NOTE Confidence: 0.829002901911736

 $00:54:53.430 \longrightarrow 00:54:56.377$ is often off trial as far as other

NOTE Confidence: 0.829002901911736

 $00{:}54{:}56.377 \dashrightarrow 00{:}54{:}57.573$ immunomodulatory modulators, yes,

NOTE Confidence: 0.829002901911736

 $00{:}54{:}57.573 \dashrightarrow 00{:}55{:}00.317$ but I'm not a good enough immunologist to

NOTE Confidence: 0.829002901911736

 $00:55:00.317 \longrightarrow 00:55:03.150$ tell you exactly what that's going to be.

NOTE Confidence: 0.829002901911736

00:55:03.150 --> 00:55:04.908 I think that it's something that

NOTE Confidence: 0.829002901911736

 $00:55:04.908 \longrightarrow 00:55:07.263$ a lot of us are watching really

NOTE Confidence: 0.829002901911736

00:55:07.263 --> 00:55:09.867 closely to try to understand what is

 $00:55:09.943 \longrightarrow 00:55:12.540$ it that the simple explanation is EGFR?

NOTE Confidence: 0.829002901911736

 $00{:}55{:}12.540 \dashrightarrow 00{:}55{:}13.932$ Mutant lung cancer doesn't

NOTE Confidence: 0.829002901911736

 $00:55:13.932 \longrightarrow 00:55:15.672$ have that many new antigens,

NOTE Confidence: 0.829002901911736

00:55:15.680 --> 00:55:17.756 'cause it's not very complex tumor,

NOTE Confidence: 0.829002901911736 $00:55:17.760 \longrightarrow 00:55:18.432$ but then, NOTE Confidence: 0.829002901911736 $00:55:18.432 \longrightarrow 00:55:18.768$ overtime.

00:55:18.768 --> 00:55:20.448 Theoretically it gets more complex,

NOTE Confidence: 0.829002901911736

NOTE Confidence: 0.829002901911736

 $00{:}55{:}20.450 \dashrightarrow 00{:}55{:}22.172$ so there should be more NEO

NOTE Confidence: 0.829002901911736

 $00:55:22.172 \longrightarrow 00:55:23.880$ antigens but really knew antigens.

NOTE Confidence: 0.829002901911736

 $00:55:23.880 \longrightarrow 00:55:26.064$ Not all Neo antigens are the same,

NOTE Confidence: 0.829002901911736

00:55:26.070 --> 00:55:28.392 and so I think that if we can figure

NOTE Confidence: 0.829002901911736

 $00:55:28.392 \longrightarrow 00:55:31.166$ out what is it that can make that

NOTE Confidence: 0.829002901911736

 $00:55:31.166 \longrightarrow 00:55:32.929$ cancer more identifiable to immune,

NOTE Confidence: 0.829002901911736

 $00:55:32.930 \longrightarrow 00:55:35.107$ system will have ways to move forward.

NOTE Confidence: 0.829002901911736

00:55:35.110 --> 00:55:36.982 But I don't think we've figured

 $00:55:36.982 \longrightarrow 00:55:38.550$ that out very well yet,

NOTE Confidence: 0.829002901911736

 $00:55:38.550 \longrightarrow 00:55:38.860$ so.

NOTE Confidence: 0.74587082862854

 $00:55:40.690 \longrightarrow 00:55:43.510$ Great, we have one from Katie

NOTE Confidence: 0.74587082862854

00:55:43.510 --> 00:55:46.239 Palitti who you know as well.

NOTE Confidence: 0.74587082862854

00:55:46.240 --> 00:55:48.304 When I don't know if we can have

NOTE Confidence: 0.74587082862854

00:55:48.304 --> 00:55:50.087 Katie give it live or question,

NOTE Confidence: 0.74587082862854

 $00:55:50.090 \longrightarrow 00:55:52.015$ I don't know how to do that,

NOTE Confidence: 0.74587082862854

 $00.55.52.020 \longrightarrow 00.55.53.390$ so I'll just ask it.

NOTE Confidence: 0.74587082862854

 $00{:}55{:}53.390 \dashrightarrow 00{:}55{:}54.896$ She, she says she's only at

NOTE Confidence: 0.74587082862854

 $00:55:54.896 \longrightarrow 00:55:56.560$ one of the other leaders of

NOTE Confidence: 0.74587082862854

 $00{:}55{:}56.560 \dashrightarrow 00{:}55{:}58.065$ our sport program in EGFR.

NOTE Confidence: 0.74587082862854

 $00{:}55{:}58.070 \dashrightarrow 00{:}55{:}59.988$ Resistance does thank you for great lecture.

NOTE Confidence: 0.74587082862854

00:55:59.990 --> 00:56:00.665 Heather grey talk.

NOTE Confidence: 0.74587082862854

 $00{:}56{:}00.665 \dashrightarrow 00{:}56{:}02.015$ I was wondering what you think

NOTE Confidence: 0.74587082862854

 $00:56:02.015 \longrightarrow 00:56:03.748$ we should be thinking about to

NOTE Confidence: 0.74587082862854

 $00:56:03.748 \longrightarrow 00:56:04.912$ address the heterogeneity of

 $00{:}56{:}04.912 \dashrightarrow 00{:}56{:}06.177$ resistance mechanisms that can

NOTE Confidence: 0.74587082862854

 $00{:}56{:}06.177 \dashrightarrow 00{:}56{:}07.417$ emerge in individual patients.

NOTE Confidence: 0.841703712940216

 $00:56:08.070 \longrightarrow 00:56:09.111$ Great question Katie.

NOTE Confidence: 0.841703712940216

00:56:09.111 --> 00:56:12.331 So I think that as much as the targeted

NOTE Confidence: 0.841703712940216

 $00:56:12.331 \longrightarrow 00:56:14.816$ story is a beautiful story and to

NOTE Confidence: 0.841703712940216

 $00:56:14.816 \longrightarrow 00:56:17.421$ be able to say we found the deletion

NOTE Confidence: 0.841703712940216

 $00:56:17.421 \longrightarrow 00:56:19.805$ 19 and we treat it with their lot

NOTE Confidence: 0.841703712940216

 $00{:}56{:}19.805 \dashrightarrow 00{:}56{:}22.121$ nib and then the T790M developed and

NOTE Confidence: 0.841703712940216

 $00:56:22.121 \longrightarrow 00:56:24.527$ then we treated with those summerton.

NOTE Confidence: 0.841703712940216

 $00:56:24.530 \longrightarrow 00:56:26.933$ If that only gets a so far and I

NOTE Confidence: 0.841703712940216

 $00:56:26.933 \longrightarrow 00:56:28.934$ don't think we're ever going to

NOTE Confidence: 0.841703712940216

 $00:56:28.934 \longrightarrow 00:56:31.077$ be able to really get completely

NOTE Confidence: 0.841703712940216

 $00{:}56{:}31.077 \dashrightarrow 00{:}56{:}33.982$ around that because as time goes on

NOTE Confidence: 0.841703712940216

 $00:56:33.982 \longrightarrow 00:56:35.852$ there's more and more heterogeneity

NOTE Confidence: 0.841703712940216

00:56:35.852 --> 00:56:37.228 in the resistance pathways.

 $00:56:37.230 \longrightarrow 00:56:38.940$ Should we have to be?

NOTE Confidence: 0.841703712940216

00:56:38.940 --> 00:56:39.726 Thinking about it,

NOTE Confidence: 0.841703712940216

 $00:56:39.726 \longrightarrow 00:56:40.774$ you know the straightforward

NOTE Confidence: 0.841703712940216

 $00:56:40.774 \longrightarrow 00:56:41.560$ answer right now.

NOTE Confidence: 0.841703712940216

 $00:56:41.560 \longrightarrow 00:56:42.580$ Is chemotherapy 'cause that

NOTE Confidence: 0.841703712940216

 $00{:}56{:}42.580 \dashrightarrow 00{:}56{:}44.110$ doesn't really care so much about

NOTE Confidence: 0.841703712940216

 $00:56:44.152 \longrightarrow 00:56:45.487$ what the resistance pathway was,

NOTE Confidence: 0.841703712940216

 $00:56:45.490 \longrightarrow 00:56:47.597$ but it gets back to Sarah's question

NOTE Confidence: 0.841703712940216

 $00{:}56{:}47.597 \dashrightarrow 00{:}56{:}49.278$ that eventually we've got to figure

NOTE Confidence: 0.841703712940216

00:56:49.278 --> 00:56:50.951 out how to get the immune system

NOTE Confidence: 0.841703712940216

 $00{:}56{:}51.008 \dashrightarrow 00{:}56{:}52.646$ to be able to recognize the cancer

NOTE Confidence: 0.841703712940216

 $00:56:52.646 \longrightarrow 00:56:54.181$ and all of its various iterations.

NOTE Confidence: 0.841703712940216

 $00:56:54.181 \longrightarrow 00:56:55.566$ As it becomes more resistant

NOTE Confidence: 0.841703712940216

 $00:56:55.566 \longrightarrow 00:56:57.017$ and be able to tackle it.

NOTE Confidence: 0.841703712940216

 $00:56:57.020 \longrightarrow 00:56:58.889$ I don't have an answer as to

NOTE Confidence: 0.841703712940216

 $00:56:58.889 \longrightarrow 00:57:00.418$ how we're going to do that,

 $00:57:00.420 \longrightarrow 00:57:02.373$ but to me that's really the only

NOTE Confidence: 0.841703712940216

 $00:57:02.373 \longrightarrow 00:57:04.348$ way we're really going to get at it,

NOTE Confidence: 0.841703712940216

 $00:57:04.350 \longrightarrow 00:57:05.868$ because were to get it early

NOTE Confidence: 0.841703712940216

 $00:57:05.868 \longrightarrow 00:57:07.736$ enough that it hasn't had time to

NOTE Confidence: 0.841703712940216

 $00:57:07.736 \longrightarrow 00:57:09.066$ develop all of the resistance,

NOTE Confidence: 0.841703712940216

 $00:57:09.070 \longrightarrow 00:57:10.194$ and that's where the.

NOTE Confidence: 0.841703712940216

00:57:10.194 --> 00:57:11.880 Earlier stage therapy is helpful so,

NOTE Confidence: 0.841703712940216

 $00:57:11.880 \longrightarrow 00:57:13.936$ but even there I think that I don't

NOTE Confidence: 0.841703712940216

 $00:57:13.936 \longrightarrow 00:57:16.059$ know that we're going to cure with

NOTE Confidence: 0.841703712940216

 $00{:}57{:}16.059 \dashrightarrow 00{:}57{:}17.960$ that approach because I think that

NOTE Confidence: 0.841703712940216

 $00:57:17.960 \longrightarrow 00:57:20.336$ it only takes a few cells that become

NOTE Confidence: 0.841703712940216

 $00:57:20.336 \longrightarrow 00:57:22.100$ resistant to then grow up again,

NOTE Confidence: 0.841703712940216

 $00{:}57{:}22.100 \dashrightarrow 00{:}57{:}24.660$ and so I think we've all got a lot of

NOTE Confidence: 0.841703712940216

 $00:57:24.730 \longrightarrow 00:57:27.162$ years left of work that we need to

NOTE Confidence: 0.841703712940216

 $00:57:27.162 \longrightarrow 00:57:29.986$ get doing to be able to make the impacts.

 $00:57:29.990 \longrightarrow 00:57:31.150$ We'd like to make.

NOTE Confidence: 0.847847104072571

 $00:57:31.750 \longrightarrow 00:57:33.690$ Right, and certainly by working

NOTE Confidence: 0.847847104072571

 $00:57:33.690 \longrightarrow 00:57:35.630$ through the cooperative groups and

NOTE Confidence: 0.847847104072571

 $00:57:35.689 \longrightarrow 00:57:37.569$ the societies and collaborating.

NOTE Confidence: 0.847847104072571

 $00:57:37.570 \longrightarrow 00:57:40.066$ Given that this is a worldwide

NOTE Confidence: 0.847847104072571

 $00.57:40.066 \longrightarrow 00.57:41.314$ disease so important.

NOTE Confidence: 0.847847104072571

 $00:57:41.320 \longrightarrow 00:57:43.120$ We I don't see too many

NOTE Confidence: 0.847847104072571

 $00:57:43.120 \longrightarrow 00:57:44.730$ more questions in the chat,

NOTE Confidence: 0.847847104072571

 $00:57:44.730 \longrightarrow 00:57:47.097$ just about the end of the hour I see

NOTE Confidence: 0.847847104072571

00:57:47.097 --> 00:57:49.502 Nancy nearing joined and I just want

NOTE Confidence: 0.847847104072571

 $00{:}57{:}49.502 \dashrightarrow 00{:}57{:}51.872$ to acknowledge her Johns wife and I'm

NOTE Confidence: 0.847847104072571

00:57:51.872 --> 00:57:54.021 sorry we can't talk in person Nancy,

NOTE Confidence: 0.847847104072571

00:57:54.030 --> 00:57:55.992 but next year hopefully will see

NOTE Confidence: 0.847847104072571

 $00:57:55.992 \longrightarrow 00:57:58.368$ you in the front row and we can.

NOTE Confidence: 0.847847104072571

00:57:58.370 --> 00:58:00.926 Zoomed Heather inferred picture but Curly

NOTE Confidence: 0.847847104072571

 $00:58:00.926 \longrightarrow 00:58:03.639$ any any final thoughts before we end?

 $00{:}58{:}03.640 \dashrightarrow 00{:}58{:}06.070$ No, there was a fantastic talk.

NOTE Confidence: 0.847847104072571

 $00.58:06.070 \longrightarrow 00.58:08.380$ I mean, get really outlines the

NOTE Confidence: 0.847847104072571

 $00:58:08.380 \longrightarrow 00:58:10.930$ but so exciting in lung cancer,

NOTE Confidence: 0.847847104072571

 $00:58:10.930 \longrightarrow 00:58:12.975$ both with respect to targeted

NOTE Confidence: 0.847847104072571

 $00:58:12.975 \longrightarrow 00:58:14.611$ therapies an hopefully integrating

NOTE Confidence: 0.847847104072571

 $00:58:14.611 \longrightarrow 00:58:16.190$ IO with these things.

NOTE Confidence: 0.847847104072571 00:58:16.190 --> 00:58:16.706 Overtime. NOTE Confidence: 0.847847104072571

 $00:58:16.706 \longrightarrow 00:58:20.318$ So thank you is really powerful work.

NOTE Confidence: 0.847847104072571

 $00:58:20.320 \longrightarrow 00:58:22.330$ And thanks for taking the time.

NOTE Confidence: 0.847847104072571

 $00:58:22.330 \longrightarrow 00:58:24.010$ Thanks to everyone who joined.

NOTE Confidence: 0.847847104072571

00.58:24.010 --> 00.58:26.348 We had a very large crowd today,

NOTE Confidence: 0.847847104072571

 $00:58:26.350 \longrightarrow 00:58:28.674$ both because of you Heather and because

NOTE Confidence: 0.847847104072571

 $00{:}58{:}28.674 \dashrightarrow 00{:}58{:}31.152$ of John and we remember him fondly

NOTE Confidence: 0.847847104072571

 $00:58:31.152 \longrightarrow 00:58:33.639$ and in his memory will continue to

NOTE Confidence: 0.847847104072571

 $00:58:33.639 \longrightarrow 00:58:35.781$ take good care of our patients and

 $00:58:35.781 \longrightarrow 00:58:37.740$ which was always number one goal

NOTE Confidence: 0.847847104072571

 $00:58:37.740 \longrightarrow 00:58:39.415$ with the most innovative therapies.

NOTE Confidence: 0.847847104072571

 $00:58:39.420 \longrightarrow 00:58:41.430$ Which is exactly what both are.

NOTE Confidence: 0.847847104072571

 $00:58:41.430 \longrightarrow 00:58:43.140$ Cancer centers are trying to

NOTE Confidence: 0.847847104072571

00:58:43.140 --> 00:58:45.110 do so thank you everyone and

NOTE Confidence: 0.838841021060944

00:58:45.110 --> 00:58:46.805 see you soon. Thank you.