WEBVTT

NOTE duration:"01:01:02"

NOTE recognizability:0.921

NOTE language:en-us

NOTE Confidence: 0.953192257142857

00:00:00.000 --> 00:00:03.038 Welcome to Yale Cancer Center Ground Rounds.

NOTE Confidence: 0.953192257142857

00:00:03.040 --> 00:00:04.394 I'm Doctor Miriam Lustberg

NOTE Confidence: 0.953192257142857

 $00{:}00{:}04{.}394 \dashrightarrow 00{:}00{:}07{.}136$ and I have the privilege of

NOTE Confidence: 0.953192257142857

00:00:07.136 --> 00:00:09.598 introducing our two speakers today.

NOTE Confidence: 0.953192257142857

 $00{:}00{:}09{.}600 \dashrightarrow 00{:}00{:}12.834$ Doctor Elena Ratner is a professor

NOTE Confidence: 0.953192257142857

 $00:00:12.834 \rightarrow 00:00:15.668$ of obstetrics and gynecology and

NOTE Confidence: 0.953192257142857

 $00:00:15.668 \rightarrow 00:00:18.618$ reproductive sciences and cochief of

NOTE Confidence: 0.953192257142857

00:00:18.618 --> 00:00:21.840 the section of Gynecological Oncology.

NOTE Confidence: 0.953192257142857

 $00:00:21.840 \rightarrow 00:00:24.510$ She's a board certified gynecologist.

NOTE Confidence: 0.953192257142857

 $00{:}00{:}24.510 \dashrightarrow 00{:}00{:}26.640$ Oncologist with special interest

NOTE Confidence: 0.953192257142857

 $00:00:26.640 \longrightarrow 00:00:27.770$ in chemotherapy,

NOTE Confidence: 0.953192257142857

 $00{:}00{:}27.770 \dashrightarrow 00{:}00{:}29.465$ Targeted drug development,

NOTE Confidence: 0.953192257142857

 $00:00:29.470 \longrightarrow 00:00:31.520$ patient quality of life programs

 $00:00:31.520 \longrightarrow 00:00:34.630$ and early cancer detection.

NOTE Confidence: 0.953192257142857

 $00{:}00{:}34.630 \dashrightarrow 00{:}00{:}36.256$ She's the current codirector

NOTE Confidence: 0.953192257142857

00:00:36.256 --> 00:00:38.868 of Discovery to Cure,

NOTE Confidence: 0.953192257142857

00:00:38.870 --> 00:00:42.176 Director of discovery to Cure Early

NOTE Confidence: 0.953192257142857

 $00{:}00{:}42.176$ --> $00{:}00{:}45.132$ ovarian cancer detection program and a

NOTE Confidence: 0.953192257142857

 $00:00:45.132 \rightarrow 00:00:47.664$ founder and director of the Sexuality,

NOTE Confidence: 0.953192257142857

00:00:47.670 --> 00:00:50.734 intimacy and Menopause program.

NOTE Confidence: 0.953192257142857

 $00:00:50.734 \longrightarrow 00:00:52.266$ Her expertise?

NOTE Confidence: 0.953192257142857

 $00:00:52.270 \rightarrow 00:00:54.880$ Is cytoreductive surgery for patients

NOTE Confidence: 0.953192257142857

 $00{:}00{:}54.880 \dashrightarrow 00{:}00{:}57.490$ with advanced ovarian cancer minimally

NOTE Confidence: 0.953192257142857

 $00:00:57.566 \longrightarrow 00:01:00.611$ invasive procedures for patients with

NOTE Confidence: 0.953192257142857

00:01:00.611 --> 00:01:03.128 gynacologic malignancies or complex

NOTE Confidence: 0.953192257142857

 $00{:}01{:}03{.}128 \dashrightarrow 00{:}01{:}05{.}724$ benign gynacologic conditions and

NOTE Confidence: 0.953192257142857

00:01:05.724 --> 00:01:08.320 robotic fertility sparing surgery

NOTE Confidence: 0.953192257142857

00:01:08.409 --> 00:01:10.989 for young patients with cancer?

NOTE Confidence: 0.953192257142857

00:01:10.990 --> 00:01:13.365 Doctor Ratner's laboratory is working

- NOTE Confidence: 0.953192257142857
- $00:01:13.365 \rightarrow 00:01:16.325$ on new targeted drugs for ovarian
- NOTE Confidence: 0.953192257142857
- $00:01:16.325 \longrightarrow 00:01:19.355$ cancer in order to provide patients
- NOTE Confidence: 0.953192257142857
- 00:01:19.355 00:01:21.330 with truly personalized care.
- NOTE Confidence: 0.953192257142857
- 00:01:21.330 --> 00:01:23.490 Her work additionally focuses on
- NOTE Confidence: 0.953192257142857
- $00:01:23.490 \rightarrow 00:01:25.350$ reversing chemotherapy resistance
- NOTE Confidence: 0.953192257142857
- $00:01:25.350 \longrightarrow 00:01:28.414$ in ovarian and uterine cancers.
- NOTE Confidence: 0.953192257142857
- 00:01:28.414 --> 00:01:31.469 Our second speaker is Doctor
- NOTE Confidence: 0.953192257142857
- 00:01:31.469 --> 00:01:32.890 Alessandro Santine.
- NOTE Confidence: 0.953192257142857
- 00:01:32.890 --> 00:01:33.310 Dr.
- NOTE Confidence: 0.953192257142857
- $00:01:33.310 \longrightarrow 00:01:36.395$ Santine is a professor of obstetrics
- NOTE Confidence: 0.953192257142857
- $00:01:36.395 \rightarrow 00:01:38.527$ and gynecology and reproductive
- NOTE Confidence: 0.953192257142857
- $00{:}01{:}38{.}527 \dashrightarrow 00{:}01{:}41{.}075$ sciences and cochief of the
- NOTE Confidence: 0.953192257142857
- 00:01:41.075 --> 00:01:43.007 section of Gynecology oncology.
- NOTE Confidence: 0.953192257142857
- 00:01:43.010 --> 00:01:44.630 His clinical interests include
- NOTE Confidence: 0.953192257142857
- $00:01:44.630 \longrightarrow 00:01:46.903$ cancer of the ovary, uterus,
- NOTE Confidence: 0.953192257142857

00:01:46.903 - > 00:01:50.168 vagina, cervix, and the vova.

NOTE Confidence: 0.953192257142857

00:01:50.170 --> 00:01:51.850 Interperitoneal chemotherapy,

NOTE Confidence: 0.953192257142857

00:01:51.850 --> 00:01:56.050 tumor genetics, immunology and immunotherapy,

NOTE Confidence: 0.953192257142857

00:01:56.050 --> 00:01:57.294 tumor angiogenesis,

NOTE Confidence: 0.953192257142857

 $00:01:57.294 \rightarrow 00:01:59.782$ radiation biology and experimental

NOTE Confidence: 0.953192257142857

 $00:01:59.782 \rightarrow 00:02:03.170$ the rapeutics, and gynecologic oncology.

NOTE Confidence: 0.953192257142857

 $00:02:03.170 \longrightarrow 00:02:03.634$ Dr.

NOTE Confidence: 0.953192257142857

 $00:02:03.634 \longrightarrow 00:02:05.490$ Santine's current research interests

NOTE Confidence: 0.953192257142857

 $00:02:05.490 \rightarrow 00:02:08.570$ include immunotherapy for ovarian,

NOTE Confidence: 0.953192257142857

 $00:02:08.570 \rightarrow 00:02:10.970$ cervical and endometrial carcinomas,

NOTE Confidence: 0.953192257142857

00:02:10.970 --> 00:02:13.882 refractory to standard treatment modalities,

NOTE Confidence: 0.953192257142857

 $00:02:13.882 \rightarrow 00:02:16.622$ and development of the rapeutic vaccines.

NOTE Confidence: 0.953192257142857

 $00:02:16.630 \dashrightarrow 00:02:18.510$ Against the human papilloma virus,

NOTE Confidence: 0.953192257142857

 $00:02:18.510 \longrightarrow 00:02:20.845$ infected genital tumors and the

NOTE Confidence: 0.953192257142857

 $00{:}02{:}20.845 \dashrightarrow 00{:}02{:}22.713$ use of monoclonal antibodies

NOTE Confidence: 0.953192257142857

 $00:02:22.713 \rightarrow 00:02:24.900$ against chemotherapy resistant

- NOTE Confidence: 0.953192257142857
- 00:02:24.900 --> 00:02:26.548 gynecologic malignancies.
- NOTE Confidence: 0.953192257142857
- $00:02:26.550 \rightarrow 00:02:26.906$ Dr.
- NOTE Confidence: 0.953192257142857
- 00:02:26.906 --> 00:02:29.042 Santine has more than 300 original
- NOTE Confidence: 0.953192257142857
- $00{:}02{:}29{.}042 \dashrightarrow 00{:}02{:}31{.}509$ research and peer reviewed publications
- NOTE Confidence: 0.953192257142857
- $00{:}02{:}31{.}510 \dashrightarrow 00{:}02{:}33{.}502$ and has written extensively on
- NOTE Confidence: 0.953192257142857
- $00{:}02{:}33{.}502 \dashrightarrow 00{:}02{:}35{.}692$ various topics including cancer of
- NOTE Confidence: 0.953192257142857
- $00:02:35.692 \rightarrow 00:02:38.195$ the ovary Endometrium cervix as well
- NOTE Confidence: 0.953192257142857
- $00:02:38.195 \rightarrow 00:02:41.200$ as on tumor immunology and immunotherapy.
- NOTE Confidence: 0.953192257142857
- $00{:}02{:}41{.}200 \dashrightarrow 00{:}02{:}43{.}433$ I thank both speakers for sharing their
- NOTE Confidence: 0.953192257142857
- $00:02:43.433 \rightarrow 00:02:45.400$ expertise with us and I welcome Dr.
- NOTE Confidence: 0.953192257142857
- $00:02:45.400 \longrightarrow 00:02:46.280$ Ratner to the podium.
- NOTE Confidence: 0.953192257142857
- 00:02:46.280 --> 00:02:46.720 Thank you.
- NOTE Confidence: 0.950317
- $00:02:51.920 \longrightarrow 00:02:54.240$ Thank you so much that Collinsburg.
- NOTE Confidence: 0.936479825
- $00{:}02{:}54{.}240 \dashrightarrow 00{:}02{:}56{.}120$ Let me bring us to the beginning the
- NOTE Confidence: 0.86821436
- $00:03:00.520 \rightarrow 00:03:02.080$ oldfashioned way. I'm just going to bring it,
- NOTE Confidence: 0.8682695

 $00:03:04.240 \rightarrow 00:03:05.560$ we're sure this more sophisticated way. NOTE Confidence: 0.8682695 00:03:05.560 -> 00:03:08.976 Here we go. So we are so excited NOTE Confidence: 0.8682695 00:03:08.976 --> 00:03:10.730 to speak with you today. NOTE Confidence: 0.8682695 00:03:10.730 --> 00:03:12.710 Thank you for giving us opportunity NOTE Confidence: 0.8682695 $00:03:12.769 \longrightarrow 00:03:14.897$ to present to you the work of our NOTE Confidence: 0.8682695 $00:03:14.897 \rightarrow 00:03:17.166$ division and to discuss the state of NOTE Confidence: 0.8682695 $00:03:17.166 \longrightarrow 00:03:18.841$ the union of gynecological cancers NOTE Confidence: 0.8682695 $00:03:18.850 \rightarrow 00:03:21.010$ especially in the State of Connecticut. NOTE Confidence: 0.8682695 00:03:21.010 --> 00:03:22.098 Just like everything else NOTE Confidence: 0.8682695 00:03:22.098 - 00:03:23.730 in the Centina and I do, NOTE Confidence: 0.8682695 $00:03:23.730 \rightarrow 00:03:25.004$ we are going to double team this. NOTE Confidence: 0.8682695 $00:03:25.010 \rightarrow 00:03:27.584$ I will start with clinical presentation NOTE Confidence: 0.8682695 $00{:}03{:}27{.}584 \dashrightarrow 00{:}03{:}29{.}874$ and Doctor Centine will discuss NOTE Confidence: 0.8682695 00:03:29.874 --> 00:03:31.890 his very impressive research, NOTE Confidence: 0.8682695 $00:03:31.890 \longrightarrow 00:03:34.599$ so ovarian cancer. NOTE Confidence: 0.8682695 $00:03:34.599 \rightarrow 00:03:39.114$ Continues to be unacceptably deadly.

- NOTE Confidence: 0.8682695
- $00:03:39.120 \longrightarrow 00:03:42.678$ It's the survival of ovarian cancer.

 $00:03:42.680 \rightarrow 00:03:43.250$ Interestingly,

NOTE Confidence: 0.8682695

00:03:43.250 - 00:03:44.960 shockingly and unacceptably

NOTE Confidence: 0.8682695

 $00:03:44.960 \rightarrow 00:03:47.810$ has really changed very little

NOTE Confidence: 0.8682695

 $00:03:47.810 \longrightarrow 00:03:50.056$ over the past 20-30 years.

NOTE Confidence: 0.8682695

 $00:03:50.056 \dashrightarrow 00:03:53.612$ The reason for this is multiple call

NOTE Confidence: 0.8682695

 $00{:}03{:}53.612 \dashrightarrow 00{:}03{:}57.375$ there is there continues to be a

NOTE Confidence: 0.8682695

 $00:03:57.375 \rightarrow 00:04:00.350$ profoundly late detection ovarian cancers.

NOTE Confidence: 0.8682695

 $00{:}04{:}00{.}350 \dashrightarrow 00{:}04{:}03{.}870$ There is a lot of research that says

NOTE Confidence: 0.8682695

00:04:03.870 --> 00:04:06.621 that ovarian cancers by the time that

NOTE Confidence: 0.8682695

00:04:06.621 $\operatorname{-->}$ 00:04:08.689 diagnosed usually have been there for

NOTE Confidence: 0.8682695

00:04:08.689 --> 00:04:11.350 as long as 24 months and frequently

NOTE Confidence: 0.8682695

 $00{:}04{:}11{.}350 \dashrightarrow 00{:}04{:}14{.}110$ women has seen six other physicians,

NOTE Confidence: 0.8682695

 $00{:}04{:}14{.}110 \dashrightarrow 00{:}04{:}17{.}220$ six other providers prior to

NOTE Confidence: 0.8682695

 $00{:}04{:}17{.}220 \dashrightarrow 00{:}04{:}19{.}708$ actually finally being diagnosed.

00:04:19.710 --> 00:04:21.990 And that is why advocacy is so important,

NOTE Confidence: 0.8682695

 $00{:}04{:}21{.}990 \dashrightarrow 00{:}04{:}25{.}204$ that's why education is so important and

NOTE Confidence: 0.8682695

 $00{:}04{:}25{.}204 \dashrightarrow 00{:}04{:}29{.}476$ that is why we so profoundly and strongly.

NOTE Confidence: 0.8682695

 $00:04:29.480 \rightarrow 00:04:33.340$ Support encourage personalized care,

NOTE Confidence: 0.8682695

 $00{:}04{:}33{.}340 \dashrightarrow 00{:}04{:}35{.}690$ personalized care to patients in

NOTE Confidence: 0.8682695

 $00{:}04{:}35{.}690 \dashrightarrow 00{:}04{:}38{.}319$ the setting of early detection,

NOTE Confidence: 0.8682695

 $00:04:38.320 \longrightarrow 00:04:40.320$ in the setting of prevention,

NOTE Confidence: 0.8682695

 $00{:}04{:}40{.}320 \dashrightarrow 00{:}04{:}42{.}608$ which as I mentioned to you is so

NOTE Confidence: 0.8682695

00:04:42.608 --> 00:04:44.422 important in the disease because

NOTE Confidence: 0.8682695

 $00:04:44.422 \longrightarrow 00:04:46.382$ early detection is so difficult

NOTE Confidence: 0.8682695

 $00{:}04{:}46{.}382 \dashrightarrow 00{:}04{:}48{.}598$ because the symptoms are so vague.

NOTE Confidence: 0.8682695

00:04:48.600 $-\!\!>$ 00:04:51.276 Personalized care in terms of surgery,

NOTE Confidence: 0.8682695

 $00{:}04{:}51{.}280 \dashrightarrow 00{:}04{:}53{.}506$ surgery in this disease and it's

NOTE Confidence: 0.8682695

 $00{:}04{:}53.506 \dashrightarrow 00{:}04{:}55.937$ incredibly important and we will talk

NOTE Confidence: 0.8682695

 $00{:}04{:}55{.}937 \dashrightarrow 00{:}04{:}57{.}709$ today about stratifying patients.

NOTE Confidence: 0.8682695

 $00{:}04{:}57{.}710 \dashrightarrow 00{:}04{:}59{.}943$ To who is able to have this

- NOTE Confidence: 0.8682695
- $00:04:59.943 \rightarrow 00:05:01.305$ surgery minimally invasive and

 $00{:}05{:}01{.}305 \dashrightarrow 00{:}05{:}03{.}025$ we'll talk briefly about hypec,

NOTE Confidence: 0.8682695

 $00:05:03.030 \longrightarrow 00:05:05.914$ something that one of our colleagues Bargain

NOTE Confidence: 0.8682695

 $00:05:05.914 \rightarrow 00:05:08.789$ and Dekean is very passionate about.

NOTE Confidence: 0.8682695

 $00{:}05{:}08.790 \dashrightarrow 00{:}05{:}10.475$ And then the Constantine will

NOTE Confidence: 0.8682695

 $00:05:10.475 \longrightarrow 00:05:12.160$ talk about the importance of

NOTE Confidence: 0.8682695

 $00:05:12.218 \rightarrow 00:05:14.308$ personalized care and the treatment,

NOTE Confidence: 0.8682695

 $00:05:14.310 \longrightarrow 00:05:15.334$ the management,

NOTE Confidence: 0.8682695

 $00{:}05{:}15{.}334 \dashrightarrow 00{:}05{:}17{.}894$ the maintenance of these women.

NOTE Confidence: 0.8682695

 $00:05:17.900 \dashrightarrow 00:05:21.100$ We are incredibly blessed to be in

NOTE Confidence: 0.8682695

 $00:05:21.100 \longrightarrow 00:05:23.100$ a wonderful supportive environment.

NOTE Confidence: 0.8682695

 $00{:}05{:}23.100 \dashrightarrow 00{:}05{:}25.164$ We are so happy that some of our

NOTE Confidence: 0.8682695

 $00:05:25.164 \longrightarrow 00:05:26.459$ team is here today.

NOTE Confidence: 0.8682695

 $00{:}05{:}26{.}460 \dashrightarrow 00{:}05{:}28{.}530$ We have the most incredible research

NOTE Confidence: 0.8682695

 $00:05:28.530 \dashrightarrow 00:05:30.731$ team and the most incredible clinical

- $00:05:30.731 \rightarrow 00:05:33.370$ team and none of this would be
- NOTE Confidence: 0.8682695
- $00:05:33.370 \dashrightarrow 00:05:35.217$ possible for sure without them.
- NOTE Confidence: 0.8682695
- $00:05:35.220 \dashrightarrow 00:05:37.380$ So very briefly, we don't have much time.
- NOTE Confidence: 0.8682695
- $00{:}05{:}37{.}380 \dashrightarrow 00{:}05{:}38{.}394$ So I'm going to be skipping
- NOTE Confidence: 0.8682695
- $00:05:38.394 \longrightarrow 00:05:39.380$ through a lot of things,
- NOTE Confidence: 0.8682695
- $00:05:39.380 \dashrightarrow 00:05:41.414$ but just kind of to set the stage today,
- NOTE Confidence: 0.8682695
- $00:05:41.420 \longrightarrow 00:05:42.645$ we're going to be talking
- NOTE Confidence: 0.8682695
- $00:05:42.645 \longrightarrow 00:05:43.380$ about epithelial cancers,
- NOTE Confidence: 0.8682695
- 00:05:43.380 --> 00:05:44.094 you know,
- NOTE Confidence: 0.8682695
- $00{:}05{:}44.094 \dashrightarrow 00{:}05{:}44.808$ ovarian cancers.
- NOTE Confidence: 0.8682695
- $00{:}05{:}44.808 \dashrightarrow 00{:}05{:}46.593$ Ovarian cells in general have
- NOTE Confidence: 0.8682695
- 00:05:46.593 --> 00:05:47.818 different kinds of cells,
- NOTE Confidence: 0.8682695
- $00:05:47.820 \rightarrow 00:05:49.098$ and epithelial cancers are the ones
- NOTE Confidence: 0.8682695
- $00:05:49.098 \dashrightarrow 00:05:50.460$ we're going to talk about today.
- NOTE Confidence: 0.8682695
- $00{:}05{:}50{.}460 \dashrightarrow 00{:}05{:}51{.}576$ Those are the most common ones.
- NOTE Confidence: 0.8682695
- $00:05:51.580 \dashrightarrow 00:05:53.140$ And when you hear about people

- NOTE Confidence: 0.8682695
- $00:05:53.140 \longrightarrow 00:05:54.180$ talking about ovarian cancer,

00:05:54.180 --> 00:05:56.260 usually it's about epithelial,

NOTE Confidence: 0.8682695

 $00:05:56.260 \rightarrow 00:05:59.380$ which is serious serous endometrioid clear

NOTE Confidence: 0.8682695

 $00:05:59.453 \rightarrow 00:06:02.155$ cell we use in this or carcinosarcoma.

NOTE Confidence: 0.8682695

 $00:06:02.160 \longrightarrow 00:06:02.624$ Nowadays,

NOTE Confidence: 0.8682695

 $00:06:02.624 \rightarrow 00:06:05.408$ things like genomics and things like

NOTE Confidence: 0.8682695

 $00{:}06{:}05{.}408 \dashrightarrow 00{:}06{:}07{.}631$ mutations that drive the cancers

NOTE Confidence: 0.8682695

 $00:06:07.631 \longrightarrow 00:06:09.917$ is so much more important actually

NOTE Confidence: 0.8682695

 $00:06:09.917 \rightarrow 00:06:12.513$ that the kind of Histology that the

NOTE Confidence: 0.8682695

 $00{:}06{:}12.513 \dashrightarrow 00{:}06{:}14.685$ cancer has and again the Constantine

NOTE Confidence: 0.8682695

 $00:06:14.685 \longrightarrow 00:06:17.115$ will share all that with us.

NOTE Confidence: 0.8682695

 $00:06:17.120 \longrightarrow 00:06:18.920$ But in terms of prevention,

NOTE Confidence: 0.8682695

00:06:18.920 --> 00:06:20.320 in terms of early detection,

NOTE Confidence: 0.94919884444445

 $00{:}06{:}20{.}320 \dashrightarrow 00{:}06{:}21{.}848$ in terms of treatment,

NOTE Confidence: 0.94919884444445

 $00:06:21.848 \dashrightarrow 00:06:23.758$ in terms of clinical presentation,

 $00:06:23.760 \rightarrow 00:06:26.096$ so much time and effort now is being

NOTE Confidence: 0.94919884444445

 $00{:}06{:}26.096 \dashrightarrow 00{:}06{:}28.639$ spent on understanding predisposition.

NOTE Confidence: 0.94919884444445

 $00:06:28.640 \rightarrow 00:06:30.344$ You know, we have not succeeded

NOTE Confidence: 0.94919884444445

 $00:06:30.344 \rightarrow 00:06:31.480$ in curing these cancers.

NOTE Confidence: 0.94919884444445

 $00{:}06{:}31.480 \dashrightarrow 00{:}06{:}33.692$ We have not succeeded not because of

NOTE Confidence: 0.94919884444445

 $00:06:33.692 \rightarrow 00:06:35.879$ lack of trying in early detection.

NOTE Confidence: 0.94919884444445

 $00:06:35.880 \longrightarrow 00:06:38.120$ So the next best thing,

NOTE Confidence: 0.94919884444445

 $00:06:38.120 \rightarrow 00:06:41.040$ and really just the best thing is prevention.

NOTE Confidence: 0.94919884444445

 $00{:}06{:}41.040 \dashrightarrow 00{:}06{:}42.605$ We know that there's certain

NOTE Confidence: 0.94919884444445

 $00:06:42.605 \rightarrow 00:06:43.857$ genetic mutations that increase

NOTE Confidence: 0.94919884444445

 $00{:}06{:}43.857 \dashrightarrow 00{:}06{:}45.400$ the risk of bearing cancers.

NOTE Confidence: 0.94919884444445

00:06:45.400 --> 00:06:48.585 BEAR, C1, BEAR, C2 and Lynch all

NOTE Confidence: 0.94919884444445

00:06:48.585 --> 00:06:50.440 profoundly increase the risk.

NOTE Confidence: 0.94919884444445

 $00:06:50.440 \longrightarrow 00:06:52.960$ And that is why it is so important

NOTE Confidence: 0.94919884444445

 $00{:}06{:}52{.}960 \dashrightarrow 00{:}06{:}55{.}478$ that women know their predisposition.

NOTE Confidence: 0.94919884444445

 $00:06:55.480 \dashrightarrow 00:06:57.877$ We do a lot of advocacy and a lot of talks

 $00:06:57.877 \rightarrow 00:07:00.157$ to talk to women about know your genes,

NOTE Confidence: 0.94919884444445

00:07:00.160 --> 00:07:01.180 know your predisposition,

NOTE Confidence: 0.94919884444445

 $00:07:01.180 \longrightarrow 00:07:02.880$ know what kind of genetic

NOTE Confidence: 0.94919884444445

00:07:02.880 --> 00:07:04.240 mutations you might carry,

NOTE Confidence: 0.94919884444445

 $00:07:04.240 \longrightarrow 00:07:06.088$ because a lot of these women

NOTE Confidence: 0.94919884444445

 $00:07:06.088 \longrightarrow 00:07:07.920$ could be saved not by cure,

NOTE Confidence: 0.94919884444445

 $00:07:07.920 \longrightarrow 00:07:10.128$ not by early detection,

NOTE Confidence: 0.94919884444445

 $00:07:10.128 \longrightarrow 00:07:12.336$ but actually by prevention.

NOTE Confidence: 0.94919884444445

 $00:07:12.340 \longrightarrow 00:07:14.578$ And so much now in treatment

NOTE Confidence: 0.94919884444445

 $00:07:14.578 \rightarrow 00:07:16.520$ depends on the pathophysiology of

NOTE Confidence: 0.94919884444445

 $00:07:16.520 \rightarrow 00:07:18.420$ these high grade serious cancers.

NOTE Confidence: 0.94919884444445

 $00{:}07{:}18.420 \dashrightarrow 00{:}07{:}21.500$ And we know so much now about different

NOTE Confidence: 0.94919884444445

 $00:07:21.500 \dashrightarrow 00:07:23.660$ DNA mechanisms that drive these cancers.

NOTE Confidence: 0.94919884444445

 $00{:}07{:}23.660 \dashrightarrow 00{:}07{:}26.060$ One of the biggest one is

NOTE Confidence: 0.94919884444445

 $00{:}07{:}26.060 \dashrightarrow 00{:}07{:}26.860$ homogeneous recombination.

 $00:07:26.860 \rightarrow 00:07:29.037$ That is something that we have really

NOTE Confidence: 0.94919884444445

 $00:07:29.037 \longrightarrow 00:07:30.778$ explored widely over the past five,

NOTE Confidence: 0.94919884444445

 $00:07:30.780 \longrightarrow 00:07:32.859$ seven years and where a lot of

NOTE Confidence: 0.94919884444445

 $00:07:32.859 \rightarrow 00:07:34.538$ the targeted drugs play a role.

NOTE Confidence: 0.949598185714286

00:07:36.600 --> 00:07:38.348 So briefly, ovarian cancer,

NOTE Confidence: 0.949598185714286

00:07:38.348 --> 00:07:40.096 second most common malignancy

NOTE Confidence: 0.949598185714286

 $00:07:40.096 \rightarrow 00:07:42.480$ of the of the general tract,

NOTE Confidence: 0.949598185714286

 $00:07:42.480 \longrightarrow 00:07:44.228$ by far the deadliest.

NOTE Confidence: 0.949598185714286

 $00:07:44.228 \rightarrow 00:07:47.960$ The incidence is actually not that high 1.3%,

NOTE Confidence: 0.949598185714286

 $00:07:47.960 \longrightarrow 00:07:52.232$ but as you see 21,000 women get

NOTE Confidence: 0.949598185714286

 $00{:}07{:}52.232 \dashrightarrow 00{:}07{:}55.611$ diagnosed and 13,000 women die.

NOTE Confidence: 0.949598185714286

00:07:55.611 --> 00:07:58.879 So incredibly deadly cancer.

NOTE Confidence: 0.949598185714286

00:07:58.880 --> 00:08:00.800 It is very interesting,

NOTE Confidence: 0.949598185714286

 $00:08:00.800 \rightarrow 00:08:03.200$ there's for sure geographic distribution,

NOTE Confidence: 0.949598185714286

 $00:08:03.200 \rightarrow 00:08:06.470$ there's for sure high risk populations.

NOTE Confidence: 0.949598185714286

 $00:08:06.470 \rightarrow 00:08:08.322$ Women with Caucasian populations,

 $00:08:08.322 \rightarrow 00:08:11.510$ women that come from smaller family size,

NOTE Confidence: 0.949598185714286

 $00:08:11.510 \rightarrow 00:08:12.938$ higher socioeconomic status and

NOTE Confidence: 0.949598185714286

 $00:08:12.938 \longrightarrow 00:08:15.470$ high fat diet all play a role.

NOTE Confidence: 0.949598185714286

 $00:08:15.470 \longrightarrow 00:08:18.137$ And what's interesting is that when women

NOTE Confidence: 0.949598185714286

 $00{:}08{:}18.137 \dashrightarrow 00{:}08{:}20.854$ move from the region that's of lower

NOTE Confidence: 0.949598185714286

 $00:08:20.854 \dashrightarrow 00:08:23.870$ incidence to a region of higher incidence,

NOTE Confidence: 0.949598185714286

 $00:08:23.870 \rightarrow 00:08:28.070$ most sadly they acquire the higher risk of.

NOTE Confidence: 0.949598185714286

00:08:28.070 - 00:08:29.546 So it's now where you're born,

NOTE Confidence: 0.949598185714286

00:08:29.550 --> 00:08:30.710 it's where you grow up,

NOTE Confidence: 0.949598185714286

 $00:08:30.710 \longrightarrow 00:08:32.070$ where you where you live,

NOTE Confidence: 0.949598185714286

 $00:08:32.070 \rightarrow 00:08:35.338$ which is very interesting.

NOTE Confidence: 0.949598185714286

00:08:35.340 --> 00:08:37.140 So there's many different studies,

NOTE Confidence: 0.949598185714286

 $00:08:37.140 \longrightarrow 00:08:39.576$ of course, that talk about predisposition.

NOTE Confidence: 0.949598185714286

 $00{:}08{:}39{.}580 \dashrightarrow 00{:}08{:}40{.}396$ And you know,

NOTE Confidence: 0.949598185714286

 $00:08:40.396 \rightarrow 00:08:42.300$ we know that obesity plays a role.

 $00:08:42.300 \longrightarrow 00:08:45.660$ We know that women who have

NOTE Confidence: 0.949598185714286

00:08:45.660 --> 00:08:47.440 their periods start early,

NOTE Confidence: 0.949598185714286

 $00:08:47.440 \longrightarrow 00:08:49.220$ their menopause come late,

NOTE Confidence: 0.949598185714286

 $00:08:49.220 \rightarrow 00:08:50.816$ women who do not have kids,

NOTE Confidence: 0.949598185714286

 $00{:}08{:}50.820 \dashrightarrow 00{:}08{:}52.220$ women who have endometriosis.

NOTE Confidence: 0.949598185714286

 $00:08:52.220 \rightarrow 00:08:55.330$ We know these women have a predisposition,

NOTE Confidence: 0.949598185714286

 $00{:}08{:}55{.}330 \dashrightarrow 00{:}08{:}57{.}339$ and the reason for that is because

NOTE Confidence: 0.949598185714286

 $00{:}08{:}57{.}339 \dashrightarrow 00{:}08{:}58{.}584$ there's two different theories

NOTE Confidence: 0.949598185714286

 $00:08:58.584 \dashrightarrow 00:09:00.249$ of how ovarian cancers happen.

NOTE Confidence: 0.949598185714286

 $00:09:00.250 \rightarrow 00:09:03.210$ One is that every time a woman ovulates,

NOTE Confidence: 0.949598185714286

 $00:09:03.210 \longrightarrow 00:09:05.250$ some sort of a process happens.

NOTE Confidence: 0.949598185714286

 $00:09:05.250 \dashrightarrow 00:09:06.606$ And the more time she ovulates,

NOTE Confidence: 0.949598185714286

 $00:09:06.610 \longrightarrow 00:09:08.662$ the higher is her risk of

NOTE Confidence: 0.949598185714286

 $00:09:08.662 \rightarrow 00:09:09.688$ developing ovarian cancer.

NOTE Confidence: 0.949598185714286

 $00:09:09.690 \rightarrow 00:09:12.406$ That's why certain things are so protective.

NOTE Confidence: 0.949598185714286

 $00:09:12.410 \longrightarrow 00:09:13.718$ So, for example,

- NOTE Confidence: 0.949598185714286
- $00:09:13.718 \rightarrow 00:09:16.334$ women who have had five pregnancies
- NOTE Confidence: 0.949598185714286
- $00:09:16.334 \rightarrow 00:09:19.080$ and breastfeed each every child for
- NOTE Confidence: 0.949598185714286
- $00:09:19.080 \longrightarrow 00:09:21.986$ one year have a 50% reduction rate.
- NOTE Confidence: 0.949598185714286
- $00:09:21.986 \rightarrow 00:09:25.820$ I've worked on the plan very, very different.
- NOTE Confidence: 0.949598185714286
- $00:09:25.820 \longrightarrow 00:09:27.788$ The most same option is the
- NOTE Confidence: 0.949598185714286
- 00:09:27.788 --> 00:09:28.772 birth control pills.
- NOTE Confidence: 0.949598185714286
- 00:09:28.780 --> 00:09:29.628 So similar,
- NOTE Confidence: 0.949598185714286
- $00:09:29.628 \rightarrow 00:09:32.596$ any woman who uses birth control pills
- NOTE Confidence: 0.949598185714286
- $00:09:32.596 \longrightarrow 00:09:35.908$ for five years has a reduction of 50%.
- NOTE Confidence: 0.949598185714286
- 00:09:35.908 --> 00:09:38.020 So very, very impressive.
- NOTE Confidence: 0.949598185714286
- 00:09:38.020 --> 00:09:41.028 Women who have used OCP's birth control pills
- NOTE Confidence: 0.949598185714286
- $00:09:41.028 \rightarrow 00:09:45.530$ for 10 years have reduction as high as 80%.
- NOTE Confidence: 0.949598185714286
- $00{:}09{:}45{.}530 \dashrightarrow 00{:}09{:}48{.}498$ And that is all cumulative and that's all
- NOTE Confidence: 0.949598185714286
- $00:09:48.498 \dashrightarrow 00:09:51.209$ respective to the woman personalized risk.
- NOTE Confidence: 0.949598185714286
- $00:09:51.210 \rightarrow 00:09:53.240$ So even somebody who has a BRC
- NOTE Confidence: 0.949598185714286

 $00:09:53.240 \longrightarrow 00:09:54.798$ mutation and has a 40% risk,

NOTE Confidence: 0.949598185714286

 $00:09:54.798 \longrightarrow 00:09:56.646$ if they have used OC PS:,

NOTE Confidence: 0.949598185714286

 $00:09:56.650 \longrightarrow 00:09:57.343$ for five years,

NOTE Confidence: 0.949598185714286

 $00:09:57.343 \longrightarrow 00:09:58.729$ the risk goes down to 20%.

NOTE Confidence: 0.949598185714286

 $00:09:58.730 \longrightarrow 00:10:00.250$ They've used it for 10:15,

NOTE Confidence: 0.949598185714286

 $00:10:00.250 \longrightarrow 00:10:01.528$ it goes to 10:00 and 8:00.

NOTE Confidence: 0.949598185714286

 $00:10:01.530 \longrightarrow 00:10:03.398$ So with different modifications,

NOTE Confidence: 0.949598185714286

 $00:10:03.398 \longrightarrow 00:10:05.733$ we actually can do risk

NOTE Confidence: 0.949598185714286

 $00:10:05.733 \dashrightarrow 00:10:07.807$ adjustment and decrease the risk.

NOTE Confidence: 0.949598185714286

 $00{:}10{:}07{.}810 \dashrightarrow 00{:}10{:}10{.}048$ And then we know there's also

NOTE Confidence: 0.949598185714286

 $00:10:10.048 \longrightarrow 00:10:11.540$ inflammatory factors such as

NOTE Confidence: 0.949598185714286

00:10:11.609 --> 00:10:13.281 pelvic inflammatory disease and

NOTE Confidence: 0.949598185714286

 $00:10:13.281 \dashrightarrow 00:10:16.180$ endometriosis that play a role as well.

NOTE Confidence: 0.949598185714286

 $00:10:16.180 \longrightarrow 00:10:18.178$ The second theory of ovarian cancer

NOTE Confidence: 0.949598185714286

 $00:10:18.178 \longrightarrow 00:10:19.920$ is actually that these cancers

NOTE Confidence: 0.949598185714286

 $00:10:19.920 \longrightarrow 00:10:21.670$ are not potentially not even

- NOTE Confidence: 0.949598185714286
- $00{:}10{:}21.670 \dashrightarrow 00{:}10{:}23.420$ ovarian cancers to begin with.
- NOTE Confidence: 0.949598185714286
- $00:10:23.420 \rightarrow 00:10:25.980$ They're actually fallopian tube cancers,
- NOTE Confidence: 0.949598185714286
- $00:10:25.980 \rightarrow 00:10:28.218$ especially in women with genetic mutations.
- NOTE Confidence: 0.949598185714286
- $00:10:28.220 \rightarrow 00:10:29.914$ And these cancers start in the Philippine
- NOTE Confidence: 0.949598185714286
- $00:10:29.914 \rightarrow 00:10:31.860$ tubes and only then spread to the ovaries.
- NOTE Confidence: 0.949598185714286
- $00{:}10{:}31.860 \dashrightarrow 00{:}10{:}34.068$ And that is why now adays there's such a
- NOTE Confidence: 0.949598185714286
- 00:10:34.068 --> 00:10:36.129 culture of actually removing fallopian
- NOTE Confidence: 0.949598185714286
- $00:10:36.129 \rightarrow 00:10:38.057$ tubes at different opportunities,
- NOTE Confidence: 0.949598185714286
- $00{:}10{:}38.060 \dashrightarrow 00{:}10{:}40.215$ because that creates a risk
- NOTE Confidence: 0.949598185714286
- $00:10:40.215 \rightarrow 00:10:43.050$ reduction as high as 50 to 70%.
- NOTE Confidence: 0.949598185714286
- $00:10:43.050 \rightarrow 00:10:45.048$ So here within our our institution,
- NOTE Confidence: 0.949598185714286
- $00:10:45.050 \longrightarrow 00:10:46.625$ we actually have been able
- NOTE Confidence: 0.949598185714286
- $00:10:46.625 \longrightarrow 00:10:47.885$ to influence the culture.
- NOTE Confidence: 0.949598185714286
- $00:10:47.890 \longrightarrow 00:10:49.890$ And in the older days,
- NOTE Confidence: 0.949598185714286
- $00:10:49.890 \longrightarrow 00:10:50.679$ 10 years ago,
- NOTE Confidence: 0.949598185714286

 $00:10:50.679 \rightarrow 00:10:52.520$ when people when women used to have

NOTE Confidence: 0.941801712307693

00:10:52.577 --> 00:10:53.489 their tubes tied,

NOTE Confidence: 0.941801712307693

 $00:10:53.490 \rightarrow 00:10:55.128$ they literally have their tubes tied.

NOTE Confidence: 0.941801712307693

 $00:10:55.130 \rightarrow 00:10:57.125$ Now we actually remove the fallopian tubes,

NOTE Confidence: 0.941801712307693

 $00{:}10{:}57{.}130 \dashrightarrow 00{:}10{:}59{.}380$ especially in women with higher

NOTE Confidence: 0.941801712307693

 $00{:}10{:}59{.}380 \dashrightarrow 00{:}11{:}00{.}730$ predisposition to cancers.

NOTE Confidence: 0.941801712307693

 $00:11:00.730 \longrightarrow 00:11:03.075$ We are able to to provide a

NOTE Confidence: 0.941801712307693

00:11:03.075 --> 00:11:04.608 very substantial risk reduction

NOTE Confidence: 0.941801712307693

 $00{:}11{:}04.608 \dashrightarrow 00{:}11{:}06.888$ just by that simple technique.

NOTE Confidence: 0.941801712307693

 $00:11:06.890 \longrightarrow 00:11:07.794$ In the older days,

NOTE Confidence: 0.941801712307693

 $00{:}11{:}07.794 \dashrightarrow 00{:}11{:}09.150$ we used to leave fallopian tubes

NOTE Confidence: 0.941801712307693

 $00:11:09.203 \longrightarrow 00:11:10.623$ anytime we left ovaries when

NOTE Confidence: 0.941801712307693

 $00:11:10.623 \rightarrow 00:11:12.043$ somebody would have a hysterectomy.

NOTE Confidence: 0.941801712307693

00:11:12.050 --> 00:11:13.885 Nowadays, for sure fallopian tubes

NOTE Confidence: 0.941801712307693

 $00{:}11{:}13.885 \dashrightarrow 00{:}11{:}16.114$ would be removed because we know

NOTE Confidence: 0.941801712307693

 $00:11:16.114 \rightarrow 00:11:18.396$ that there's a huge component of risk

 $00:11:18.396 \rightarrow 00:11:20.178$ reduction by removing the fallopian

NOTE Confidence: 0.941801712307693

 $00:11:20.178 \rightarrow 00:11:22.670$ tube and it's a very easy adjustment.

NOTE Confidence: 0.941801712307693

00:11:22.670 --> 00:11:24.190 But of course, out of all those things,

NOTE Confidence: 0.941801712307693

 $00:11:24.190 \rightarrow 00:11:28.310$ the most highest risk is the family history.

NOTE Confidence: 0.941801712307693

 $00:11:28.310 \longrightarrow 00:11:31.572$ Anybody who has a family first degree

NOTE Confidence: 0.941801712307693

 $00:11:31.572 \rightarrow 00:11:34.374$ family member who has ovarian cancer,

NOTE Confidence: 0.941801712307693

 $00:11:34.374 \longrightarrow 00:11:36.238$ that increases that woman's

NOTE Confidence: 0.941801712307693

 $00:11:36.238 \longrightarrow 00:11:38.029$ lifetime risk to 4 to 5%.

NOTE Confidence: 0.941801712307693

 $00{:}11{:}38{.}030 \dashrightarrow 00{:}11{:}39{.}548$ And as I showed you before,

NOTE Confidence: 0.941801712307693

00:11:39.550 --> 00:11:42.310 this hereditary breast and ovarian syndromes,

NOTE Confidence: 0.941801712307693

 $00:11:42.310 \longrightarrow 00:11:44.907$ which account for 20% of ovarian cancers,

NOTE Confidence: 0.941801712307693

 $00:11:44.910 \longrightarrow 00:11:45.970$ but we really think,

NOTE Confidence: 0.941801712307693

 $00{:}11{:}45{.}970 \dashrightarrow 00{:}11{:}47{.}030$ account for much more.

NOTE Confidence: 0.941801712307693

 $00{:}11{:}47.030 \dashrightarrow 00{:}11{:}49.990$ We just don't yet know exactly which ones.

NOTE Confidence: 0.941801712307693

 $00{:}11{:}49{.}990 \dashrightarrow 00{:}11{:}51{.}748$ They of course carry the the

- $00:11:51.748 \rightarrow 00:11:52.627 \text{ most highest risk.}$
- NOTE Confidence: 0.9553487
- $00:11:54.710 \longrightarrow 00:11:55.624$ Very briefly.
- NOTE Confidence: 0.9553487
- 00:11:55.624 --> 00:11:58.366 There's a very profound and very
- NOTE Confidence: 0.9553487
- $00:11:58.366 \rightarrow 00:12:01.022$ meaningful and very important racial
- NOTE Confidence: 0.9553487
- $00:12:01.022 \rightarrow 00:12:03.787$ health disparities in ovarian cancer.
- NOTE Confidence: 0.9553487
- $00{:}12{:}03.790 \dashrightarrow 00{:}12{:}04.830$ We have known this forever.
- NOTE Confidence: 0.9553487
- $00{:}12{:}04.830 \dashrightarrow 00{:}12{:}07.310$ We have known for ever that
- NOTE Confidence: 0.9553487
- $00:12:07.310 \longrightarrow 00:12:09.550$ women of color do worse.
- NOTE Confidence: 0.9553487
- $00{:}12{:}09{.}550 \dashrightarrow 00{:}12{:}11{.}854$ And we used to think that it's just
- NOTE Confidence: 0.9553487
- $00:12:11.854 \rightarrow 00:12:13.748$ because of the Histology of the
- NOTE Confidence: 0.9553487
- 00:12:13.748 --> 00:12:15.985 cancer that the women who are non
- NOTE Confidence: 0.9553487
- 00:12:15.985 --> 00:12:17.760 Caucasian get Histology that are
- NOTE Confidence: 0.9553487
- $00{:}12{:}17.760 \dashrightarrow 00{:}12{:}19.829$ more aggressive and that is indeed.
- NOTE Confidence: 0.9553487
- 00:12:19.830 --> 00:12:22.370 Case, but unquestionably and deniably
- NOTE Confidence: 0.9553487
- 00:12:22.370 --> 00:12:24.910 confirmed by many different studies,
- NOTE Confidence: 0.9553487
- $00:12:24.910 \longrightarrow 00:12:25.732$ especially recently.

- NOTE Confidence: 0.9553487
- $00:12:25.732 \longrightarrow 00:12:28.198$ We also know the women of

 $00:12:28.198 \longrightarrow 00:12:29.909$ color get worse care.

NOTE Confidence: 0.9553487

 $00:12:29.910 \longrightarrow 00:12:33.210$ They have more profound delays

NOTE Confidence: 0.9553487

 $00:12:33.210 \rightarrow 00:12:35.256$ in diagnosis and chemotherapy.

NOTE Confidence: 0.9553487

00:12:35.256 --> 00:12:38.028 They don't get necessarily this term

NOTE Confidence: 0.9553487

 $00:12:38.028 \rightarrow 00:12:41.052$ of care surgery now with the joint

NOTE Confidence: 0.9553487

 $00{:}12{:}41.052 \dashrightarrow 00{:}12{:}43.298$ on cologist and another very important

NOTE Confidence: 0.9553487

00:12:43.298 --> 00:12:46.614 component is that women of color are

NOTE Confidence: 0.9553487

 $00{:}12{:}46.614 \dashrightarrow 00{:}12{:}49.069$ under enrolled in clinical trials.

NOTE Confidence: 0.9553487

00:12:49.070 --> 00:12:52.670 Only 17% of clinical trial participants

NOTE Confidence: 0.9553487

 $00{:}12{:}52.670 \dashrightarrow 00{:}12{:}56.070$ represent racial or ethnic minorities

NOTE Confidence: 0.9553487

 $00{:}12{:}56.070 \dashrightarrow 00{:}12{:}58.230$ and African American women's

NOTE Confidence: 0.9553487

 $00:12:58.230 \longrightarrow 00:13:00.390$ participation only reached 9/11,

NOTE Confidence: 0.9553487

 $00{:}13{:}00{.}390 \dashrightarrow 00{:}13{:}01{.}350$ 10%, nine percent.

NOTE Confidence: 0.9553487

 $00{:}13{:}01{.}350 \dashrightarrow 00{:}13{:}03{.}955$ So we believe that all of that plays

 $00:13:03.955 \longrightarrow 00:13:06.055$ a very important role and that is

NOTE Confidence: 0.9553487

00:13:06.055 --> 00:13:08.681 why so much effort is being is being

NOTE Confidence: 0.9553487

 $00{:}13{:}08{.}681 \dashrightarrow 00{:}13{:}10{.}790$ made to improve this cruel into

NOTE Confidence: 0.9553487

 $00:13:10.790 \longrightarrow 00:13:13.990$ trials and open it to women of color.

NOTE Confidence: 0.9553487

 $00:13:13.990 \rightarrow 00:13:16.924$ So very briefly just the state of our union.

NOTE Confidence: 0.9553487

00:13:16.930 --> 00:13:18.970 You an ecology here at Yale is very,

NOTE Confidence: 0.9553487

 $00:13:18.970 \longrightarrow 00:13:19.674$ very busy.

NOTE Confidence: 0.9553487

 $00:13:19.674 \rightarrow 00:13:22.490$ We have a great number of surgical cases,

NOTE Confidence: 0.9553487

 $00{:}13{:}22{.}490 \dashrightarrow 00{:}13{:}25{.}575$ new patients and it continues

NOTE Confidence: 0.9553487

 $00:13:25.575 \rightarrow 00:13:28.450$ to increase every year.

NOTE Confidence: 0.9553487

 $00{:}13{:}28{.}450 \dashrightarrow 00{:}13{:}30{.}106$ Our biggest we cover the entire

NOTE Confidence: 0.9553487

00:13:30.106 --> 00:13:31.733 state of Connecticut all the way

NOTE Confidence: 0.9553487

 $00{:}13{:}31{.}733 \dashrightarrow 00{:}13{:}33{.}385$ from Greenwich all the way to New

NOTE Confidence: 0.9553487

 $00{:}13{:}33{.}385 \dashrightarrow 00{:}13{:}34{.}982$ London with the great majority of

NOTE Confidence: 0.9553487

 $00{:}13{:}34{.}982 \dashrightarrow 00{:}13{:}37{.}226$ our care that happens here in New

NOTE Confidence: 0.9553487

 $00:13:37.226 \rightarrow 00:13:39.566$ Haven and then Bridgeport Hospital.

- NOTE Confidence: 0.9553487
- $00{:}13{:}39{.}570 \dashrightarrow 00{:}13{:}42{.}846$ And as you see the number of our new

 $00:13:42.846 \rightarrow 00:13:45.537$ patient visits and surgeries grows.

NOTE Confidence: 0.9553487

 $00:13:45.540 \longrightarrow 00:13:46.968$ Each and every year,

NOTE Confidence: 0.9553487

 $00{:}13{:}46{.}968 \dashrightarrow 00{:}13{:}49{.}110$ and this is the territory that

NOTE Confidence: 0.9553487

 $00:13:49.180 \longrightarrow 00:13:49.966$ we cover again,

NOTE Confidence: 0.9553487

 $00:13:49.966 \longrightarrow 00:13:51.548$ all the way from New York all

NOTE Confidence: 0.9553487

 $00:13:51.548 \rightarrow 00:13:52.779$ the way to New London,

NOTE Confidence: 0.9553487

 $00:13:52.780 \longrightarrow 00:13:56.612$ so very profoundly so there.

NOTE Confidence: 0.9553487

 $00:13:56.612 \longrightarrow 00:13:58.004$ One of the things that have

NOTE Confidence: 0.9553487

 $00{:}13{:}58{.}004 \dashrightarrow 00{:}13{:}59{.}213$ changed drastically in the care

NOTE Confidence: 0.9553487

00:13:59.213 --> 00:14:00.378 of joining College of Patients,

NOTE Confidence: 0.9553487

 $00{:}14{:}00{.}380 \dashrightarrow 00{:}14{:}01{.}430$ something that I'm going to

NOTE Confidence: 0.9553487

 $00:14:01.430 \longrightarrow 00:14:02.060$ talk about briefly,

NOTE Confidence: 0.9553487

 $00{:}14{:}02.060 \dashrightarrow 00{:}14{:}04.916$ is how much we are able to do

NOTE Confidence: 0.9553487

 $00:14:04.916 \rightarrow 00:14:07.790$ surgically in a minimally invasive way.

 $00{:}14{:}07{.}790 \dashrightarrow 00{:}14{:}09{.}490$ So when you look at that data that I just

NOTE Confidence: 0.9553487

 $00:14:09.534 \rightarrow 00:14:11.070$ pulled up and the data that you know,

NOTE Confidence: 0.9553487

00:14:11.070 --> 00:14:13.428 I study very closely and extensively,

NOTE Confidence: 0.9553487

 $00:14:13.430 \longrightarrow 00:14:16.070$ the culture of this field is

NOTE Confidence: 0.9553487

 $00{:}14{:}16.070 \dashrightarrow 00{:}14{:}17.830$ changing to become almost

NOTE Confidence: 0.9553487

 $00{:}14{:}17{.}913 \dashrightarrow 00{:}14{:}20{.}439$ predominantly outpatient field.

NOTE Confidence: 0.9553487

00:14:20.440 --> 00:14:22.462 Great majority of our women used

NOTE Confidence: 0.9553487

 $00:14:22.462 \longrightarrow 00:14:24.621$ to have the surgeries and have

NOTE Confidence: 0.9553487

 $00{:}14{:}24.621 \dashrightarrow 00{:}14{:}26.117$ a prolonged inpatient stay.

NOTE Confidence: 0.9553487

 $00{:}14{:}26{.}120 \dashrightarrow 00{:}14{:}27{.}779$ And now great majority of women are

NOTE Confidence: 0.9553487

 $00{:}14{:}27{.}779 \dashrightarrow 00{:}14{:}30{.}005$ able to have the same kind of radical

NOTE Confidence: 0.9553487

 $00:14:30.005 \rightarrow 00:14:31.465$ debalking surgeries in a minimally

NOTE Confidence: 0.9553487

 $00{:}14{:}31{.}514 \dashrightarrow 00{:}14{:}33{.}138$ invasive way that I'll show you in

NOTE Confidence: 0.9553487

 $00{:}14{:}33{.}138 \dashrightarrow 00{:}14{:}36{.}120$ a little bit and are able to be

NOTE Confidence: 0.9553487

 $00{:}14{:}36{.}120 \dashrightarrow 00{:}14{:}38{.}400$ discharged home literally same day.

NOTE Confidence: 0.9553487

 $00:14:38.400 \longrightarrow 00:14:41.923$ And that is why so much of our financial

- NOTE Confidence: 0.9553487
- 00:14:41.923 --> 00:14:44.738 representation has now become an

 $00{:}14{:}44.738 \dashrightarrow 00{:}14{:}47.860$ outpatient service versus the inpatient.

NOTE Confidence: 0.9553487

00:14:47.860 --> 00:14:50.632 So surgery is very,

NOTE Confidence: 0.9553487

 $00:14:50.632 \rightarrow 00:14:52.018$ very important.

NOTE Confidence: 0.9553487

 $00:14:52.020 \longrightarrow 00:14:54.298$ Let me just get rid of that.

NOTE Confidence: 0.9553487

 $00:14:54.300 \rightarrow 00:14:56.739$ I don't know if that's supposed to be here.

NOTE Confidence: 0.9553487

 $00:14:56.740 \longrightarrow 00:14:57.892$ There we go.

NOTE Confidence: 0.9553487

 $00{:}14{:}57{.}892 \dashrightarrow 00{:}14{:}59{.}812$ So surgery in gynecology is

NOTE Confidence: 0.9553487

 $00:14:59.812 \longrightarrow 00:15:01.460$ of paramount importance.

NOTE Confidence: 0.9553487

 $00:15:01.460 \longrightarrow 00:15:03.215$ We know for many different

NOTE Confidence: 0.9553487

 $00{:}15{:}03.215 \dashrightarrow 00{:}15{:}05.449$ reasons that women who have had

NOTE Confidence: 0.9553487

 $00{:}15{:}05{.}449 \dashrightarrow 00{:}15{:}07{.}469$ very radical surgery and their

NOTE Confidence: 0.9553487

00:15:07.469 --> 00:15:09.085 cancer was removed entirely

NOTE Confidence: 0.940355180416667

 $00:15:09.151 \longrightarrow 00:15:10.779$ with no residual disease,

NOTE Confidence: 0.940355180416667

 $00:15:10.780 \longrightarrow 00:15:13.948$ have done profoundly better.

 $00{:}15{:}13.950 \dashrightarrow 00{:}15{:}16.533$ You can see here the overall survival

NOTE Confidence: 0.940355180416667

 $00:15:16.533 \rightarrow 00:15:18.332$ is significantly better for women

NOTE Confidence: 0.940355180416667

 $00:15:18.332 \rightarrow 00:15:20.530$ who have had 0 millimeters of disease

NOTE Confidence: 0.940355180416667

 $00:15:20.530 \longrightarrow 00:15:22.669$ left at the completion of surgery.

NOTE Confidence: 0.940355180416667

 $00:15:22.670 \longrightarrow 00:15:23.708$ This is kind of what things

NOTE Confidence: 0.940355180416667

00:15:23.708 --> 00:15:24.670 look like when we begin.

NOTE Confidence: 0.940355180416667

 $00:15:24.670 \longrightarrow 00:15:27.630$ This is kind of how ovarian cancer presents.

NOTE Confidence: 0.940355180416667

 $00:15:27.630 \rightarrow 00:15:30.446$ This is the cancer that spreads over the

NOTE Confidence: 0.940355180416667

 $00{:}15{:}30{.}446 \dashrightarrow 00{:}15{:}33{.}819$ bowel or momentum over the pertinal surfaces.

NOTE Confidence: 0.940355180416667

 $00:15:33.820 \rightarrow 00:15:36.100$ And very frequently we hear especially

NOTE Confidence: 0.940355180416667

00:15:36.100 --> 00:15:38.500 Yale give new adjuvant chemotherapy,

NOTE Confidence: 0.940355180416667

 $00{:}15{:}38{.}500 \dashrightarrow 00{:}15{:}40{.}840$ which is chemotherapy before the surgery

NOTE Confidence: 0.940355180416667

 $00{:}15{:}40{.}840 \dashrightarrow 00{:}15{:}43{.}290$ to increase the chances of successful

NOTE Confidence: 0.940355180416667

 $00:15:43.290 \rightarrow 00:15:45.340$ debalking to no residual disease.

NOTE Confidence: 0.940355180416667

 $00:15:45.340 \longrightarrow 00:15:48.324$ So I want to talk briefly about approach

NOTE Confidence: 0.940355180416667

 $00:15:48.324 \rightarrow 00:15:51.607$ to surgery and again why this has to be

- NOTE Confidence: 0.940355180416667
- $00:15:51.607 \rightarrow 00:15:54.059$ so personalized and so individualized.
- NOTE Confidence: 0.940355180416667
- $00:15:54.060 \rightarrow 00:15:56.220$ This was a big poster that
- NOTE Confidence: 0.940355180416667
- $00:15:56.220 \longrightarrow 00:15:58.300$ was hanging in the office.
- NOTE Confidence: 0.940355180416667
- 00:15:58.300 --> 00:16:00.898 Of my mentor and previous boss.
- NOTE Confidence: 0.940355180416667
- 00:16:00.900 --> 00:16:02.700 And when I was younger,
- NOTE Confidence: 0.940355180416667
- $00:16:02.700 \longrightarrow 00:16:04.024$ not that much younger.
- NOTE Confidence: 0.940355180416667
- 00:16:04.024 --> 00:16:06.620 It was a big point of debate.
- NOTE Confidence: 0.940355180416667
- $00:16:06.620 \rightarrow 00:16:08.200$ The greater the surgeon,
- NOTE Confidence: 0.940355180416667
- $00:16:08.200 \longrightarrow 00:16:09.780$ the bigger the incision.
- NOTE Confidence: 0.940355180416667
- $00:16:09.780 \longrightarrow 00:16:11.940$ Certainly not the case.
- NOTE Confidence: 0.940355180416667
- 00:16:11.940 --> 00:16:14.019 Granted, I now have his office and his job,
- NOTE Confidence: 0.940355180416667
- $00{:}16{:}14.020 \dashrightarrow 00{:}16{:}17.770$ so there's that.
- NOTE Confidence: 0.940355180416667
- $00{:}16{:}17.770 \dashrightarrow 00{:}16{:}20.626$ So many studies have showed the
- NOTE Confidence: 0.940355180416667
- $00{:}16{:}20.626$ --> $00{:}16{:}22.530$ laparoscopy and minimally invasive
- NOTE Confidence: 0.940355180416667
- $00{:}16{:}22.607 \dashrightarrow 00{:}16{:}24.455$ approach to ovarian cancers
- NOTE Confidence: 0.940355180416667

00:16:24.455 --> 00:16:26.765 and gercologic cancers not only

NOTE Confidence: 0.940355180416667

 $00:16:26.765 \rightarrow 00:16:29.248$ safe but completely appropriate.

NOTE Confidence: 0.940355180416667

 $00:16:29.250 \rightarrow 00:16:32.010$ We know that complications are less,

NOTE Confidence: 0.940355180416667

 $00:16:32.010 \rightarrow 00:16:34.369$ We know that efficiency is the same.

NOTE Confidence: 0.940355180416667

 $00{:}16{:}34{.}370 \dashrightarrow 00{:}16{:}37{.}408$ We know that effectiveness is the same.

NOTE Confidence: 0.940355180416667

00:16:37.410 --> 00:16:37.996 So again,

NOTE Confidence: 0.940355180416667

 $00:16:37.996 \rightarrow 00:16:40.340$ everything has to be done in a very

NOTE Confidence: 0.940355180416667

 $00:16:40.405 \rightarrow 00:16:43.009$ personalized way that has to make sense,

NOTE Confidence: 0.940355180416667

 $00:16:43.010 \rightarrow 00:16:44.882$ the biggest thing in joint oncology

NOTE Confidence: 0.940355180416667

 $00:16:44.882 \rightarrow 00:16:46.489$ surgery and something that we

NOTE Confidence: 0.940355180416667

 $00:16:46.489 \longrightarrow 00:16:47.697$ care so passionately about.

NOTE Confidence: 0.940355180416667

 $00:16:47.700 \longrightarrow 00:16:49.828$ Is that masses do not get spilled

NOTE Confidence: 0.940355180416667

 $00:16:49.828 \longrightarrow 00:16:51.659$ because when the mass ruptures,

NOTE Confidence: 0.940355180416667

 $00:16:51.660 \rightarrow 00:16:53.590$ that's up upstages the patient

NOTE Confidence: 0.940355180416667

 $00:16:53.590 \longrightarrow 00:16:55.134$ and worsens their prognosis.

NOTE Confidence: 0.940355180416667

 $00:16:55.140 \longrightarrow 00:16:57.898$ So all kinds of things are done

- NOTE Confidence: 0.940355180416667
- $00:16:57.900 \longrightarrow 00:17:00.288$ to prevent that.
- NOTE Confidence: 0.940355180416667
- $00:17:00.288 \longrightarrow 00:17:00.876$ And again,
- NOTE Confidence: 0.940355180416667
- 00:17:00.876 --> 00:17:03.665 in the older days used to be a big point
- NOTE Confidence: 0.940355180416667
- $00:17:03.665 \rightarrow 00:17:05.820$ of debate and big point of discussion.
- NOTE Confidence: 0.940355180416667
- $00{:}17{:}05.820 \dashrightarrow 00{:}17{:}07.876$ So I want to share some of the
- NOTE Confidence: 0.940355180416667
- $00{:}17{:}07.876 \dashrightarrow 00{:}17{:}10.378$ videos of mine and some of our
- NOTE Confidence: 0.940355180416667
- $00{:}17{:}10.378 \dashrightarrow 00{:}17{:}13.740$ partners Sudo Zodi that show how we
- NOTE Confidence: 0.940355180416667
- $00{:}17{:}13.740 \dashrightarrow 00{:}17{:}16.884$ deal with some of these concerns.
- NOTE Confidence: 0.940355180416667
- 00:17:16.884 --> 00:17:19.920 So you know in ovarian cancer,
- NOTE Confidence: 0.940355180416667
- $00:17:19.920 \longrightarrow 00:17:21.520$ because our patients are women,
- NOTE Confidence: 0.940355180416667
- $00:17:21.520 \rightarrow 00:17:24.640$ we use frequently the vaginal
- NOTE Confidence: 0.940355180416667
- $00{:}17{:}24.640 \dashrightarrow 00{:}17{:}28.540$ orifice as a site of of removal.
- NOTE Confidence: 0.940355180416667
- $00:17:28.540 \rightarrow 00:17:34.300$ So this is a young patient who had Carson
- NOTE Confidence: 0.940355180416667
- $00{:}17{:}34{.}300 \dashrightarrow 00{:}17{:}38{.}935$ Sarcoma that had isolated periodic mass.
- NOTE Confidence: 0.940355180416667
- 00:17:38.935 --> 00:17:41.770 That you know in the older days,
- NOTE Confidence: 0.940355180416667

00:17:41.770 --> 00:17:44.535 again few years back would have just

NOTE Confidence: 0.940355180416667

00:17:44.535 - 00:17:47.314 been removed with a big incision,

NOTE Confidence: 0.940355180416667

 $00:17:47.314 \rightarrow 00:17:50.290$ with a huge resection and huge

NOTE Confidence: 0.940355180416667

 $00:17:50.391 \longrightarrow 00:17:52.090$ recovery because of the concern

NOTE Confidence: 0.940355180416667

 $00:17:52.090 \rightarrow 00:17:53.730$ both for the complexity of the case.

NOTE Confidence: 0.940355180416667

 $00:17:53.730 \rightarrow 00:17:56.570$ So as you can see this is the great vessels,

NOTE Confidence: 0.940355180416667

 $00:17:56.570 \longrightarrow 00:17:57.746$ this is the ureter,

NOTE Confidence: 0.940355180416667

 $00:17:57.746 \longrightarrow 00:18:00.134$ this is the mask that was entirely

NOTE Confidence: 0.940355180416667

 $00{:}18{:}00{.}134 \dashrightarrow 00{:}18{:}02{.}394$ resected and then the concern would have

NOTE Confidence: 0.940355180416667

 $00:18:02.394 \rightarrow 00:18:05.040$ been how do you remove a mask like this.

NOTE Confidence: 0.940355180416667

 $00:18:05.040 \longrightarrow 00:18:06.750$ And this is something that

NOTE Confidence: 0.940355180416667

 $00{:}18{:}06{.}750 \dashrightarrow 00{:}18{:}08{.}250$ we use a great deal.

NOTE Confidence: 0.940355180416667

 $00{:}18{:}08{.}250 \dashrightarrow 00{:}18{:}10{.}156$ And something that we talk about a

NOTE Confidence: 0.940355180416667

 $00:18:10.156 \rightarrow 00:18:12.380$ lot is that we're able to just play

NOTE Confidence: 0.940355180416667

 $00:18:12.452 \rightarrow 00:18:14.717$ something right through the vagina,

NOTE Confidence: 0.940355180416667

 $00:18:14.720 \longrightarrow 00:18:17.516$ place the back into the vagina

- NOTE Confidence: 0.940355180416667
- $00:18:17.520 \longrightarrow 00:18:20.880$ that is only very, very small,
- NOTE Confidence: 0.940355180416667
- $00:18:20.880 \longrightarrow 00:18:22.624$ small 10 millimeter incision
- NOTE Confidence: 0.940355180416667
- $00:18:22.624 \longrightarrow 00:18:25.240$ and then remove it like this.
- NOTE Confidence: 0.940355180416667
- $00:18:25.240 \longrightarrow 00:18:26.536$ And this patient was able to
- NOTE Confidence: 0.940355180416667
- $00{:}18{:}26{.}536 \dashrightarrow 00{:}18{:}27{.}400$ go home same day.
- NOTE Confidence: 0.911545551333333
- $00{:}18{:}27{.}400 \dashrightarrow 00{:}18{:}30{.}235$ This patient also had a metal diverticulum
- NOTE Confidence: 0.911545551333333
- $00{:}18{:}30{.}235 \dashrightarrow 00{:}18{:}33{.}516$ that we we resected at the same time.
- NOTE Confidence: 0.911545551333333
- 00:18:33.520 --> 00:18:35.650 And also was removed similarly
- NOTE Confidence: 0.911545551333333
- $00:18:35.650 \longrightarrow 00:18:38.200$ like that as you can see.
- NOTE Confidence: 0.911545551333333
- $00:18:38.200 \longrightarrow 00:18:40.314$ So this is the bowel surgery,
- NOTE Confidence: 0.911545551333333
- $00:18:40.320 \rightarrow 00:18:43.239$ the reticulum similarly placed in the bag,
- NOTE Confidence: 0.911545551333333
- $00:18:43.240 \longrightarrow 00:18:46.040$ similarly placed in the vagina
- NOTE Confidence: 0.911545551333333
- $00{:}18{:}46{.}040 \dashrightarrow 00{:}18{:}48{.}640$ and similarly patient went home
- NOTE Confidence: 0.911545551333333
- $00:18:48.640 \rightarrow 00:18:50.938$ same day and had minimal minimal
- NOTE Confidence: 0.911545551333333
- $00{:}18{:}50{.}938 \dashrightarrow 00{:}18{:}52{.}914$ recovery from from something that
- NOTE Confidence: 0.911545551333333

 $00:18:52.914 \rightarrow 00:18:54.799$ would have been very extensive.

NOTE Confidence: 0.911545551333333

00:18:54.800 --> 00:18:57.404 Natural or laparoscopy is so important

NOTE Confidence: 0.911545551333333

 $00{:}18{:}57{.}404 \dashrightarrow 00{:}19{:}01{.}035$ and again we are lucky to be able to use

NOTE Confidence: 0.911545551333333

 $00:19:01.040 \rightarrow 00:19:04.246$ vaginal orifice in our patients for this.

NOTE Confidence: 0.911545551333333

 $00:19:04.250 \longrightarrow 00:19:06.488$ So at times, again, you know,

NOTE Confidence: 0.911545551333333

 $00{:}19{:}06{.}490 \dashrightarrow 00{:}19{:}07{.}650$ everything has to make sense.

NOTE Confidence: 0.911545551333333

 $00{:}19{:}07.650 \dashrightarrow 00{:}19{:}09.922$ You know, this is the older patient who

NOTE Confidence: 0.911545551333333

 $00:19:09.922 \rightarrow 00:19:12.822$ had a very many medical comorbidities

NOTE Confidence: 0.911545551333333

 $00{:}19{:}12.822 \dashrightarrow 00{:}19{:}15.737$ who would not have with standed.

NOTE Confidence: 0.911545551333333

00:19:15.740 --> 00:19:17.112 Overall, big large laparotomy,

NOTE Confidence: 0.911545551333333

 $00{:}19{:}17{.}112 \dashrightarrow 00{:}19{:}19{.}579$ which is what we would have done

NOTE Confidence: 0.911545551333333

 $00:19:19.579 \rightarrow 00:19:21.655$ usually for this 12 centimeter mass.

NOTE Confidence: 0.911545551333333

 $00:19:21.660 \longrightarrow 00:19:22.860$ But at the same time,

NOTE Confidence: 0.911545551333333

 $00:19:22.860 \rightarrow 00:19:24.948$ we of course want to be very careful and

NOTE Confidence: 0.911545551333333

 $00:19:24.948 \rightarrow 00:19:26.980$ not rupture the mass and not leak it.

NOTE Confidence: 0.911545551333333

 $00:19:26.980 \longrightarrow 00:19:29.708$ So we do all kinds of special techniques

 $00:19:29.708 \rightarrow 00:19:32.239$ again in a very truly personalized way

NOTE Confidence: 0.911545551333333

 $00{:}19{:}32{.}239 \dashrightarrow 00{:}19{:}35{.}341$ to be able to provide our patients with

NOTE Confidence: 0.911545551333333

 $00:19:35.341 \longrightarrow 00:19:37.849$ the best care specifically for them.

NOTE Confidence: 0.911545551333333

 $00:19:37.850 \longrightarrow 00:19:38.585$ So this patient,

NOTE Confidence: 0.911545551333333

00:19:38.585 --> 00:19:39.565 as you can see,

NOTE Confidence: 0.911545551333333

 $00{:}19{:}39{.}570 \dashrightarrow 00{:}19{:}41{.}110$ everything was protected and then

NOTE Confidence: 0.911545551333333

 $00{:}19{:}41{.}110 \dashrightarrow 00{:}19{:}43{.}203$ the mask was insulated and we were

NOTE Confidence: 0.911545551333333

 $00:19:43.203 \longrightarrow 00:19:44.685$ actually able to see inside the

NOTE Confidence: 0.911545551333333

 $00{:}19{:}44.685 \dashrightarrow 00{:}19{:}46.461$ mask and then we were able to see

NOTE Confidence: 0.911545551333333

 $00{:}19{:}46{.}461 \dashrightarrow 00{:}19{:}48{.}106$ that this was a malignant mask.

NOTE Confidence: 0.911545551333333

 $00:19:48.106 \longrightarrow 00:19:50.962$ Once the fluid was removed in the

NOTE Confidence: 0.911545551333333

 $00:19:50.962 \rightarrow 00:19:53.725$ contained way and that nothing was removed,

NOTE Confidence: 0.911545551333333

 $00{:}19{:}53.730 \dashrightarrow 00{:}19{:}55.970$ nothing was nothing leaked.

NOTE Confidence: 0.911545551333333

 $00{:}19{:}55{.}970 \dashrightarrow 00{:}19{:}59{.}130$ And then same same way we were able

NOTE Confidence: 0.911545551333333

 $00{:}19{:}59{.}130 \dashrightarrow 00{:}20{:}01{.}312$ to just tie something around it

 $00:20:01.312 \longrightarrow 00:20:03.544$ to make sure that nothing leaks.

NOTE Confidence: 0.911545551333333

 $00:20:03.550 \rightarrow 00:20:05.223$ Remove it and then remove it similarly

NOTE Confidence: 0.911545551333333

00:20:05.223 --> 00:20:07.550 to the vagina like I showed you before

NOTE Confidence: 0.911545551333333

 $00:20:07.550 \rightarrow 00:20:09.083$ and all the lymph nodes and everything

NOTE Confidence: 0.911545551333333

 $00:20:09.083 \rightarrow 00:20:10.709$ else could be done laparoscopically.

NOTE Confidence: 0.911545551333333

 $00{:}20{:}10.710 \dashrightarrow 00{:}20{:}13.326$ And again the spatial with multiple

NOTE Confidence: 0.911545551333333

 $00{:}20{:}13.326 \dashrightarrow 00{:}20{:}16.206$ comorbidities was able to go home

NOTE Confidence: 0.911545551333333

 $00:20:16.206 \rightarrow 00:20:19.642$ next morning with minimal recovery

NOTE Confidence: 0.911545551333333

 $00:20:19.642 \longrightarrow 00:20:23.708$ time and was able to have her

NOTE Confidence: 0.911545551333333

00:20:23.708 - > 00:20:25.307 cancer appropriately removed.

NOTE Confidence: 0.911545551333333

 $00:20:25.310 \rightarrow 00:20:27.305$ We nowadays use something called high pack,

NOTE Confidence: 0.911545551333333

 $00:20:27.310 \rightarrow 00:20:30.366$ which is what I mentioned to you before.

NOTE Confidence: 0.911545551333333

 $00:20:30.370 \longrightarrow 00:20:33.667$ Which is there's a lot of conceptual

NOTE Confidence: 0.911545551333333

 $00{:}20{:}33.667 \dashrightarrow 00{:}20{:}35.735$ thinking that hyper hypothermia

NOTE Confidence: 0.911545551333333

 $00{:}20{:}35{.}735 \dashrightarrow 00{:}20{:}39{.}161$ and neoplasia at the time of

NOTE Confidence: 0.911545551333333

00:20:39.161 --> 00:20:41.597 aggressive debalking really improves

- NOTE Confidence: 0.911545551333333
- 00:20:41.597 --> 00:20:43.487 the overall survival.
- NOTE Confidence: 0.911545551333333
- $00:20:43.490 \longrightarrow 00:20:45.290$ And this is again, this is before debalking.
- NOTE Confidence: 0.911545551333333
- $00:20:45.290 \longrightarrow 00:20:46.702$ This is after debalking,
- NOTE Confidence: 0.911545551333333
- $00{:}20{:}46.702 \dashrightarrow 00{:}20{:}49.186$ we remove all the disease and then
- NOTE Confidence: 0.911545551333333
- $00{:}20{:}49{.}186 \dashrightarrow 00{:}20{:}51{.}770$ we're able to place a cyst plan into
- NOTE Confidence: 0.911545551333333
- $00:20:51.770 \longrightarrow 00:20:55.410$ the belly for 90 minutes that improves
- NOTE Confidence: 0.911545551333333
- $00:20:55.410 \rightarrow 00:21:00.180$ direct permeation of the tumors.
- NOTE Confidence: 0.911545551333333
- $00:21:00.180 \longrightarrow 00:21:01.552$ And there's some very,
- NOTE Confidence: 0.911545551333333
- $00:21:01.552 \rightarrow 00:21:03.267$ very strong literature that supports
- NOTE Confidence: 0.911545551333333
- $00:21:03.267 \rightarrow 00:21:05.380$ it in this population with pretty,
- NOTE Confidence: 0.911545551333333
- 00:21:05.380 --> 00:21:07.120 pretty significant improvement
- NOTE Confidence: 0.911545551333333
- $00{:}21{:}07{.}120 \dashrightarrow 00{:}21{:}09{.}440$ and progression free survival
- NOTE Confidence: 0.911545551333333
- $00:21:09.440 \longrightarrow 00:21:11.180$ and overall survival.
- NOTE Confidence: 0.911545551333333
- 00:21:11.180 --> 00:21:13.052 In the next two minutes that I have left,
- NOTE Confidence: 0.911545551333333
- 00:21:13.060 --> 00:21:14.210 I promised I consenting that
- NOTE Confidence: 0.911545551333333

 $00:21:14.210 \rightarrow 00:21:15.819$ I was going to be on time.

NOTE Confidence: 0.911545551333333

00:21:15.820 --> 00:21:17.934 I want to talk briefly about survivorship.

NOTE Confidence: 0.911545551333333

00:21:17.940 --> 00:21:19.860 So survivorship is so important.

NOTE Confidence: 0.911545551333333

 $00:21:19.860 \rightarrow 00:21:22.440$ We for generations have undervalued

NOTE Confidence: 0.911545551333333

 $00{:}21{:}22{.}440 \dashrightarrow 00{:}21{:}25{.}020$ under address the importance of

NOTE Confidence: 0.911545551333333

 $00{:}21{:}25.103 \dashrightarrow 00{:}21{:}27.567$ survivorship in this population.

NOTE Confidence: 0.911545551333333

00:21:27.570 --> 00:21:28.710 It's because forever we,

NOTE Confidence: 0.911545551333333

 $00:21:28.710 \longrightarrow 00:21:30.420$ we thought the women get this

NOTE Confidence: 0.911545551333333

 $00:21:30.476 \longrightarrow 00:21:31.846$ cancers and then they die.

NOTE Confidence: 0.911545551333333

 $00:21:31.850 \longrightarrow 00:21:33.050$ That is not the case.

NOTE Confidence: 0.911545551333333

 $00{:}21{:}33.050 \dashrightarrow 00{:}21{:}35.482$ As you see that as we're getting better

NOTE Confidence: 0.911545551333333

00:21:35.482 --> 00:21:37.510 in diagnosis is we're getting better

NOTE Confidence: 0.911545551333333

 $00:21:37.510 \rightarrow 00:21:39.544$ in surgery as we're getting better

NOTE Confidence: 0.871449994090909

00:21:39.609 --> 00:21:41.156 in treatment, women are living

NOTE Confidence: 0.871449994090909

 $00:21:41.156 \rightarrow 00:21:42.728$ longer and when they live longer,

NOTE Confidence: 0.871449994090909

 $00:21:42.730 \rightarrow 00:21:45.898$ it is so important that we provide them with

 $00:21:45.898 \rightarrow 00:21:47.987$ survivorship techniques to support them.

NOTE Confidence: 0.871449994090909

 $00:21:47.990 \longrightarrow 00:21:50.118$ I'm actually so happy to see that we

NOTE Confidence: 0.871449994090909

 $00:21:50.118 \longrightarrow 00:21:52.742$ have our teal team here that help

NOTE Confidence: 0.871449994090909

 $00:21:52.742 \longrightarrow 00:21:54.982$ us concentrate on nutrition this

NOTE Confidence: 0.871449994090909

 $00:21:54.982 \longrightarrow 00:21:57.200$ population because this is yet another

NOTE Confidence: 0.871449994090909

 $00:21:57.200 \rightarrow 00:22:00.092$ one of those aspects that really has

NOTE Confidence: 0.871449994090909

 $00:22:00.092 \longrightarrow 00:22:02.188$ been undervalued and underaddressed.

NOTE Confidence: 0.871449994090909

 $00:22:02.190 \longrightarrow 00:22:06.794$ So in in Galactic Cancers survivorship

NOTE Confidence: 0.871449994090909

 $00:22:06.794 \rightarrow 00:22:09.949$ has so many different aspects,

NOTE Confidence: 0.871449994090909

00:22:09.950 --> 00:22:12.710 fear recurrence, toxicities from surgery,

NOTE Confidence: 0.871449994090909

 $00{:}22{:}12.710 \dashrightarrow 00{:}22{:}14.618$ toxicities of chemo and radiation and

NOTE Confidence: 0.871449994090909

 $00{:}22{:}14.618$ --> $00{:}22{:}16.989$ that's what they got financial toxicity.

NOTE Confidence: 0.871449994090909

 $00:22:16.990 \longrightarrow 00:22:18.550$ Psychological body image and

NOTE Confidence: 0.871449994090909

 $00:22:18.550 \longrightarrow 00:22:20.110$ then sex and intimacy,

NOTE Confidence: 0.871449994090909

 $00{:}22{:}20{.}110 \dashrightarrow 00{:}22{:}23{.}702$ which is a very profound aspect of women's

 $00:22:23.702 \longrightarrow 00:22:26.733$ experiences that here at Yale we are

NOTE Confidence: 0.871449994090909

 $00{:}22{:}26.733 \dashrightarrow 00{:}22{:}29.262$ very passionate about the dressing so

NOTE Confidence: 0.871449994090909

 $00:22:29.262 \longrightarrow 00:22:32.718$ I'll let this I'll leave this with Doctor NOTE Confidence: 0.871449994090909

 $00:22:32.718 \rightarrow 00:22:36.110$ Sentine to with one minute to spare.

NOTE Confidence: 0.871449994090909

00:22:36.110 --> 00:22:37.940 So in the one minute then I have I'm not

NOTE Confidence: 0.871449994090909

00:22:37.995 --> 00:22:39.894 going to I'm not going to give away the NOTE Confidence: 0.871449994090909

 $00:22:39.894 \rightarrow 00:22:42.786$ minute what separates us here at Yale.

NOTE Confidence: 0.871449994090909

 $00:22:42.790 \longrightarrow 00:22:45.198$ Is that we are truly patient driven

NOTE Confidence: 0.871449994090909

 $00:22:45.198 \longrightarrow 00:22:48.286$ and we are truly believe in the

NOTE Confidence: 0.871449994090909

 $00:22:48.286 \rightarrow 00:22:50.266$ importance of personalized care.

NOTE Confidence: 0.871449994090909

 $00:22:50.270 \longrightarrow 00:22:51.726$ You know, we nobody,

NOTE Confidence: 0.871449994090909

 $00{:}22{:}51.726 \dashrightarrow 00{:}22{:}53.546$ nobody treats the cancer anymore.

NOTE Confidence: 0.871449994090909

 $00{:}22{:}53.550 \dashrightarrow 00{:}22{:}56.150$ We truly treat the women that we see.

NOTE Confidence: 0.871449994090909

 $00:22:56.150 \rightarrow 00:22:57.362$ We see their lives,

NOTE Confidence: 0.871449994090909

 $00{:}22{:}57{.}362 \dashrightarrow 00{:}22{:}59{.}579$ we see their experiences and then we do.

NOTE Confidence: 0.871449994090909

 $00:22:59.580 \rightarrow 00:23:02.172$ Everything we can to address their

 $00{:}23{:}02{.}172 \dashrightarrow 00{:}23{:}04{.}301$ personal experience from diagnosis to

NOTE Confidence: 0.871449994090909

 $00{:}23{:}04{.}301 \dashrightarrow 00{:}23{:}06{.}667$ surgery to treatment and we'll we'll talk

NOTE Confidence: 0.871449994090909

 $00{:}23{:}06.667 \dashrightarrow 00{:}23{:}09.472$ about and then to the survivorship and

NOTE Confidence: 0.871449994090909

 $00:23:09.472 \rightarrow 00:23:11.537$ supporting them through their journey

NOTE Confidence: 0.871449994090909

 $00:23:11.540 \longrightarrow 00:23:13.616$ both during treatment and then after.

NOTE Confidence: 0.871449994090909

 $00:23:13.620 \longrightarrow 00:23:15.452$ And again we're only able to do it

NOTE Confidence: 0.871449994090909

 $00:23:15.452 \rightarrow 00:23:17.498$ because of the amazing team that we have.

NOTE Confidence: 0.871449994090909

 $00:23:17.500 \longrightarrow 00:23:17.820$ Thank you

NOTE Confidence: 0.8023952

 $00:23:27.020 \longrightarrow 00:23:31.650$ so. Alina has already done a very

NOTE Confidence: 0.8023952

 $00:23:31.650 \longrightarrow 00:23:33.860$ nice introduction and this light

NOTE Confidence: 0.8023952

 $00:23:33.860 \longrightarrow 00:23:36.612$ for me is only to remind you that

NOTE Confidence: 0.8023952

00:23:36.620 --> 00:23:39.175 regardless to the how sophisticated

NOTE Confidence: 0.8023952

00:23:39.175 --> 00:23:42.647 our surgery can be and Alina show

NOTE Confidence: 0.8023952

00:23:42.647 --> 00:23:44.332 you minimally invasive approaches,

NOTE Confidence: 0.8023952

 $00:23:44.332 \rightarrow 00:23:47.300$ sometimes we have to do ultra radical NOTE Confidence: 0.8023952

 $00{:}23{:}47{.}371 \dashrightarrow 00{:}23{:}49{.}618$ surgery to to remove again the tumor NOTE Confidence: 0.8023952 $00:23:49.620 \rightarrow 00:23:51.328$ regardless regardless the adjuvant NOTE Confidence: 0.8023952 $00{:}23{:}51{.}328 \dashrightarrow 00{:}23{:}53{.}890$ treatment they were able to offer NOTE Confidence: 0.8023952 $00:23:53.961 \longrightarrow 00:23:56.169$ to this patient such as radiation NOTE Confidence: 0.8023952 $00:23:56.169 \rightarrow 00:23:58.360$ therapy as well as chemotherapy. NOTE Confidence: 0.8023952 00:23:58.360 --> 00:24:00.635 A significant number of our NOTE Confidence: 0.8023952 $00{:}24{:}00{.}635 \dashrightarrow 00{:}24{:}02{.}902$ patients still again develop NOTE Confidence: 0.8023952 $00:24:02.902 \rightarrow 00:24:05.206$ progression and recurrent disease. NOTE Confidence: 0.8023952 $00{:}24{:}05{.}210 \dashrightarrow 00{:}24{:}06{.}650$ And as I show you here, NOTE Confidence: 0.8023952 $00:24:06.650 \rightarrow 00:24:09.434$ of the 100,000 diagnosis with a NOTE Confidence: 0.8023952 00:24:09.434 --> 00:24:11.610 gynecological cancer in the US, NOTE Confidence: 0.8023952 $00:24:11.610 \rightarrow 00:24:14.810$ about 1/3 of women still die every year NOTE Confidence: 0.8023952 $00{:}24{:}14.810 \dashrightarrow 00{:}24{:}18.290$ because of this difficult to treat cancer. NOTE Confidence: 0.8023952 $00:24:18.290 \rightarrow 00:24:22.520$ So the question is what can we do to NOTE Confidence: 0.8023952 $00:24:22.520 \rightarrow 00:24:25.260$ improve the outcome of this patient? NOTE Confidence: 0.8023952 00:24:25.260 --> 00:24:27.868 And this again and what I'm going to

- NOTE Confidence: 0.8023952
- $00:24:27.868 \rightarrow 00:24:30.609$ show you here is our scientific vision
- NOTE Confidence: 0.8023952
- $00{:}24{:}30{.}609 \dashrightarrow 00{:}24{:}33{.}910$ and goals that we really we we we
- NOTE Confidence: 0.8023952
- $00:24:33.910 \longrightarrow 00:24:36.522$ started thinking about how we could
- NOTE Confidence: 0.8023952
- $00:24:36.522 \rightarrow 00:24:39.978$ really face up with some medical need many,
- NOTE Confidence: 0.8023952
- $00{:}24{:}39{.}980 \dashrightarrow 00{:}24{:}42{.}856$ many years ago and we developed
- NOTE Confidence: 0.8023952
- $00:24:42.856 \longrightarrow 00:24:45.180$ or we tried to develop a program,
- NOTE Confidence: 0.8023952
- $00:24:45.180 \rightarrow 00:24:47.208$ a personalized treatment program.
- NOTE Confidence: 0.8023952
- $00:24:47.208 \longrightarrow 00:24:50.250$ Adding to what Alina show you,
- NOTE Confidence: 0.8023952
- $00{:}24{:}50{.}250 \dashrightarrow 00{:}24{:}53{.}925$ so this excellent clinical and surgical care,
- NOTE Confidence: 0.8023952
- 00:24:53.930 --> 00:24:56.555 adding a translational research program
- NOTE Confidence: 0.8023952
- 00:24:56.555 --> 00:25:00.454 above that in particular targeting the
- NOTE Confidence: 0.8023952
- $00:25:00.454 \rightarrow 00:25:04.009$ difficult to treat gynecological cancer.
- NOTE Confidence: 0.8023952
- $00{:}25{:}04.010 \dashrightarrow 00{:}25{:}06.378$ And when I say difficult to treat it
- NOTE Confidence: 0.8023952
- $00{:}25{:}06{.}378 \dashrightarrow 00{:}25{:}08{.}600$ means that the problem is really in
- NOTE Confidence: 0.8023952
- $00{:}25{:}08.600 \dashrightarrow 00{:}25{:}11.125$ our specialties not to take care of
- NOTE Confidence: 0.8023952

 $00:25:11.125 \rightarrow 00:25:13.185$ a well or moderately differentiated

NOTE Confidence: 0.8023952

 $00:25:13.185 \longrightarrow 00:25:14.009$ uterine cancer.

NOTE Confidence: 0.8023952

 $00:25:14.010 \longrightarrow 00:25:16.537$ Is really to take care of the

NOTE Confidence: 0.8023952

 $00{:}25{:}16.537 \dashrightarrow 00{:}25{:}18.039$ biologically aggressive such as

NOTE Confidence: 0.8023952

 $00{:}25{:}18.039 \dashrightarrow 00{:}25{:}20.133$ the uterine serous carcinoma or the

NOTE Confidence: 0.8023952

 $00{:}25{:}20{.}133 \dashrightarrow 00{:}25{:}22{.}218$ carcinos or comma or they grade

NOTE Confidence: 0.8023952

 $00{:}25{:}22{.}218 \dashrightarrow 00{:}25{:}24{.}486$ ovarian that are wides pread and so on.

NOTE Confidence: 0.8023952

 $00:25:24.490 \longrightarrow 00:25:28.231$ So we thought to really the one of

NOTE Confidence: 0.8023952

 $00{:}25{:}28{.}231 \dashrightarrow 00{:}25{:}30{.}972$ the of the the the winning strategies NOTE Confidence: 0.8023952

 $00{:}25{:}30{.}972 \dashrightarrow 00{:}25{:}34{.}130$ was to provide access to our patient NOTE Confidence: 0.8023952

00:25:34.130 --> 00:25:36.902 in particular this group of patient

NOTE Confidence: 0.8023952

 $00{:}25{:}36{.}902 \dashrightarrow 00{:}25{:}39{.}191$ to precision medicine through a

NOTE Confidence: 0.8023952

 $00{:}25{:}39{.}191 \dashrightarrow 00{:}25{:}41{.}405$ pipeline of both pharma as well

NOTE Confidence: 0.8023952

 $00{:}25{:}41{.}405 \dashrightarrow 00{:}25{:}43{.}300$ as yield developed treatment.

NOTE Confidence: 0.8023952

 $00{:}25{:}43{.}300 \dashrightarrow 00{:}25{:}45{.}010$ Again validated within rigorous clinical

NOTE Confidence: 0.8023952

 $00:25:45.010 \rightarrow 00:25:47.897$ trial that I'm going to show you in a minute.

- NOTE Confidence: 0.8023952
- $00{:}25{:}47{.}900 \dashrightarrow 00{:}25{:}50{.}716$ But the other thing was really to to
- NOTE Confidence: 0.8023952
- $00:25:50.716 \rightarrow 00:25:52.922$ be able to build a multidisciplinary
- NOTE Confidence: 0.8023952
- 00:25:52.922 --> 00:25:55.659 approach because if you want to be
- NOTE Confidence: 0.8023952
- $00:25:55.659 \rightarrow 00:25:57.500$ successful in a big Cancer Center,
- NOTE Confidence: 0.8023952
- $00{:}25{:}57{.}500 \dashrightarrow 00{:}25{:}59{.}620$ again the surgical is important,
- NOTE Confidence: 0.8023952
- $00:25:59.620 \longrightarrow 00:26:00.366$ is important,
- NOTE Confidence: 0.8023952
- $00:26:00.366 \longrightarrow 00:26:01.112$ the the,
- NOTE Confidence: 0.8023952
- $00:26:01.112 \longrightarrow 00:26:02.977$ the chemotherapy or addition therapy
- NOTE Confidence: 0.8023952
- $00:26:02.977 \longrightarrow 00:26:05.114$ is important by if you want to
- NOTE Confidence: 0.8023952
- $00{:}26{:}05{.}114 \dashrightarrow 00{:}26{:}07{.}784$ do one step farther and going to
- NOTE Confidence: 0.8023952
- 00:26:07.784 --> 00:26:09.260 translational personalized medicine.
- NOTE Confidence: 0.8023952
- $00:26:09.260 \longrightarrow 00:26:11.288$ This is really a team approach
- NOTE Confidence: 0.8023952
- $00{:}26{:}11.288 \dashrightarrow 00{:}26{:}13.290$ that goes outside our division.
- NOTE Confidence: 0.8023952
- $00{:}26{:}13.290 \dashrightarrow 00{:}26{:}15.207$ And that is what we try to build here.
- NOTE Confidence: 0.8023952
- $00{:}26{:}15{.}210 \dashrightarrow 00{:}26{:}18{.}480$ We involve and interact with other
- NOTE Confidence: 0.8023952

 $00:26:18.480 \longrightarrow 00:26:21.758$ successful clinical as well as the

NOTE Confidence: 0.8023952

 $00:26:21.758 \rightarrow 00:26:24.474$ research program present in our Yale

NOTE Confidence: 0.8023952

 $00{:}26{:}24{.}474 \dashrightarrow 00{:}26{:}27{.}555$ Cancer Center and also of course take NOTE Confidence: 0.8023952

 $00{:}26{:}27{.}555 \dashrightarrow 00{:}26{:}30{.}075$ advantage of the cutting edge resources

NOTE Confidence: 0.8023952

 $00:26:30.075 \longrightarrow 00:26:32.445$ that we have available at Yale.

NOTE Confidence: 0.8023952

 $00{:}26{:}32{.}450 \dashrightarrow 00{:}26{:}35{.}915$ And I want to mention the the

NOTE Confidence: 0.8023952

 $00:26:35.915 \rightarrow 00:26:37.691$ genomic course sequencing facility

NOTE Confidence: 0.8023952

00:26:37.691 - > 00:26:40.680 that we have in the West Campus.

NOTE Confidence: 0.8023952

 $00{:}26{:}40.680 \dashrightarrow 00{:}26{:}43.440$ As well as again the the pathology corps,

NOTE Confidence: 0.8023952

 $00{:}26{:}43.440 \dashrightarrow 00{:}26{:}45.568$ Doctor RIM is here that is doing an

NOTE Confidence: 0.8023952

 $00{:}26{:}45{.}568 \dashrightarrow 00{:}26{:}47{.}241$ excellent job is providing tissue to

NOTE Confidence: 0.8023952

 $00{:}26{:}47{.}241 \dashrightarrow 00{:}26{:}49{.}407$ to all the research area of the Yale

NOTE Confidence: 0.8023952

 $00{:}26{:}49{.}407 \dashrightarrow 00{:}26{:}51{.}319$ Cancer Center to be able to do what

NOTE Confidence: 0.93421556

 $00{:}26{:}51{.}320 \dashrightarrow 00{:}26{:}54{.}850$ I'm going to show you internal

NOTE Confidence: 0.93421556

 $00{:}26{:}54.850 \dashrightarrow 00{:}26{:}56.870$ personalized treatment.

NOTE Confidence: 0.93421556

 $00:26:56.870 \longrightarrow 00:27:00.302$ So a lot of people talk about personalized

- NOTE Confidence: 0.93421556
- $00{:}27{:}00{.}302 \dashrightarrow 00{:}27{:}02{.}960$ treatment but what is personalized treatment,

 $00:27:02.960 \longrightarrow 00:27:05.834$ what is precision medicine in the

NOTE Confidence: 0.93421556

00:27:05.834 --> 00:27:08.180 division of gynecological oncology Yale.

NOTE Confidence: 0.93421556

00:27:08.180 --> 00:27:09.972 As Alina show you,

NOTE Confidence: 0.93421556

 $00{:}27{:}09{.}972 \dashrightarrow 00{:}27{:}12.660$ everything start with the surgical part.

NOTE Confidence: 0.93421556

 $00{:}27{:}12.660 \dashrightarrow 00{:}27{:}15.936$ So we are surgeon but ovarian uterine

NOTE Confidence: 0.93421556

 $00:27:15.936 \rightarrow 00:27:18.459$ cancer are surgically staged tumor.

NOTE Confidence: 0.93421556

 $00:27:18.460 \longrightarrow 00:27:20.140$ So we that is the first time,

NOTE Confidence: 0.93421556

 $00:27:20.140 \longrightarrow 00:27:21.960$ the first moment in the

NOTE Confidence: 0.93421556

 $00:27:21.960 \longrightarrow 00:27:23.416$ treatment of our patient.

NOTE Confidence: 0.93421556

00:27:23.420 --> 00:27:24.620 And as Alina told you,

NOTE Confidence: 0.93421556

 $00{:}27{:}24.620 \dashrightarrow 00{:}27{:}27.550$ we try to remove all the disease that we can

NOTE Confidence: 0.93421556

 $00:27:27.620 \rightarrow 00:27:29.696$ see to improve the outcome of this patient.

NOTE Confidence: 0.93421556

 $00{:}27{:}29{.}700 \dashrightarrow 00{:}27{:}32{.}092$ But one of the thing that we've been

NOTE Confidence: 0.93421556

 $00{:}27{:}32.092 \dashrightarrow 00{:}27{:}34.188$ doing now for quite a long time.

00:27:34.190 --> 00:27:36.612 Starting probably as one of the First

NOTE Confidence: 0.93421556

 $00{:}27{:}36{.}612 \dashrightarrow 00{:}27{:}38{.}628$ Division in the US and worldwide

NOTE Confidence: 0.93421556

 $00{:}27{:}38{.}628 \dashrightarrow 00{:}27{:}41{.}330$ was to consent our patient at the NOTE Confidence: 0.93421556

00:27:41.330 - 00:27:43.430 time of surgery before surgery

NOTE Confidence: 0.93421556

 $00{:}27{:}43.430 \dashrightarrow 00{:}27{:}46.167$ also to donate a piece of tissue

NOTE Confidence: 0.93421556

 $00{:}27{:}46.167 \dashrightarrow 00{:}27{:}49.629$ to go to our research laboratory NOTE Confidence: 0.93421556

 $00{:}27{:}49{.}630 \dashrightarrow 00{:}27{:}52{.}598$ and this piece of tissue was used

NOTE Confidence: 0.93421556

 $00:27:52.598 \longrightarrow 00:27:55.030$ to sequence to sequence the

NOTE Confidence: 0.930604272727273

 $00{:}27{:}57{.}310 \dashrightarrow 00{:}27{:}59{.}314$ DNA of the tumor of this

NOTE Confidence: 0.930604272727273

 $00{:}27{:}59{.}314 \dashrightarrow 00{:}28{:}01{.}060$ patient with the ultimate goal.

NOTE Confidence: 0.930604272727273

 $00{:}28{:}01{.}060 \dashrightarrow 00{:}28{:}04{.}175$ To have a better understanding of the

NOTE Confidence: 0.930604272727273

 $00:28:04.175 \longrightarrow 00:28:06.624$ genetic landscape of that specific

NOTE Confidence: 0.930604272727273

 $00{:}28{:}06{.}624 \dashrightarrow 00{:}28{:}09{.}700$ patient tumor and as I told you

NOTE Confidence: 0.930604272727273

 $00{:}28{:}09{.}700 \dashrightarrow 00{:}28{:}12{.}780$ we started close to 2010 to do that.

NOTE Confidence: 0.930604272727273

 $00:28:12.780 \longrightarrow 00:28:15.934$ At that time point the commercial entity

NOTE Confidence: 0.930604272727273

 $00:28:15.934 \rightarrow 00:28:19.737$ that we were using to receive a some

 $00{:}28{:}19.737 \dashrightarrow 00{:}28{:}22.992$ sort of FDAGMP approved report was

NOTE Confidence: 0.930604272727273

 $00{:}28{:}22{.}992 \dashrightarrow 00{:}28{:}26{.}200$ Foundation Medicine but at the same time

NOTE Confidence: 0.930604272727273

 $00:28:26.200 \rightarrow 00:28:29.649$ in parallel we are also able to perform.

NOTE Confidence: 0.930604272727273

 $00:28:29.650 \rightarrow 00:28:33.050$ All Axon sequencing at the West campus,

NOTE Confidence: 0.930604272727273

00:28:33.050 --> 00:28:35.690 as you know all Axon sequencing

NOTE Confidence: 0.930604272727273

 $00:28:35.690 \rightarrow 00:28:37.490$ is quite expensive,

NOTE Confidence: 0.930604272727273

00:28:37.490 --> 00:28:39.919 was very expensive about 10-15 years ago

NOTE Confidence: 0.930604272727273

 $00:28:39.919 \longrightarrow 00:28:42.448$ when we started is still expensive now.

NOTE Confidence: 0.930604272727273

 $00{:}28{:}42{.}450 \dashrightarrow 00{:}28{:}44{.}810$ So the question that some of you can ask me,

NOTE Confidence: 0.930604272727273

 $00{:}28{:}44{.}810 \dashrightarrow 00{:}28{:}46{.}819$ but there's something how have you been

NOTE Confidence: 0.930604272727273

 $00:28:46.819 \longrightarrow 00:28:49.200$ able to do that 100 or gynecological

NOTE Confidence: 0.930604272727273

00:28:49.200 --> 00:28:51.250 cancer patient without asking you

NOTE Confidence: 0.930604272727273

 $00:28:51.250 \rightarrow 00:28:53.567$ know the patient to pay for it.

NOTE Confidence: 0.930604272727273

 $00{:}28{:}53{.}570 \dashrightarrow 00{:}28{:}57{.}240$ The answer is that we.

NOTE Confidence: 0.930604272727273

00:28:57.240 --> 00:28:58.138 You know,

 $00:28:58.138 \rightarrow 00:29:00.832$ we had this collaborative research program

NOTE Confidence: 0.930604272727273

 $00{:}29{:}00{.}832 \dashrightarrow 00{:}29{:}04{.}200$ at Yale between Yale and Gilead Science.

NOTE Confidence: 0.930604272727273

00:29:04.200 --> 00:29:05.718 And this was a huge program,

NOTE Confidence: 0.930604272727273

 $00:29:05.720 \longrightarrow 00:29:09.859$ \$70 million in 10 years provided

NOTE Confidence: 0.930604272727273

 $00{:}29{:}09{.}859 \dashrightarrow 00{:}29{:}11{.}954$ to us through Doctor Schlesinger

NOTE Confidence: 0.930604272727273

 $00:29:11.954 \longrightarrow 00:29:14.156$ was the Chair of pharmacology

NOTE Confidence: 0.930604272727273

 $00{:}29{:}14.156 \dashrightarrow 00{:}29{:}16.880$ that was able to establish this

NOTE Confidence: 0.930604272727273

 $00:29:16.880 \longrightarrow 00:29:18.314$ collaborative research agreement.

NOTE Confidence: 0.930604272727273

 $00{:}29{:}18{.}314 \dashrightarrow 00{:}29{:}20{.}704$ And thanks to this significant

NOTE Confidence: 0.930604272727273

00:29:20.704 --> 00:29:22.120 amount of money,

NOTE Confidence: 0.930604272727273

 $00{:}29{:}22{.}120 \dashrightarrow 00{:}29{:}24{.}142$ we were able again to sequence

NOTE Confidence: 0.930604272727273

 $00:29:24.142 \longrightarrow 00:29:26.853$ over 800 gynecological cancer.

NOTE Confidence: 0.930604272727273

00:29:26.853 --> 00:29:31.310 Again all Axon sequencing, 21,000 gene,

NOTE Confidence: 0.930604272727273

 $00:29:31.310 \longrightarrow 00:29:34.535$ all the gene encoded protein.

NOTE Confidence: 0.930604272727273

 $00{:}29{:}34{.}540 \dashrightarrow 00{:}29{:}36{.}976$ Another thing that we have been doing

NOTE Confidence: 0.930604272727273

 $00:29:36.976 \longrightarrow 00:29:39.438$ for the last 10 years is a little

 $00{:}29{:}39{.}438 \dashrightarrow 00{:}29{:}41{.}461$ piece of this fresh tissue was going

NOTE Confidence: 0.930604272727273

00:29:41.461 - 00:29:44.035 to my lab and in my lab we were

NOTE Confidence: 0.930604272727273

 $00:29:44.035 \rightarrow 00:29:45.980$ trying to establish primary cell line

NOTE Confidence: 0.930604272727273

 $00:29:45.980 \dashrightarrow 00:29:49.256$ as well as patient derives senokra.

NOTE Confidence: 0.930604272727273

 $00{:}29{:}49{.}260 \dashrightarrow 00{:}29{:}52{.}036$ So in few words the viable tumor tissue

NOTE Confidence: 0.930604272727273

 $00{:}29{:}52{.}036 \dashrightarrow 00{:}29{:}54{.}619$ was both placed in the Petri dish.

NOTE Confidence: 0.930604272727273

 $00{:}29{:}54.620 \dashrightarrow 00{:}29{:}57.516$ As well as in an animal with the

NOTE Confidence: 0.930604272727273

 $00{:}29{:}57{.}516$ --> $00{:}29{:}59{.}768$ ultimate goal again to establish

NOTE Confidence: 0.930604272727273

 $00{:}29{:}59{.}768 \dashrightarrow 00{:}30{:}02{.}654$ the disease in these avatar animals.

NOTE Confidence: 0.930604272727273

 $00:30:02.660 \rightarrow 00:30:04.850$ Why is that important is because

NOTE Confidence: 0.930604272727273

 $00:30:04.850 \longrightarrow 00:30:06.740$ as I'm showing you here,

NOTE Confidence: 0.930604272727273

 $00{:}30{:}06{.}740 \dashrightarrow 00{:}30{:}09{.}785$ the ultimate goal was to develop the

NOTE Confidence: 0.930604272727273

00:30:09.785 --> 00:30:11.795 patient to get the pathology report

NOTE Confidence: 0.930604272727273

00:30:11.795 --> 00:30:14.199 to start the patient on the standard

NOTE Confidence: 0.930604272727273

 $00{:}30{:}14.199 \dashrightarrow 00{:}30{:}16.099$ treatment that as we discussed

 $00:30:16.100 \rightarrow 00:30:18.280$ unfortunately sometime is unable

NOTE Confidence: 0.930604272727273

 $00{:}30{:}18.280 \dashrightarrow 00{:}30{:}22.515$ to cure the patient and having the

NOTE Confidence: 0.930604272727273

 $00:30:22.515 \rightarrow 00:30:25.255$ genetic sequencing data available.

NOTE Confidence: 0.930604272727273

 $00:30:25.260 \longrightarrow 00:30:28.230$ To be able to identify potential

NOTE Confidence: 0.930604272727273

00:30:28.230 --> 00:30:29.220 draggable marker,

NOTE Confidence: 0.930604272727273

 $00{:}30{:}29{.}220 \dashrightarrow 00{:}30{:}31{.}845$ test this potential drug in the avatar

NOTE Confidence: 0.930604272727273

 $00{:}30{:}31{.}845 \dashrightarrow 00{:}30{:}34{.}659$ animal on the tumor of the patient

NOTE Confidence: 0.930604272727273

 $00{:}30{:}34.659 \dashrightarrow 00{:}30{:}37.083$ when she was receiving the standard

NOTE Confidence: 0.930604272727273

 $00{:}30{:}37{.}155 \dashrightarrow 00{:}30{:}39{.}697$ of care and be ready for her in case

NOTE Confidence: 0.930604272727273

 $00{:}30{:}39{.}697 \dashrightarrow 00{:}30{:}42{.}260$ the disease come back or become resistant.

NOTE Confidence: 0.930604272727273

 $00{:}30{:}42.260 \dashrightarrow 00{:}30{:}45.276$ We already had tested in for that specific

NOTE Confidence: 0.930604272727273

 $00:30:45.276 \longrightarrow 00:30:48.313$ patient a certain number of drug to

NOTE Confidence: 0.930604272727273

 $00{:}30{:}48{.}313 \dashrightarrow 00{:}30{:}50{.}523$ provide her again with personalized

NOTE Confidence: 0.930604272727273

 $00:30:50.599 \rightarrow 00:30:53.209$ treatment at the time of occurrence.

NOTE Confidence: 0.930604272727273

 $00:30:53.210 \rightarrow 00:30:55.220$ Another thing that we've been able

NOTE Confidence: 0.930604272727273

 $00:30:55.220 \longrightarrow 00:30:58.412$ to to do in the last 5-6 years

- NOTE Confidence: 0.930604272727273
- $00{:}30{:}58{.}412 \dashrightarrow 00{:}31{:}00{.}632$ is also to provide personalized
- NOTE Confidence: 0.930604272727273
- $00:31:00.632 \longrightarrow 00:31:02.248$ diagnostic to our patient.
- NOTE Confidence: 0.930604272727273
- $00:31:02.250 \longrightarrow 00:31:03.722$ What does it mean?
- NOTE Confidence: 0.930604272727273
- $00:31:03.722 \longrightarrow 00:31:06.455$ It mean to detect the present of
- NOTE Confidence: 0.930604272727273
- $00{:}31{:}06{.}455 \dashrightarrow 00{:}31{:}08{.}650$ circulating tumor DNA that is
- NOTE Confidence: 0.930604272727273
- $00{:}31{:}08.650 \dashrightarrow 00{:}31{:}11.450$ mutated DNA coming from the tumor
- NOTE Confidence: 0.930604272727273
- $00:31:11.450 \rightarrow 00:31:14.450$ and this was again possible because
- NOTE Confidence: 0.930604272727273
- $00:31:14.450 \longrightarrow 00:31:15.950$ through the sequencing.
- NOTE Confidence: 0.930604272727273
- $00{:}31{:}15{.}950 \dashrightarrow 00{:}31{:}18{.}380$ We add knowledge about the potential
- NOTE Confidence: 0.930604272727273
- $00:31:18.380 \rightarrow 00:31:20.909$ driver mutation present in this patient,
- NOTE Confidence: 0.930604272727273
- $00:31:20.910 \longrightarrow 00:31:22.830$ present the tumor of this patient.
- NOTE Confidence: 0.930604272727273
- $00:31:22.830 \longrightarrow 00:31:26.310$ We design specific probe able to
- NOTE Confidence: 0.930604272727273
- $00:31:26.310 \longrightarrow 00:31:29.358$ amplify this mutation present in the
- NOTE Confidence: 0.930604272727273
- $00{:}31{:}29{.}358 \dashrightarrow 00{:}31{:}32{.}190$ circulation of this patient and using
- NOTE Confidence: 0.915300244615385
- $00{:}31{:}32{.}190 \dashrightarrow 00{:}31{:}35{.}406$ a not invasive method that is a simple
- NOTE Confidence: 0.915300244615385

 $00:31:35.406 \longrightarrow 00:31:39.290$ collection of of five to 10CC we were

NOTE Confidence: 0.915300244615385

 $00{:}31{:}39{.}290 \dashrightarrow 00{:}31{:}42{.}864$ able and this is a tumor inform as say to

NOTE Confidence: 0.915300244615385

 $00:31:42.864 \rightarrow 00:31:45.710$ detect the presence of the mutated DNA.

NOTE Confidence: 0.915300244615385

 $00:31:45.710 \longrightarrow 00:31:46.670$ That correlate with the

NOTE Confidence: 0.915300244615385

 $00:31:46.670 \longrightarrow 00:31:48.290$ presence of this in the patient.

NOTE Confidence: 0.915300244615385

00:31:48.290 --> 00:31:52.142 So this is a new approach of course to

NOTE Confidence: 0.915300244615385

 $00:31:52.142 \longrightarrow 00:31:54.610$ monitor and is called again personal

NOTE Confidence: 0.915300244615385

00:31:54.610 --> 00:31:56.650 like diagnostic type of approach

NOTE Confidence: 0.915300244615385

 $00{:}31{:}56.650 \dashrightarrow 00{:}31{:}59.191$ because allow us to see the presence

NOTE Confidence: 0.915300244615385

00:31:59.191 - > 00:32:01.770 of tumor DNA in the circulation.

NOTE Confidence: 0.915300244615385

 $00{:}32{:}01{.}770 \dashrightarrow 00{:}32{:}04{.}815$ But even more important being a tumor

NOTE Confidence: 0.915300244615385

 $00:32:04.815 \rightarrow 00:32:07.702$ inform approach this type of technology

NOTE Confidence: 0.915300244615385

 $00{:}32{:}07.702 \dashrightarrow 00{:}32{:}12.310$ become about 10 to 100 fold more sensitive.

NOTE Confidence: 0.915300244615385

 $00{:}32{:}12{.}310 \dashrightarrow 00{:}32{:}14.776$ Then the standard approach of looking

NOTE Confidence: 0.915300244615385

 $00:32:14.776 \rightarrow 00:32:18.260$ to again the Press of DNA when you don't

NOTE Confidence: 0.915300244615385

 $00:32:18.260 \rightarrow 00:32:20.909$ know what type of mutation is pressed.

 $00:32:20.910 \longrightarrow 00:32:22.206$ So using this approach,

NOTE Confidence: 0.915300244615385

 $00:32:22.206 \rightarrow 00:32:25.174$ let me show you a little bit the progress

NOTE Confidence: 0.915300244615385

 $00{:}32{:}25{.}174 \dashrightarrow 00{:}32{:}27{.}860$ that we have done for the last again

NOTE Confidence: 0.915300244615385

 $00:32:27.860 \rightarrow 00:32:30.212$ few years taking care in particular

NOTE Confidence: 0.915300244615385

 $00{:}32{:}30{.}212 \dashrightarrow 00{:}32{:}35{.}432$ the difficult to treat patient and one

NOTE Confidence: 0.915300244615385

 $00:32:35.432 \dashrightarrow 00:32:39.524$ of the most difficult uterine cancer.

NOTE Confidence: 0.915300244615385

 $00:32:39.530 \longrightarrow 00:32:41.882$ They were dealing with in the

NOTE Confidence: 0.915300244615385

 $00:32:41.882 \longrightarrow 00:32:43.450$ clinic are uterine seros.

NOTE Confidence: 0.915300244615385

 $00:32:43.450 \longrightarrow 00:32:47.210$ They are rare tumor 3:00 to 9:50 to

NOTE Confidence: 0.915300244615385

00:32:47.210 --> 00:32:50.528 10% over all of all uterine cancer,

NOTE Confidence: 0.915300244615385

 $00:32:50.530 \longrightarrow 00:32:53.469$ but they kill over 40% of our patient.

NOTE Confidence: 0.915300244615385

 $00:32:53.469 \longrightarrow 00:32:55.347$ Why is that is because they

NOTE Confidence: 0.915300244615385

 $00{:}32{:}55{.}347 \dashrightarrow 00{:}32{:}57{.}330$ are biologically aggressive,

NOTE Confidence: 0.915300244615385

 $00:32:57.330 \rightarrow 00:33:00.180$ they can start spreading very early

NOTE Confidence: 0.915300244615385

 $00:33:00.180 \dashrightarrow 00:33:02.639$ and they're usually difficult to

00:33:02.639 - > 00:33:04.567 treat in thermal chemotherapy.

NOTE Confidence: 0.915300244615385

 $00:33:04.570 \rightarrow 00:33:07.377$ One of the important thing is that.

NOTE Confidence: 0.915300244615385

 $00:33:07.380 \longrightarrow 00:33:10.208$ The Gynong division at Yale has been

NOTE Confidence: 0.915300244615385

 $00:33:10.208 \rightarrow 00:33:13.940$ for a long time a really a place

NOTE Confidence: 0.915300244615385

 $00:33:13.940 \dashrightarrow 00:33:16.540$ where this patient were affair.

NOTE Confidence: 0.915300244615385

 $00{:}33{:}16{.}540 \dashrightarrow 00{:}33{:}18{.}628$ And this is because of course and this

NOTE Confidence: 0.915300244615385

 $00:33:18.628 \rightarrow 00:33:20.857$ is because of the work of Doctor Schwartz,

NOTE Confidence: 0.915300244615385

 $00:33:20.860 \longrightarrow 00:33:23.220$ that work here for the last 50 years.

NOTE Confidence: 0.915300244615385

 $00{:}33{:}23{.}220 \dashrightarrow 00{:}33{:}25{.}385$ Doctor Schwartz has a specific

NOTE Confidence: 0.915300244615385

 $00:33:25.385 \longrightarrow 00:33:27.550$ interest in studying this tumor.

NOTE Confidence: 0.915300244615385

 $00:33:27.550 \longrightarrow 00:33:29.517$ And when he asked me to join

NOTE Confidence: 0.915300244615385

00:33:29.517 --> 00:33:30.710 him 16 years ago,

NOTE Confidence: 0.915300244615385

 $00:33:30.710 \longrightarrow 00:33:32.908$ I kept on going and and work

NOTE Confidence: 0.915300244615385

 $00:33:32.908 \longrightarrow 00:33:34.443$ in specifically against this

NOTE Confidence: 0.915300244615385

 $00:33:34.443 \rightarrow 00:33:36.030$ biologically aggressive tumor.

NOTE Confidence: 0.915300244615385

 $00:33:36.030 \longrightarrow 00:33:38.800$ So this is to say we see a lot of this

 $00{:}33{:}38{.}800 \dashrightarrow 00{:}33{:}41{.}262$ rare tumor and because of that we were

NOTE Confidence: 0.915300244615385

 $00{:}33{:}41{.}262 \dashrightarrow 00{:}33{:}44{.}052$ able to collect a lot of specimen and

NOTE Confidence: 0.915300244615385

 $00:33:44.052 \rightarrow 00:33:48.126$ we perform and we reported the first

NOTE Confidence: 0.915300244615385

00:33:48.126 --> 00:33:50.643 comprehensive genetic landscape analysis

NOTE Confidence: 0.915300244615385

 $00:33:50.643 \rightarrow 00:33:54.638$ of the rare manigancy and this was published.

NOTE Confidence: 0.915300244615385

00:33:54.638 --> 00:33:55.010 Again,

NOTE Confidence: 0.915300244615385

 $00:33:55.010 \rightarrow 00:33:56.822$ over 10 years ago,

NOTE Confidence: 0.915300244615385

 $00:33:56.822 \longrightarrow 00:33:59.087$ as you can see here,

NOTE Confidence: 0.915300244615385

 $00:33:59.090 \dashrightarrow 00:34:01.652$ what we found in this study there

NOTE Confidence: 0.915300244615385

 $00:34:01.652 \rightarrow 00:34:04.530$ was by the way before the TC

NOTE Confidence: 0.915300244615385

 $00:34:04.530 \longrightarrow 00:34:06.796$ published before the TCGA data,

NOTE Confidence: 0.915300244615385

00:34:06.796 --> 00:34:07.209 okay,

NOTE Confidence: 0.915300244615385

 $00{:}34{:}07{.}209 \dashrightarrow 00{:}34{:}09{.}687$ about six months before the first

NOTE Confidence: 0.915300244615385

 $00:34:09.690 \rightarrow 00:34:11.882$ major comprehensive analysis done

NOTE Confidence: 0.915300244615385

00:34:11.882 --> 00:34:15.170 by the Tumor Cancer Atlas Network,

 $00:34:15.170 \rightarrow 00:34:17.290$ important chronologically to say that

NOTE Confidence: 0.915300244615385

 $00{:}34{:}17.290 \dashrightarrow 00{:}34{:}20.152$ this is what we found, we sound,

NOTE Confidence: 0.915300244615385

 $00:34:20.152 \rightarrow 00:34:22.007$ we found something very interesting,

NOTE Confidence: 0.915300244615385

 $00{:}34{:}22.010 \dashrightarrow 00{:}34{:}22.548$ namely.

NOTE Confidence: 0.915300244615385

 $00{:}34{:}22{.}548 \dashrightarrow 00{:}34{:}25{.}776$ A small minority of this biological

NOTE Confidence: 0.915300244615385

00:34:25.776 --> 00:34:28.173 aggressive cancer called uterine series NOTE Confidence: 0.915300244615385

 $00{:}34{:}28{.}173 \dashrightarrow 00{:}34{:}30{.}300$ 8 to 9% had this characteristic year.

NOTE Confidence: 0.915300244615385

 $00:34:30.300 \longrightarrow 00:34:33.258$ They had a huge number of mutation.

NOTE Confidence: 0.915300244615385

00:34:33.260 --> 00:34:35.164 They were ultra mutated,

NOTE Confidence: 0.915300244615385

 $00:34:35.164 \rightarrow 00:34:38.700$ but they were Microsoft and lie stable.

NOTE Confidence: 0.915300244615385

 $00{:}34{:}38{.}700 \dashrightarrow 00{:}34{:}41{.}406$ They didn't have any mismatch repair

NOTE Confidence: 0.915300244615385

00:34:41.406 - 00:34:43.540 deficiency. So what were this tumor?

NOTE Confidence: 0.915300244615385

 $00:34:43.540 \dashrightarrow 00:34:44.938$ We didn't know at that time,

NOTE Confidence: 0.915300244615385

 $00{:}34{:}44{.}940 \dashrightarrow 00{:}34{:}47{.}418$ but they had this genetic characteristic,

NOTE Confidence: 0.915300244615385

 $00:34:47.420 \longrightarrow 00:34:49.085$ the remaining group.

NOTE Confidence: 0.915300244615385

 $00:34:49.085 \rightarrow 00:34:52.415$ There was again the striking majority

- NOTE Confidence: 0.915300244615385
- $00:34:52.420 \rightarrow 00:34:55.140 9095\%$ where normal mutated tumor,
- NOTE Confidence: 0.915300244615385
- $00:34:55.140 \longrightarrow 00:34:57.460$ very low number of mutation,
- NOTE Confidence: 0.915300244615385
- 00:34:57.460 --> 00:35:00.770 MSI stable and cold meaning
- NOTE Confidence: 0.915300244615385
- 00:35:00.770 -> 00:35:02.094 minimal inflammation,
- NOTE Confidence: 0.915300244615385
- $00:35:02.100 \rightarrow 00:35:04.460$ minimum inflammatory cell present
- NOTE Confidence: 0.915300244615385
- $00:35:04.460 \dashrightarrow 00:35:06.820$ in the tumor microenvironment.
- NOTE Confidence: 0.8958366125
- $00:35:06.820 \longrightarrow 00:35:10.154$ So of course we start looking to the 92%,
- NOTE Confidence: 0.8958366125
- 00:35:10.154 --> 00:35:11.824 right. And what I'm showing
- NOTE Confidence: 0.8958366125
- $00:35:11.824 \longrightarrow 00:35:13.780$ you here was our finding,
- NOTE Confidence: 0.8958366125
- $00{:}35{:}13.780 \dashrightarrow 00{:}35{:}16.816$ we look to this 21,000 gene
- NOTE Confidence: 0.8958366125
- $00:35:16.820 \rightarrow 00:35:19.410$ and we found specific pathway.
- NOTE Confidence: 0.8958366125
- 00:35:19.410 --> 00:35:21.960 That this aggressive tumor was using
- NOTE Confidence: 0.8958366125
- 00:35:21.960 --> 00:35:24.854 to survive hemotherapy and then come
- NOTE Confidence: 0.8958366125
- $00:35:24.854 \rightarrow 00:35:28.519$ back after treatment and one of these
- NOTE Confidence: 0.8958366125
- 00:35:28.519 --> 00:35:31.442 again pathway that I'm you know I'm NOTE Confidence: 0.8958366125

 $00:35:31.442 \dashrightarrow 00:35:33.570$ highlighting here was the hair to new

NOTE Confidence: 0.930428735769231

00:35:35.610 - 00:35:39.174 PAKAKT enter pathway over 1/3 of

NOTE Confidence: 0.930428735769231

00:35:39.174 --> 00:35:42.406 this patient had amplification on on NOTE Confidence: 0.930428735769231

 $00{:}35{:}42{.}406 \dashrightarrow 00{:}35{:}45{.}422$ the C RB2 gene that encode for the

NOTE Confidence: 0.930428735769231

 $00:35:45.523 \rightarrow 00:35:49.105$ epidermal glow factor type 2 receptor.

NOTE Confidence: 0.930428735769231

 $00{:}35{:}49{.}110 \dashrightarrow 00{:}35{:}51{.}348$ This is well known in oncology.

NOTE Confidence: 0.930428735769231

 $00{:}35{:}51{.}350 \dashrightarrow 00{:}35{:}53{.}750$ Why? Because there are specific

NOTE Confidence: 0.930428735769231

 $00:35:53.750 \rightarrow 00:35:56.150$ treatment they target this pathway.

NOTE Confidence: 0.930428735769231

 $00:35:56.150 \longrightarrow 00:35:59.123$ The target have to new trust is

NOTE Confidence: 0.930428735769231

 $00{:}35{:}59{.}123 \dashrightarrow 00{:}36{:}01{.}588$ monoclonal antibodies one of those.

NOTE Confidence: 0.930428735769231

 $00{:}36{:}01.590 \dashrightarrow 00{:}36{:}04.902$ But the problem is that this was

NOTE Confidence: 0.930428735769231

 $00:36:04.902 \longrightarrow 00:36:06.882$ an antibody available for the

NOTE Confidence: 0.930428735769231

 $00:36:06.882 \rightarrow 00:36:09.013$ treatment of breast cancer patient

NOTE Confidence: 0.930428735769231

 $00:36:09.013 \rightarrow 00:36:11.623$ and later on gastric cancer patient.

NOTE Confidence: 0.930428735769231

 $00:36:11.630 \longrightarrow 00:36:13.826$ But there was really no report,

NOTE Confidence: 0.930428735769231

 $00{:}36{:}13.830 \dashrightarrow 00{:}36{:}16.050$ no literature in the treatment

- NOTE Confidence: 0.930428735769231
- $00{:}36{:}16.050 \dashrightarrow 00{:}36{:}17.826$ of uterine serous carcinoma.

 $00:36:17.830 \longrightarrow 00:36:20.166$ So we decided to fill that gap and

NOTE Confidence: 0.930428735769231

 $00:36:20.166 \rightarrow 00:36:22.146$ we designed a study here at Yale,

NOTE Confidence: 0.930428735769231

 $00:36:22.150 \rightarrow 00:36:25.280$ we designed an investigator initiated

NOTE Confidence: 0.930428735769231

 $00{:}36{:}25{.}280 \dashrightarrow 00{:}36{:}28{.}385$ trial there was again through a

NOTE Confidence: 0.930428735769231

 $00{:}36{:}28{.}385 \dashrightarrow 00{:}36{:}30{.}518$ consortium of institution because as

NOTE Confidence: 0.930428735769231

 $00:36:30.518 \rightarrow 00:36:33.670$ I just told you this is a rare cancer

NOTE Confidence: 0.930428735769231

 $00:36:33.670 \rightarrow 00:36:35.945$ and we were and we were targeting

NOTE Confidence: 0.930428735769231

 $00:36:35.945 \longrightarrow 00:36:37.992$ about 30% of this rare cancer the

NOTE Confidence: 0.930428735769231

 $00:36:37.992 \longrightarrow 00:36:39.590$ one over expressing her to new.

NOTE Confidence: 0.930428735769231

 $00:36:39.590 \longrightarrow 00:36:42.350$ So we needed multiple side to do that.

NOTE Confidence: 0.930428735769231

 $00{:}36{:}42{.}350 \dashrightarrow 00{:}36{:}44{.}793$ So we had eleven side going from

NOTE Confidence: 0.930428735769231

00:36:44.793 --> 00:36:47.258 John Hopkins to higher state and.

NOTE Confidence: 0.930428735769231

 $00{:}36{:}47.258 \dashrightarrow 00{:}36{:}49.130$ The primary goal here,

NOTE Confidence: 0.930428735769231

 $00{:}36{:}49{.}130 \dashrightarrow 00{:}36{:}51{.}248$ the primary objective was to understand

 $00:36:51.248 \longrightarrow 00:36:53.748$ if adding trust to the map this

NOTE Confidence: 0.930428735769231

 $00:36:53.748 \rightarrow 00:36:55.453$ monoclonal antibody to our standard

NOTE Confidence: 0.930428735769231

00:36:55.453 --> 00:36:57.921 of care that is based on carbo packet NOTE Confidence: 0.930428735769231

 $00:36:57.921 \rightarrow 00:37:00.550$ tax could make the difference in term

NOTE Confidence: 0.930428735769231

 $00:37:00.550 \rightarrow 00:37:02.730$ of increasing progression free survival,

NOTE Confidence: 0.930428735769231

 $00:37:02.730 \rightarrow 00:37:04.314$ increasing the objective response

NOTE Confidence: 0.930428735769231

 $00{:}37{:}04{.}314 \dashrightarrow 00{:}37{:}06{.}294$ rate and the overall survival

NOTE Confidence: 0.930428735769231

 $00:37:06.294 \rightarrow 00:37:08.130$ of this patient and so on.

NOTE Confidence: 0.930428735769231

 $00{:}37{:}08{.}130 \dashrightarrow 00{:}37{:}10{.}330$ And these are our result.

NOTE Confidence: 0.930428735769231

 $00:37:10.330 \longrightarrow 00:37:12.766$ So we were able to show that

NOTE Confidence: 0.930428735769231

 $00:37:12.766 \rightarrow 00:37:14.500$ adding this monoclonal antibody.

NOTE Confidence: 0.930428735769231

 $00:37:14.500 \rightarrow 00:37:16.148$ Without any significant increase

NOTE Confidence: 0.930428735769231

 $00:37:16.148 \longrightarrow 00:37:18.208$ in toxicity when compared to

NOTE Confidence: 0.930428735769231

00:37:18.208 --> 00:37:20.338 the Carbo Parkly tax regiment,

NOTE Confidence: 0.930428735769231

 $00:37:20.340 \longrightarrow 00:37:23.028$ we were able to increase of about 5

NOTE Confidence: 0.930428735769231

 $00:37:23.028 \rightarrow 00:37:24.849$ month progression free survival and

- NOTE Confidence: 0.930428735769231
- $00:37:24.849 \longrightarrow 00:37:27.376$ close to six month the overall survival
- NOTE Confidence: 0.930428735769231
- $00:37:27.446 \dashrightarrow 00:37:29.536$ of the entire patient population.
- NOTE Confidence: 0.930428735769231
- $00{:}37{:}29{.}540 \dashrightarrow 00{:}37{:}32{.}228$ They were both advanced stage as well
- NOTE Confidence: 0.930428735769231
- $00{:}37{:}32.228 \dashrightarrow 00{:}37{:}34.417$ as recurrent patient with uterine
- NOTE Confidence: 0.930428735769231
- $00{:}37{:}34{.}417 \dashrightarrow 00{:}37{:}36{.}586$ serous carcinoma with a benefit
- NOTE Confidence: 0.930428735769231
- 00:37:36.586 --> 00:37:39.400 even much higher in the chemo naive
- NOTE Confidence: 0.930428735769231
- $00:37:39.486 \longrightarrow 00:37:42.220$ stage 3 and stage 4 disease so.
- NOTE Confidence: 0.930428735769231
- $00:37:42.220 \rightarrow 00:37:45.020$ This study is important because
- NOTE Confidence: 0.930428735769231
- 00:37:45.020 --> 00:37:48.310 not only ask recognized important,
- NOTE Confidence: 0.930428735769231
- 00:37:48.310 --> 00:37:49.206 you know,
- NOTE Confidence: 0.930428735769231
- 00:37:49.206 --> 00:37:52.746 listing it as a major advancement in 2019,
- NOTE Confidence: 0.930428735769231
- $00{:}37{:}52.746 \dashrightarrow 00{:}37{:}55.518$ one of the five major advancement of
- NOTE Confidence: 0.930428735769231
- $00:37:55.518 \rightarrow 00:37:58.914$ that year in the oncology literature,
- NOTE Confidence: 0.930428735769231
- $00{:}37{:}58{.}914 \dashrightarrow 00{:}38{:}02{.}862$ but also because this increase in
- NOTE Confidence: 0.930428735769231
- $00:38:02.862 \rightarrow 00:38:05.642$ overall survival caused the revision
- NOTE Confidence: 0.930428735769231

 $00:38:05.642 \rightarrow 00:38:07.578$ of the National Comprehensive

NOTE Confidence: 0.930428735769231

 $00{:}38{:}07{.}578 \dashrightarrow 00{:}38{:}10{.}052$ Cancer Network guideline that are

NOTE Confidence: 0.930428735769231

 $00:38:10.052 \longrightarrow 00:38:12.096$ widely used as standard.

NOTE Confidence: 0.930428735769231

00:38:12.100 -> 00:38:14.914 Of care again in oncology by both

NOTE Confidence: 0.930428735769231

 $00{:}38{:}14{.}914 \dashrightarrow 00{:}38{:}17{.}780$ clinician as well as insurance company

NOTE Confidence: 0.930428735769231

 $00:38:17.780 \longrightarrow 00:38:21.756$ and since that time 2019, Carboplatin,

NOTE Confidence: 0.930428735769231

00:38:21.756 --> 00:38:24.380 Parkly, Taxol and Trastozoma.

NOTE Confidence: 0.930428735769231

 $00:38:24.380 \longrightarrow 00:38:27.110$ So the yield regimen is now again

NOTE Confidence: 0.930428735769231

 $00{:}38{:}27{.}110 \dashrightarrow 00{:}38{:}29{.}337$ recommended in old patient with

NOTE Confidence: 0.930428735769231

00:38:29.337 --> 00:38:31.792 utransitous carcinoma or expressing her

NOTE Confidence: 0.930428735769231

 $00{:}38{:}31.792 \dashrightarrow 00{:}38{:}36.700$ to you in the US as well as worldwide,

NOTE Confidence: 0.930428735769231

 $00:38:36.700 \longrightarrow 00:38:39.288$ let me do now.

NOTE Confidence: 0.930428735769231

 $00:38:39.290 \longrightarrow 00:38:41.058$ Let me tell you a little bit more

NOTE Confidence: 0.930428735769231

 $00:38:41.058 \longrightarrow 00:38:42.489$ about the Poly ultramutated.

NOTE Confidence: 0.930428735769231

 $00{:}38{:}42.490 \dashrightarrow 00{:}38{:}47.044$ So we look to this uterines of carcinoma.

NOTE Confidence: 0.930428735769231

00:38:47.050 - 00:38:48.130 As I told you before,

 $00:38:48.130 \longrightarrow 00:38:49.621$ a small minority,

NOTE Confidence: 0.930428735769231

 $00:38:49.621 \rightarrow 00:38:53.100$ a nine percent had this abnormal alteration

NOTE Confidence: 0.930428735769231

 $00:38:53.185 \rightarrow 00:38:56.008$ in a gene called DNA polymerase epsilon.

NOTE Confidence: 0.930428735769231

 $00:38:56.008 \rightarrow 00:38:58.970$ This is a DNA polymerase at that time.

NOTE Confidence: 0.930428735769231

00:38:58.970 - 00:39:01.328 I remind you this was over 10 years ago.

NOTE Confidence: 0.930428735769231

 $00:39:01.330 \rightarrow 00:39:04.210$ Nobody knew what this really gene was doing,

NOTE Confidence: 0.930428735769231

 $00:39:04.210 \longrightarrow 00:39:04.714$ but.

NOTE Confidence: 0.930428735769231

00:39:04.714 --> 00:39:06.730 Look into this tumor,

NOTE Confidence: 0.930428735769231

 $00:39{:}06.730 \dashrightarrow 00{:}39{:}08.785$ we immediately realize that was

NOTE Confidence: 0.930428735769231

00:39:08.785 --> 00:39:10.840 really playing a major role

NOTE Confidence: 0.939061272941177

 $00:39:10.918 \rightarrow 00:39:13.844$ in the proof reading capability of the DNA.

NOTE Confidence: 0.939061272941177

 $00{:}39{:}13.850 \dashrightarrow 00{:}39{:}15.978$ Why is that? Is because this tumor

NOTE Confidence: 0.939061272941177

 $00{:}39{:}15{.}978 \dashrightarrow 00{:}39{:}18{.}248$ had a huge number of mutation.

NOTE Confidence: 0.939061272941177

00:39:18.250 --> 00:39:21.064 So polymerase epsilon in normal cell again

NOTE Confidence: 0.939061272941177

 $00{:}39{:}21.064 \dashrightarrow 00{:}39{:}24.010$ is endowed with proof reading capability.

 $00:39:24.010 \longrightarrow 00:39:26.458$ When the cell divide and the

NOTE Confidence: 0.939061272941177

 $00:39:26.458 \rightarrow 00:39:28.666$ DNA duplicate the DNA polymerase

NOTE Confidence: 0.939061272941177

 $00{:}39{:}28.666 \dashrightarrow 00{:}39{:}31.116$ epsilon proof read the sequence and

NOTE Confidence: 0.939061272941177

 $00:39:31.116 \longrightarrow 00:39:34.684$ if there is an error, a mutation.

NOTE Confidence: 0.939061272941177

 $00{:}39{:}34{.}684 \dashrightarrow 00{:}39{:}38{.}303$ Stop and repair and fix the error.

NOTE Confidence: 0.939061272941177

 $00:39:38.310 \longrightarrow 00:39:41.091$ But if one of the two allele present in NOTE Confidence: 0.939061272941177

00:39:41.091 --> 00:39:43.869 the cell encoding for the polymerase

NOTE Confidence: 0.939061272941177

00:39:43.869 - > 00:39:46.000 epsilon mutate and the majority of

NOTE Confidence: 0.939061272941177

00:39:46.000 $-\!>$ 00:39:48.140 the special are also multi mutation

NOTE Confidence: 0.939061272941177

 $00:39:48.140 \longrightarrow 00:39:50.265$ so acquired during the lifetime,

NOTE Confidence: 0.939061272941177

 $00{:}39{:}50{.}270 \dashrightarrow 00{:}39{:}52{.}928$ the cell survive but start acquiring

NOTE Confidence: 0.939061272941177

00:39:52.928 --> 00:39:55.209 error mistakes and start accumulating

NOTE Confidence: 0.939061272941177

 $00{:}39{:}55{.}209 \dashrightarrow 00{:}39{:}58{.}254$ again mutation to the point where you

NOTE Confidence: 0.939061272941177

 $00:39:58.254 \dashrightarrow 00:40:00.998$ have this monster tumor they have.

NOTE Confidence: 0.939061272941177

 $00{:}40{:}00{.}998 \dashrightarrow 00{:}40{:}03{.}966$ 1000 and 1000 mutation much higher than

NOTE Confidence: 0.939061272941177

 $00:40:03.970 \rightarrow 00:40:07.490$ any other human cancer including lung,

00:40:07.490 --> 00:40:08.130 Melanoma,

NOTE Confidence: 0.939061272941177

 $00:40:08.130 \rightarrow 00:40:10.050$ bladder cancer that are well known to be,

NOTE Confidence: 0.939061272941177

 $00{:}40{:}10.050 \dashrightarrow 00{:}40{:}12.570$ I mean have a high number of mutation.

NOTE Confidence: 0.939061272941177

 $00:40:12.570 \rightarrow 00:40:14.989$ So when we look to this tumor specifically

NOTE Confidence: 0.939061272941177

 $00{:}40{:}14.989 \dashrightarrow 00{:}40{:}17.263$ we saw that in the exonuclease

NOTE Confidence: 0.939061272941177

 $00{:}40{:}17.263 \dashrightarrow 00{:}40{:}19.729$ domain of the polymerase epsilon,

NOTE Confidence: 0.939061272941177

 $00:40:19.730 \longrightarrow 00:40:21.905$ the majority of recurrent mutation

NOTE Confidence: 0.939061272941177

 $00:40:21.905 \longrightarrow 00:40:24.018$ with localizing that area and

NOTE Confidence: 0.939061272941177

 $00:40:24.018 \longrightarrow 00:40:25.490$ why is that important?

NOTE Confidence: 0.939061272941177

 $00:40:25.490 \longrightarrow 00:40:28.472$ Is because even if the majority

NOTE Confidence: 0.939061272941177

00:40:28.472 --> 00:40:30.381 of the polyultramutated USC,

NOTE Confidence: 0.939061272941177

 $00{:}40{:}30{.}381 \dashrightarrow 00{:}40{:}33{.}258$ they do well when compared to the

NOTE Confidence: 0.939061272941177

 $00{:}40{:}33.258 \dashrightarrow 00{:}40{:}35.644$ other that I told you still some

NOTE Confidence: 0.939061272941177

 $00{:}40{:}35{.}644 \dashrightarrow 00{:}40{:}37{.}546$ of this patient where they come

NOTE Confidence: 0.939061272941177

 $00{:}40{:}37{.}546 \dashrightarrow 00{:}40{:}39{.}746$ back with recurring disease and

 $00:40:39.746 \rightarrow 00:40:41.570$ these tumor are relentless,

NOTE Confidence: 0.939061272941177

00:40:41.570 - 00:40:43.690 they are resistant to chemotherapy,

NOTE Confidence: 0.939061272941177

 $00:40:43.690 \longrightarrow 00:40:45.680$ they are resistant to radiation

NOTE Confidence: 0.939061272941177

 $00:40:45.680 \rightarrow 00:40:48.551$ treatment and they kill our patient so.

NOTE Confidence: 0.939061272941177

 $00{:}40{:}48.551 \dashrightarrow 00{:}40{:}50.938$ Few years ago looking to the set

NOTE Confidence: 0.939061272941177

 $00{:}40{:}50{.}938 \dashrightarrow 00{:}40{:}53{.}530$ to the genetic characteristic,

NOTE Confidence: 0.939061272941177

 $00:40:53.530 \rightarrow 00:40:55.728$ to the fact they were all permutated,

NOTE Confidence: 0.939061272941177

 $00:40:55.730 \rightarrow 00:40:58.538$ we thought they could be exquisitively

NOTE Confidence: 0.939061272941177

 $00{:}40{:}58.538 \dashrightarrow 00{:}41{:}01.130$ sensitive to new checkpoint inhibitor

NOTE Confidence: 0.939061272941177

00:41:01.130 --> 00:41:03.968 and and again few years ago and

NOTE Confidence: 0.939061272941177

 $00{:}41{:}03.968 \dashrightarrow 00{:}41{:}05.489$ here I'm showing you a case report.

NOTE Confidence: 0.939061272941177

 $00:41:05.490 \longrightarrow 00:41:07.882$ This is one of my patient treated in

NOTE Confidence: 0.939061272941177

 $00:41:07.882 \rightarrow 00:41:10.699$ two or seven initially for an advanced

NOTE Confidence: 0.939061272941177

 $00:41:10.699 \rightarrow 00:41:15.331$ stage type 2 mix again uterine carcinoma.

NOTE Confidence: 0.939061272941177

 $00{:}41{:}15{.}331 \dashrightarrow 00{:}41{:}18{.}466$ She was treated with surgery.

NOTE Confidence: 0.939061272941177

00:41:18.470 --> 00:41:19.702 Chemotherapy or addition treatment,

- NOTE Confidence: 0.939061272941177
- $00{:}41{:}19.702 \dashrightarrow 00{:}41{:}21.870$ she did well for a few year,
- NOTE Confidence: 0.939061272941177
- 00:41:21.870 --> 00:41:24.467 came back in 2012 with recurring disease,
- NOTE Confidence: 0.939061272941177
- $00:41:24.470 \longrightarrow 00:41:28.650$ multiple metastatic lesion as
- NOTE Confidence: 0.939061272941177
- $00:41:28.650 \longrightarrow 00:41:32.810$ usual and at that time point
- NOTE Confidence: 0.939061272941177
- $00:41:32.810 \longrightarrow 00:41:34.185$ she was retrieved with surgery,
- NOTE Confidence: 0.939061272941177
- $00:41:34.190 \longrightarrow 00:41:37.070$ additional bowel resection,
- NOTE Confidence: 0.939061272941177
- $00:41:37.070 \rightarrow 00:41:40.426$ additional chemo and she start progressing.
- NOTE Confidence: 0.939061272941177
- $00{:}41{:}40{.}426 \dashrightarrow 00{:}41{:}43{.}982$ So she was referred to us by an outside
- NOTE Confidence: 0.939061272941177
- $00{:}41{:}43{.}982 \dashrightarrow 00{:}41{:}46{.}622$ medical on cologist for a second opinion.
- NOTE Confidence: 0.939061272941177
- 00:41:46.622 --> 00:41:47.498 In reality,
- NOTE Confidence: 0.939061272941177
- $00{:}41{:}47{.}500 \dashrightarrow 00{:}41{:}49{.}509$ of course the referral was to for
- NOTE Confidence: 0.939061272941177
- $00:41:49.509 \rightarrow 00:41:51.529$ the patient to accept Hospice because
- NOTE Confidence: 0.939061272941177
- $00:41:51.529 \longrightarrow 00:41:53.354$ there was really nothing that
- NOTE Confidence: 0.939061272941177
- $00:41:53.354 \rightarrow 00:41:55.427$ we could offer internal systemic
- NOTE Confidence: 0.939061272941177
- $00:41:55.427 \longrightarrow 00:41:57.135$ treatment to this patient.
- NOTE Confidence: 0.939061272941177

 $00:41:57.140 \longrightarrow 00:41:58.580$ When she came to see us,

NOTE Confidence: 0.939061272941177

 $00:41:58.580 \longrightarrow 00:42:00.854$ we sequenced the tumor with foundation

NOTE Confidence: 0.939061272941177

 $00{:}42{:}00{.}854 \dashrightarrow 00{:}42{:}03{.}292$ medicine and what we found was that

NOTE Confidence: 0.939061272941177

 $00:42:03.292 \longrightarrow 00:42:05.760$ she had a Poly hotspot mutation

NOTE Confidence: 0.939061272941177

 $00:42:05.760 \rightarrow 00:42:07.660$ in the exonuclearized domain,

NOTE Confidence: 0.939061272941177

 $00{:}42{:}07.660 \dashrightarrow 00{:}42{:}12.752$ the P 2028 to 286 are this huge

NOTE Confidence: 0.939061272941177

 $00:42:12.752 \longrightarrow 00:42:17.090$ number of mutation present so.

NOTE Confidence: 0.939061272941177

00:42:17.090 --> 00:42:19.358 This is was a lucky patient because

NOTE Confidence: 0.939061272941177

 $00{:}42{:}19{.}358 \dashrightarrow 00{:}42{:}23{.}994$ at that time was about 2014 we had the

NOTE Confidence: 0.939061272941177

 $00:42:23.994 \rightarrow 00:42:26.570$ program on going at Yale with nivolumab.

NOTE Confidence: 0.939061272941177

 $00{:}42{:}26.570 \dashrightarrow 00{:}42{:}28.684$ This was a program that doctor Roy

NOTE Confidence: 0.939061272941177

 $00{:}42{:}28.684 \dashrightarrow 00{:}42{:}30.944$ Herbs as active in lung cancer at

NOTE Confidence: 0.939061272941177

 $00:42:30.944 \rightarrow 00:42:32.549$ that time point Nivolumab there

NOTE Confidence: 0.939061272941177

 $00{:}42{:}32{.}549 \dashrightarrow 00{:}42{:}34{.}927$ was the first immune checkpoint

NOTE Confidence: 0.939061272941177

 $00:42:34.927 \rightarrow 00:42:36.923$ inhibitor before before pembrolizumab.

NOTE Confidence: 0.939061272941177

 $00:42:36.930 \rightarrow 00:42:39.422$ OK true that everybody now is using

- NOTE Confidence: 0.939061272941177
- $00{:}42{:}39{.}422 \dashrightarrow 00{:}42{:}41{.}425$ was only approving Melanoma but
- NOTE Confidence: 0.939061272941177
- 00:42:41.425 --> 00:42:43.770 because of this program I was able
- NOTE Confidence: 0.9201268
- $00{:}42{:}43.770 \dashrightarrow 00{:}42{:}47.360$ to. To the OR through a compassionate
- NOTE Confidence: 0.9201268
- $00{:}42{:}47{.}360 \dashrightarrow 00{:}42{:}49{.}567$ base approval to offer to this
- NOTE Confidence: 0.9201268
- $00{:}42{:}49{.}567 \dashrightarrow 00{:}42{:}51{.}217$ patient NI voluma for free.
- NOTE Confidence: 0.9201268
- $00:42:51.220 \longrightarrow 00:42:52.472$ She signed a consent.
- NOTE Confidence: 0.9201268
- $00:42:52.472 \longrightarrow 00:42:54.776$ I got the approval for the chief
- NOTE Confidence: 0.9201268
- $00:42:54.776 \longrightarrow 00:42:56.541$ medical officer of the Yale
- NOTE Confidence: 0.9201268
- $00:42:56.541 \longrightarrow 00:42:58.846$ Cancer Center was or Jerry or
- NOTE Confidence: 0.9201268
- 00:42:58.846 --> 00:43:01.340 Lillenbaum and the rest of history.
- NOTE Confidence: 0.9201268
- $00{:}43{:}01{.}340 \dashrightarrow 00{:}43{:}03{.}300$ We start treating this patient.
- NOTE Confidence: 0.9201268
- 00:43:03.300 --> 00:43:06.108 I don't know if you can see this is
- NOTE Confidence: 0.9201268
- $00:43:06.108 \rightarrow 00:43:08.756$ one of the metastatic deposit as big
- NOTE Confidence: 0.9201268
- 00:43:08.756 --> 00:43:12.047 as a spleen as you can see here 7-8
- NOTE Confidence: 0.9201268
- $00:43:12.047 \rightarrow 00:43:14.609$ centimeter masses in the Donal cavity.
- NOTE Confidence: 0.9201268

00:43:14.610 --> 00:43:18.130 This is her bladder nodule or two 3

NOTE Confidence: 0.9201268

 $00{:}43{:}18.130 \dashrightarrow 00{:}43{:}21.450$ centimeter growing in the wall of the

NOTE Confidence: 0.9201268

 $00{:}43{:}21{.}450 \dashrightarrow 00{:}43{:}23{.}370$ bladder causing constant bleeding.

NOTE Confidence: 0.9201268

 $00{:}43{:}23{.}370 \dashrightarrow 00{:}43{:}26{.}646$ In a week the bleeding disappear

NOTE Confidence: 0.9201268

 $00{:}43{:}26.650 \dashrightarrow 00{:}43{:}31.774$ and in about 6 weeks pain disappear.

NOTE Confidence: 0.9201268

 $00{:}43{:}31{.}774$ --> $00{:}43{:}35{.}922$ It took about 18 months to completely NOTE Confidence: 0.9201268

 $00{:}43{:}35{.}922 \dashrightarrow 00{:}43{:}38{.}834$ melt out these pounds of cancer that

NOTE Confidence: 0.9201268

 $00:43:38.834 \rightarrow 00:43:41.516$ this patient had on the volume up.

NOTE Confidence: 0.9201268

00:43:41.520 --> 00:43:43.585 The patient is still doing is cure

NOTE Confidence: 0.9201268

00:43:43.585 --> 00:43:45.667 of the disease and I swear two

NOTE Confidence: 0.9201268

 $00{:}43{:}45.667 \dashrightarrow 00{:}43{:}47.365$ weeks ago in clinic eight years

NOTE Confidence: 0.9201268

00:43:47.430 --> 00:43:49.356 after what I'm showing you here.

NOTE Confidence: 0.9201268

 $00{:}43{:}49{.}360 \dashrightarrow 00{:}43{:}51{.}748$ So this was a terminal patient

NOTE Confidence: 0.9201268

 $00{:}43{:}51.748 \dashrightarrow 00{:}43{:}54.559$ referred to us for again Hospice.

NOTE Confidence: 0.9201268

 $00:43:54.560 \longrightarrow 00:43:56.000$ We sequenced the tumor,

NOTE Confidence: 0.9201268

 $00:43:56.000 \rightarrow 00:43:58.160$ we found that we are mutation,

- NOTE Confidence: 0.9201268
- $00:43:58.160 \longrightarrow 00:44:00.225$ we treat the immune checkpoint

 $00:44:00.225 \longrightarrow 00:44:01.877$ inhibitor as visit response

NOTE Confidence: 0.9201268

00:44:01.877 --> 00:44:04.160 and of course we reported this,

NOTE Confidence: 0.9201268

 $00:44:04.160 \longrightarrow 00:44:07.160$ this is the first report of

NOTE Confidence: 0.9201268

 $00:44:07.160 \longrightarrow 00:44:08.660$ the successful treatment.

NOTE Confidence: 0.9201268

00:44:08.660 --> 00:44:11.444 Of a polyultra
mutated endometrial

NOTE Confidence: 0.9201268

 $00{:}44{:}11{.}444 \dashrightarrow 00{:}44{:}13{.}970$ carcinoma and now the word know

NOTE Confidence: 0.9201268

 $00:44:13.970 \longrightarrow 00:44:16.331$ and everybody is looking to this

NOTE Confidence: 0.9201268

 $00:44:16.331 \longrightarrow 00:44:18.540$ patient and when they found the

NOTE Confidence: 0.9201268

00:44:18.540 --> 00:44:19.948 polyultra
mutation for their treatment

NOTE Confidence: 0.9201268

 $00{:}44{:}19{.}948 \dashrightarrow 00{:}44{:}21{.}860$ with immune checkpoint inhibitor.

NOTE Confidence: 0.9201268

00:44:21.860 --> 00:44:24.037 Now the second tumor that I want

NOTE Confidence: 0.9201268

 $00{:}44{:}24.037 \dashrightarrow 00{:}44{:}26.540$ to tell you in the last 5-6 minutes

NOTE Confidence: 0.9201268

 $00{:}44{:}26{.}540 \dashrightarrow 00{:}44{:}28{.}844$ are carcinos or Comma, the uterus.

NOTE Confidence: 0.9201268

 $00{:}44{:}28{.}844 \dashrightarrow 00{:}44{:}30{.}140$ Carcinos or Comma,

 $00:44:30.140 \longrightarrow 00:44:32.690$ the uterus are also biologically

NOTE Confidence: 0.9201268

00:44:32.690 --> 00:44:35.468 aggressive are rare 3 to 5%.

NOTE Confidence: 0.9201268

00:44:35.468 --> 00:44:38.430 But they account for over 20% of

NOTE Confidence: 0.9201268

 $00:44:38.430 \longrightarrow 00:44:40.730$ all death in uterine carcinoma.

NOTE Confidence: 0.9201268

 $00{:}44{:}40{.}730 \dashrightarrow 00{:}44{:}43{.}430$ And one of the peculiarity about

NOTE Confidence: 0.9201268

00:44:43.430 --> 00:44:45.603 carcinofacoma of the uterine is

NOTE Confidence: 0.9201268

 $00{:}44{:}45{.}603 \dashrightarrow 00{:}44{:}48{.}186$ that for over 150 years this tumor

NOTE Confidence: 0.9201268

 $00:44:48.186 \longrightarrow 00:44:50.730$ have been a subject of debate.

NOTE Confidence: 0.9201268

 $00:44:50.730 \longrightarrow 00:44:51.930$ And why is that?

NOTE Confidence: 0.9201268

 $00:44:51.930 \longrightarrow 00:44:53.730$ Is because they are mixed human,

NOTE Confidence: 0.9201268

 $00{:}44{:}53{.}730 \dashrightarrow 00{:}44{:}56{.}208$ there is both an epithelial component

NOTE Confidence: 0.9201268

 $00{:}44{:}56{.}210 \dashrightarrow 00{:}44{:}58{.}688$ as well as a sarcometus component.

NOTE Confidence: 0.9201268

 $00{:}44{:}58.690 \dashrightarrow 00{:}45{:}01.175$ And the question is what are these

NOTE Confidence: 0.9201268

 $00:45:01.175 \rightarrow 00:45:03.387$ mixed cancers? Are they carcinoma?

NOTE Confidence: 0.9201268

 $00{:}45{:}03{.}387 \dashrightarrow 00{:}45{:}05{.}383$ That undergoes are comatus

NOTE Confidence: 0.9201268

 $00{:}45{:}05{.}383 \dashrightarrow 00{:}45{:}08{.}061$ transformation or are sarcoma that

- NOTE Confidence: 0.9201268
- $00:45:08.061 \rightarrow 00:45:10.265$ undergo epitilla differentiation such

 $00{:}45{:}10{.}265 \dashrightarrow 00{:}45{:}13{.}130$ As for instance synovial sarcoma.

NOTE Confidence: 0.9201268

 $00:45:13.130 \longrightarrow 00:45:15.450$ So we answer that question.

NOTE Confidence: 0.9201268

 $00:45:15.450 \longrightarrow 00:45:17.634$ This is another paper that we

NOTE Confidence: 0.9201268

 $00:45:17.634 \rightarrow 00:45:19.090$ published few years ago.

NOTE Confidence: 0.9201268

00:45:19.090 --> 00:45:22.018 Again we sequenced over 70 carcino

NOTE Confidence: 0.9201268

 $00:45:22.018 \rightarrow 00:45:25.281$ sarcoma of the uterus and we published

NOTE Confidence: 0.9201268

 $00:45:25.281 \rightarrow 00:45:27.903$ again this high impact journal and

NOTE Confidence: 0.9201268

 $00{:}45{:}27{.}903 \dashrightarrow 00{:}45{:}30{.}527$ what we found here is number one

NOTE Confidence: 0.9201268

 $00:45:30.530 \rightarrow 00:45:33.670$ we were able through sequencing.

NOTE Confidence: 0.9201268

 $00{:}45{:}33.670 \dashrightarrow 00{:}45{:}36.910$ We also perform multiple sequencing,

NOTE Confidence: 0.9201268

 $00:45:36.910 \longrightarrow 00:45:39.570$ so not only we sequence the 70

NOTE Confidence: 0.9201268

 $00:45:39.570 \longrightarrow 00:45:41.905$ specimen from the 70 patient but

NOTE Confidence: 0.9201268

 $00{:}45{:}41{.}905 \dashrightarrow 00{:}45{:}44{.}509$ about five or six of this tumor.

NOTE Confidence: 0.9201268

 $00{:}45{:}44{.}510 \dashrightarrow 00{:}45{:}47{.}054$ We were doing macro dissection and

 $00{:}45{:}47.054 \dashrightarrow 00{:}45{:}49.217$ sequence area of epithelial tumor

NOTE Confidence: 0.9201268

 $00{:}45{:}49{.}217 \dashrightarrow 00{:}45{:}51{.}197$ and other sarcometus humor tumor

NOTE Confidence: 0.9201268

00:45:51.197 --> 00:45:53.873 in the same patient and we were

NOTE Confidence: 0.9201268

 $00{:}45{:}53.873 \dashrightarrow 00{:}45{:}55.678$ overlapping the old Axon sequencing

NOTE Confidence: 0.9201268

 $00{:}45{:}55{.}678 \dashrightarrow 00{:}45{:}58{.}015$ data to try to understand are

NOTE Confidence: 0.9201268

00:45:58.015 --> 00:46:00.140 these tumor epithelial or this

NOTE Confidence: 0.9201268

 $00{:}46{:}00{.}140 \dashrightarrow 00{:}46{:}03{.}294$ tumor mesenkimal and the answer is.

NOTE Confidence: 0.9201268

 $00:46:03.294 \rightarrow 00:46:03.870$ Unequivocally,

NOTE Confidence: 0.9201268

 $00{:}46{:}03{.}870 \dashrightarrow 00{:}46{:}06{.}470$ these are all epithelial cancer.

NOTE Confidence: 0.9201268

 $00:46:06.470 \longrightarrow 00:46:08.546$ They all start as an epithelia,

NOTE Confidence: 0.9201268

00:46:08.550 --> 00:46:09.584 uterine carcinoma,

NOTE Confidence: 0.9201268

 $00:46:09.584 \rightarrow 00:46:13.203$ but they go through clonal evolution and

NOTE Confidence: 0.9201268

 $00:46:13.203 \rightarrow 00:46:16.467$ part of the clone during the lifespan,

NOTE Confidence: 0.95534864

 $00{:}46{:}16{.}470 \dashrightarrow 00{:}46{:}18{.}762$ lifetime of the disease.

NOTE Confidence: 0.95534864

00:46:18.762 --> 00:46:19.908 They differentiate.

NOTE Confidence: 0.95534864

 $00:46:19.910 \longrightarrow 00:46:21.167$ They acquire epithelium,

- NOTE Confidence: 0.95534864
- $00{:}46{:}21.167 \dashrightarrow 00{:}46{:}22.424$ mesenchemal transition and
- NOTE Confidence: 0.95534864
- 00:46:22.424 --> 00:46:24.670 there's as I'm showing you here,
- NOTE Confidence: 0.95534864
- $00{:}46{:}24.670 \dashrightarrow 00{:}46{:}27.652$ some of these carcinos or coma acquire
- NOTE Confidence: 0.95534864
- $00{:}46{:}27.652 \dashrightarrow 00{:}46{:}29.391$ this epithelium mesenchemal transition
- NOTE Confidence: 0.95534864
- $00:46:29.391 \rightarrow 00:46:32.156$ in some of the clone relatively early.
- NOTE Confidence: 0.95534864
- $00:46:32.160 \longrightarrow 00:46:34.878$ Some very late during their lifetime.
- NOTE Confidence: 0.95534864
- $00{:}46{:}34.880 \dashrightarrow 00{:}46{:}37.295$ The 2nd and very important finding of
- NOTE Confidence: 0.95534864
- $00:46:37.295 \longrightarrow 00:46:39.787$ the study is that we have identify
- NOTE Confidence: 0.95534864
- $00{:}46{:}39.787 \dashrightarrow 00{:}46{:}42.140$ for the first time some recurrent
- NOTE Confidence: 0.95534864
- 00:46:42.140 --> 00:46:45.653 mutation in Eastern core gene that
- NOTE Confidence: 0.95534864
- $00{:}46{:}45{.}653 \dashrightarrow 00{:}46{:}47{.}718$ regulate the transcription of the
- NOTE Confidence: 0.95534864
- 00:46:47.720 --> 00:46:51.848 DNA and these were correlated with
- NOTE Confidence: 0.95534864
- $00{:}46{:}51{.}848 \dashrightarrow 00{:}46{:}53{.}912$ the mesenchemal transformation.
- NOTE Confidence: 0.95534864
- $00{:}46{:}53{.}920 \dashrightarrow 00{:}46{:}55{.}840$ Now we are clinician, right.
- NOTE Confidence: 0.95534864
- $00{:}46{:}55{.}840 \dashrightarrow 00{:}46{:}57{.}718$ So we are very interesting science,
- NOTE Confidence: 0.95534864

00:46:57.720 --> 00:46:59.995 but what we want is translation science,

NOTE Confidence: 0.95534864

 $00{:}47{:}00{.}000 \dashrightarrow 00{:}47{:}02{.}223$ we want to be able to help our patient.

NOTE Confidence: 0.95534864

 $00:47:02.230 \longrightarrow 00:47:04.222$ One of the thing that we found in

NOTE Confidence: 0.95534864

00:47:04.222 --> 00:47:06.210 carcinofer Com as well as you trying

NOTE Confidence: 0.95534864

00:47:06.210 --> 00:47:07.910 serous carcinoma doing RNA sequencing

NOTE Confidence: 0.95534864

 $00:47:07.910 \rightarrow 00:47:10.430$ was that there was a appregulation

NOTE Confidence: 0.95534864

00:47:10.430 --> 00:47:13.974 of the messenger RNA encoding for a

NOTE Confidence: 0.95534864

 $00:47:13.974 \rightarrow 00:47:16.074$ specific antigen called trophoblast

NOTE Confidence: 0.95534864

 $00{:}47{:}16.074 \dashrightarrow 00{:}47{:}18.628$ cells or face antigen or Trop 2.

NOTE Confidence: 0.95534864

 $00{:}47{:}18.630 \dashrightarrow 00{:}47{:}20.870$ And this is the work of two of

NOTE Confidence: 0.95534864

 $00{:}47{:}20.870 \dashrightarrow 00{:}47{:}22.189$ our recent fellow Dr.

NOTE Confidence: 0.95534864

 $00{:}47{:}22.190 \dashrightarrow 00{:}47{:}24.174$ Borazebek and Shannen Han.

NOTE Confidence: 0.95534864

 $00{:}47{:}24.174 \dashrightarrow 00{:}47{:}26.158$ So we look to.

NOTE Confidence: 0.95534864

 $00:47:26.160 \longrightarrow 00:47:27.978$ Hundreds of this tumor and here

NOTE Confidence: 0.95534864

 $00{:}47{:}27{.}978 \dashrightarrow 00{:}47{:}30{.}290$ of course we have done this with

NOTE Confidence: 0.95534864

 $00:47:30.290 \longrightarrow 00:47:32.366$ the help of our pathologist in

- NOTE Confidence: 0.95534864
- $00{:}47{:}32{.}366 \dashrightarrow 00{:}47{:}34{.}640$ particular Doctor Natalia Busa and Dr.
- NOTE Confidence: 0.95534864
- $00{:}47{:}34.640 \dashrightarrow 00{:}47{:}35.520$ Pei Wei.
- NOTE Confidence: 0.95534864
- $00{:}47{:}35{.}520 \dashrightarrow 00{:}47{:}36{.}400$ We did,
- NOTE Confidence: 0.95534864
- $00{:}47{:}36{.}400 \dashrightarrow 00{:}47{:}39{.}732$ we did tissue microarray and we found
- NOTE Confidence: 0.95534864
- $00{:}47{:}39{.}732 \dashrightarrow 00{:}47{:}43{.}456$ that indeed 60 to 90% of uterine
- NOTE Confidence: 0.95534864
- $00{:}47{:}43.456 \dashrightarrow 00{:}47{:}46.528$ serous carcinoma and the carcinosar coma,
- NOTE Confidence: 0.95534864
- $00:47:46.528 \rightarrow 00:47:48.256$ they were really overexpressed
- NOTE Confidence: 0.95534864
- $00{:}47{:}48.256 \dashrightarrow 00{:}47{:}49.440$ in tropoblast too.
- NOTE Confidence: 0.95534864
- $00:47:49.440 \longrightarrow 00:47:51.664$ So in the way that I show you
- NOTE Confidence: 0.95534864
- $00:47:51.664 \rightarrow 00:47:53.765$ before we generate patient that
- NOTE Confidence: 0.95534864
- $00{:}47{:}53.765 \dashrightarrow 00{:}47{:}56.215$ like sinograph from this patient.
- NOTE Confidence: 0.95534864
- $00:47:56.220 \rightarrow 00:47:59.279$ And we start testing in the animal
- NOTE Confidence: 0.95534864
- $00{:}47{:}59{.}279 \dashrightarrow 00{:}48{:}02{.}260$ the activity of a new agent called NOTE Confidence: 0.95534864
- $00{:}48{:}02{.}260 \dashrightarrow 00{:}48{:}04{.}984$ Rodelvi that is now a commercial NOTE Confidence: 0.95534864
- 00:48:04.984 --> 00:48:07.629 agent approved for breast as well
- NOTE Confidence: 0.95534864

 $00{:}48{:}07.629 \dashrightarrow 00{:}48{:}09.504$ as bladder cancer that target

NOTE Confidence: 0.95534864

 $00:48:09.504 \longrightarrow 00:48:11.420$ the DROP 2 receptor.

NOTE Confidence: 0.95534864

 $00{:}48{:}11{.}420 \dashrightarrow 00{:}48{:}13{.}580$ And this is an antibody track conjugate.

NOTE Confidence: 0.95534864

00:48:13.580 --> 00:48:15.680 So it's a monoclonal antibody

NOTE Confidence: 0.95534864

 $00{:}48{:}15.680 \dashrightarrow 00{:}48{:}18.500$ targeting DROP 2 as a cleavable

NOTE Confidence: 0.95534864

 $00{:}48{:}18{.}500 \dashrightarrow 00{:}48{:}21{.}100$ linker and attached to the

NOTE Confidence: 0.95534864

 $00:48:21.100 \longrightarrow 00:48:24.454$ antibody there is SN38 that is.

NOTE Confidence: 0.95534864

00:48:24.454 --> 00:48:28.750 A topo isomerase 1 inhibitor and doing

NOTE Confidence: 0.95534864

 $00{:}48{:}28{.}750 \dashrightarrow 00{:}48{:}32{.}538$ that we were able to show that again

NOTE Confidence: 0.95534864

 $00:48:32.538 \longrightarrow 00:48:35.290$ both in vitro as well as in vivo,

NOTE Confidence: 0.95534864

 $00:48:35.290 \rightarrow 00:48:36.930$ this tumor were highly sensitive.

NOTE Confidence: 0.95534864

00:48:36.930 --> 00:48:38.757 This is actually the tumor in the

NOTE Confidence: 0.95534864

 $00:48:38.757 \longrightarrow 00:48:40.336$ animal that complete as you can

NOTE Confidence: 0.95534864

 $00{:}48{:}40{.}336 \dashrightarrow 00{:}48{:}41{.}818$ see disappear when our control of

NOTE Confidence: 0.95534864

00:48:41.818 --> 00:48:43.680 course we're growing and this is the

NOTE Confidence: 0.95534864

 $00:48:43.680 \longrightarrow 00:48:45.210$ overall survival of this animal.

- NOTE Confidence: 0.95534864
- 00:48:45.210 --> 00:48:47.786 So the question is PDX work,

 $00:48:47.786 \longrightarrow 00:48:50.082$ is it going to work in patient?

NOTE Confidence: 0.95534864

00:48:50.090 - 00:48:51.810 Let me give you an example of that.

NOTE Confidence: 0.95534864

 $00:48:51.810 \rightarrow 00:48:54.470$ This is again another of my patient.

NOTE Confidence: 0.95534864

 $00{:}48{:}54{.}470 \dashrightarrow 00{:}48{:}57{.}070$ As you can see are 74 years old

NOTE Confidence: 0.95534864

00:48:57.070 --> 00:48:59.284 uterine serous carcinoma treated with

NOTE Confidence: 0.95534864

 $00:48:59.284 \rightarrow 00:49:01.606$ a gold standard surgery followed by

NOTE Confidence: 0.95534864

00:49:01.606 --> 00:49:03.782 chemo by radiation recurring disease,

NOTE Confidence: 0.95534864

00:49:03.782 --> 00:49:06.274 more chemo to the point where the

NOTE Confidence: 0.95534864

 $00{:}49{:}06{.}274$ --> $00{:}49{:}08{.}349$ patient was progressing on anything.

NOTE Confidence: 0.95534864

 $00{:}49{:}08.350 \dashrightarrow 00{:}49{:}11.680$ So at that time we have a Phase 1B

NOTE Confidence: 0.95534864

 $00{:}49{:}11.680 \dashrightarrow 00{:}49{:}14.830$ trial ongoing a Yale with Trodelvi,

NOTE Confidence: 0.95534864

 $00{:}49{:}14.830 \dashrightarrow 00{:}49{:}18.150$ so as you know calcium sarcoma rare tumor.

NOTE Confidence: 0.95534864

 $00{:}49{:}18.150 \dashrightarrow 00{:}49{:}20.796$ So I had to convince the the

NOTE Confidence: 0.95534864

 $00{:}49{:}20.796 \dashrightarrow 00{:}49{:}22.819$ medical monitor of the study.

- $00:49:22.820 \rightarrow 00:49:23.504$ That time,
- NOTE Confidence: 0.95534864
- $00{:}49{:}23.504 \dashrightarrow 00{:}49{:}25.560$ at that time was Immunomedics to
- NOTE Confidence: 0.95534864
- $00:49:25.560 \longrightarrow 00:49:27.260$ approve the treatment of this,
- NOTE Confidence: 0.95534864
- $00{:}49{:}27{.}260 \dashrightarrow 00{:}49{:}29{.}624$ you know of this uterine sero carcinoma
- NOTE Confidence: 0.95534864
- $00:49:29.624 \rightarrow 00:49:32.900$ patient rare tumor in the trial.
- NOTE Confidence: 0.95534864
- $00{:}49{:}32{.}900 \dashrightarrow 00{:}49{:}35{.}860$ But I was able to do so patients
- NOTE Confidence: 0.95534864
- 00:49:35.860 --> 00:49:36.600 start receiving
- NOTE Confidence: 0.926333050909091
- 00:49:36.682 --> 00:49:39.015 Trudelvi and again as you can see here,
- NOTE Confidence: 0.926333050909091
- $00{:}49{:}39{.}020 \dashrightarrow 00{:}49{:}40{.}220$ this is the baseline.
- NOTE Confidence: 0.926333050909091
- $00:49:40.220 \rightarrow 00:49:43.132$ She had huge liver, meth as well as
- NOTE Confidence: 0.926333050909091
- $00:49:43.132 \rightarrow 00:49:44.844$ carcinomatosis everywhere in the
- NOTE Confidence: 0.926333050909091
- $00{:}49{:}44.844 \dashrightarrow 00{:}49{:}48.452$ donor cavity and the patient start
- NOTE Confidence: 0.926333050909091
- $00{:}49{:}48{.}452 \dashrightarrow 00{:}49{:}51{.}140$ responding and we gave this patient.
- NOTE Confidence: 0.926333050909091
- $00{:}49{:}51{.}140 \dashrightarrow 00{:}49{:}53{.}618$ As you can see here dramatic shrinkage
- NOTE Confidence: 0.926333050909091
- $00:49:53.618 \longrightarrow 00:49:56.842$ of all the disease over 66% reduction
- NOTE Confidence: 0.926333050909091
- $00:49:56.842 \longrightarrow 00:50:01.092$ by this is 1.1 criteria in the in the

- NOTE Confidence: 0.926333050909091
- $00:50:01.092 \rightarrow 00:50:03.321$ inner disease and we have this patient
- NOTE Confidence: 0.926333050909091
- $00:50:03.321 \longrightarrow 00:50:05.883$ ten month of very good life with
- NOTE Confidence: 0.926333050909091
- $00:50:05.883 \rightarrow 00:50:08.454$ again because this is also very well
- NOTE Confidence: 0.926333050909091
- $00:50:08.454 \rightarrow 00:50:11.740$ tolerated antibody in term of treatment.
- NOTE Confidence: 0.926333050909091
- $00:50:11.740 \longrightarrow 00:50:14.740$ So this was one patient, right.
- NOTE Confidence: 0.926333050909091
- $00:50:14.740 \longrightarrow 00:50:16.315$ You cannot change a guideline
- NOTE Confidence: 0.926333050909091
- $00:50:16.315 \longrightarrow 00:50:17.260$ with one patient.
- NOTE Confidence: 0.926333050909091
- $00:50:17.260 \longrightarrow 00:50:19.072$ You need to design the proper
- NOTE Confidence: 0.926333050909091
- $00:50:19.072 \longrightarrow 00:50:20.856$ study and that is what we've done.
- NOTE Confidence: 0.926333050909091
- $00:50:20.860 \longrightarrow 00:50:22.580$ We had designed an IIT,
- NOTE Confidence: 0.926333050909091
- $00:50:22.580 \longrightarrow 00:50:24.445$ so an investigator in shady
- NOTE Confidence: 0.926333050909091
- $00{:}50{:}24.445 \dashrightarrow 00{:}50{:}25.937$ trial with his agent,
- NOTE Confidence: 0.926333050909091
- 00:50:25.940 --> 00:50:27.625 not easy because Immuno Medics
- NOTE Confidence: 0.926333050909091
- 00:50:27.625 --> 00:50:29.380 was purchased by Gilead, right.
- NOTE Confidence: 0.926333050909091
- $00{:}50{:}29{.}380 \dashrightarrow 00{:}50{:}31{.}660$ So that create a little bit of a problem.
- NOTE Confidence: 0.926333050909091

00:50:31.660 -> 00:50:34.348 But we were able to convince or so

NOTE Confidence: 0.926333050909091

 $00:50:34.348 \rightarrow 00:50:37.354$ Gilead to provide us with drug and some

NOTE Confidence: 0.926333050909091

 $00{:}50{:}37{.}354 \dashrightarrow 00{:}50{:}40{.}206$ money to design and of course perform

NOTE Confidence: 0.926333050909091

 $00:50:40.206 \longrightarrow 00:50:43.485$ this phase two clinker trial with

NOTE Confidence: 0.926333050909091

 $00{:}50{:}43{.}485 \ldots > 00{:}50{:}45{.}900$ the ultimate goal to evaluate again a

NOTE Confidence: 0.926333050909091

 $00:50:45.900 \rightarrow 00:50:48.080$ response rate as a primary objective.

NOTE Confidence: 0.926333050909091

 $00:50:48.080 \rightarrow 00:50:51.278$ As well as overall overall survival,

NOTE Confidence: 0.926333050909091

 $00:50:51.280 \rightarrow 00:50:53.960$ progression free survival and the

NOTE Confidence: 0.926333050909091

 $00:50:53.960 \rightarrow 00:50:56.640$ safety as a secondary objective.

NOTE Confidence: 0.926333050909091

 $00:50:56.640 \longrightarrow 00:50:58.494$ And these are the result of

NOTE Confidence: 0.926333050909091

 $00:50:58.494 \rightarrow 00:51:00.411$ this trial that I'm going to

NOTE Confidence: 0.926333050909091

00:51:00.411 - 00:51:02.560 present in about 10 days at ASCO.

NOTE Confidence: 0.926333050909091

 $00:51:02.560 \longrightarrow 00:51:04.140$ This is a presentation.

NOTE Confidence: 0.926333050909091

 $00:51:04.140 \longrightarrow 00:51:05.720$ This is a preview.

NOTE Confidence: 0.926333050909091

 $00{:}51{:}05{.}720 \dashrightarrow 00{:}51{:}08{.}798$ I shouldn't do that, but I'll do it anyway.

NOTE Confidence: 0.926333050909091

 $00:51:08.800 \longrightarrow 00:51:10.756$ And as you can see here,

 $00:51:10.760 \longrightarrow 00:51:13.360$ this study was designed as

NOTE Confidence: 0.926333050909091

 $00{:}51{:}13{.}360 \dashrightarrow 00{:}51{:}15{.}412$ a Simon two stage design.

NOTE Confidence: 0.926333050909091

 $00{:}51{:}15{.}412 \dashrightarrow 00{:}51{:}17{.}710$ That it means to enroll the

NOTE Confidence: 0.926333050909091

 $00:51:17.787 \longrightarrow 00:51:19.737$ 1st 21 a valuable patient,

NOTE Confidence: 0.926333050909091

 $00{:}51{:}19{.}740 \dashrightarrow 00{:}51{:}22{.}386$ see if there was any response and

NOTE Confidence: 0.926333050909091

00:51:22.386 --> 00:51:25.327 in case of three or more of PROCR,

NOTE Confidence: 0.926333050909091

 $00{:}51{:}25{.}327 \dashrightarrow 00{:}51{:}27{.}409$ so either complete or partial response

NOTE Confidence: 0.926333050909091

 $00{:}51{:}27{.}409 \dashrightarrow 00{:}51{:}29{.}794$ would have been able to move to the

NOTE Confidence: 0.926333050909091

 $00{:}51{:}29{.}794 \dashrightarrow 00{:}51{:}31{.}820$ second stage and enroll a total of 50.

NOTE Confidence: 0.926333050909091

 $00{:}51{:}31{.}820 \dashrightarrow 00{:}51{:}36{.}130$ So as you can see here over two third

NOTE Confidence: 0.926333050909091

 $00:51:36.130 \rightarrow 00:51:39.000$ of the patient enroll in this trial.

NOTE Confidence: 0.926333050909091

 $00{:}51{:}39{.}000 \dashrightarrow 00{:}51{:}41{.}648$ Where specifically uterine serous

NOTE Confidence: 0.926333050909091

 $00{:}51{:}41.648 \dashrightarrow 00{:}51{:}43.634$ carcinoma and carcinosarcoma,

NOTE Confidence: 0.926333050909091

 $00{:}51{:}43.640 \dashrightarrow 00{:}51{:}46.412$ the most difficult to treat uterine

NOTE Confidence: 0.926333050909091

 $00{:}51{:}46{.}412 \dashrightarrow 00{:}51{:}49{.}540$ carcinoma and we had seven response

 $00:51:49.540 \longrightarrow 00:51:53.162$ not three so 33 response rate either

NOTE Confidence: 0.926333050909091

 $00:51:53.162 \rightarrow 00:51:56.906$ CR or PR and so positive stage one and

NOTE Confidence: 0.926333050909091

 $00:51:56.906 \rightarrow 00:52:00.799$ we are now a rolling the stage two,

NOTE Confidence: 0.926333050909091

 $00:52:00.800 \rightarrow 00:52:03.103$ so another 30 patient to try to

NOTE Confidence: 0.926333050909091

 $00:52:03.103 \longrightarrow 00:52:05.937$ see in an overall of 50 patient.

NOTE Confidence: 0.926333050909091

 $00:52:05.940 \longrightarrow 00:52:07.980$ How effective this agent is be,

NOTE Confidence: 0.926333050909091

 $00{:}52{:}07{.}980 \dashrightarrow 00{:}52{:}10{.}038$ but there is no doubt that there

NOTE Confidence: 0.926333050909091

 $00:52:10.038 \rightarrow 00:52:12.328$ is a strong signal and again this

NOTE Confidence: 0.926333050909091

 $00{:}52{:}12{.}328 \dashrightarrow 00{:}52{:}14{.}740$ is an IIT trial developed at Yale.

NOTE Confidence: 0.926333050909091

 $00:52:14.740 \longrightarrow 00:52:16.934$ So I'd like to conclude analogy

NOTE Confidence: 0.926333050909091

 $00:52:16.934 \rightarrow 00:52:18.818$ and thanking a lot of people.

NOTE Confidence: 0.926333050909091

 $00{:}52{:}18.820 \dashrightarrow 00{:}52{:}21.430$ The particular Elena that is here

NOTE Confidence: 0.926333050909091

 $00{:}52{:}21{.}430 \dashrightarrow 00{:}52{:}23{.}832$ my cochief they attending of the

NOTE Confidence: 0.926333050909091

00:52:23.832 --> 00:52:26.130 Juan on
cology division that I listed

NOTE Confidence: 0.926333050909091

 $00:52:26.197 \rightarrow 00:52:29.280$ here and here in particular I want

NOTE Confidence: 0.926333050909091

 $00:52:29.280 \longrightarrow 00:52:32.685$ to to to emphasize the the

- NOTE Confidence: 0.926333050909091
- 00:52:32.685 --> 00:52:34.369 role of Doctor Schwartz.
- NOTE Confidence: 0.926333050909091
- $00{:}52{:}34{.}370 \dashrightarrow 00{:}52{:}37{.}580$ A mentor for all of us that has been work
- NOTE Confidence: 0.926333050909091
- 00:52:37.666 --> 00:52:40.570 here for over 50 years is just retire
- NOTE Confidence: 0.926333050909091
- $00:52:40.570 \rightarrow 00:52:43.330$ as well as of course the group here.
- NOTE Confidence: 0.926333050909091
- $00:52:43.330 \rightarrow 00:52:46.180$ The support staff and the clinical
- NOTE Confidence: 0.926333050909091
- $00:52:46.180 \longrightarrow 00:52:47.852$ trial coordinators again that
- NOTE Confidence: 0.926333050909091
- $00{:}52{:}47.852 \dashrightarrow 00{:}52{:}50.379$ I'm showing you here that are the
- NOTE Confidence: 0.926333050909091
- $00:52:50.379 \rightarrow 00:52:52.722$ reason why again we are successful
- NOTE Confidence: 0.926333050909091
- $00:52:52.722 \longrightarrow 00:52:54.632$ in provided to our patient
- NOTE Confidence: 0.943943513333333
- $00:52:54.706 \longrightarrow 00:52:56.616$ clinical trial because they are
- NOTE Confidence: 0.943943513333333
- $00:52:56.616 \longrightarrow 00:52:58.945$ the one that make this happen.
- NOTE Confidence: 0.943943513333333
- $00:52:58.945 \longrightarrow 00:53:01.435$ The clinical trial office in the
- NOTE Confidence: 0.943943513333333
- $00:53:01.435 \rightarrow 00:53:03.310$ division of gynecological oncology.
- NOTE Confidence: 0.943943513333333
- $00{:}53{:}03{.}310 \dashrightarrow 00{:}53{:}05{.}690$ And finally last man or least the
- NOTE Confidence: 0.943943513333333
- 00:53:05.690 --> 00:53:07.777 researcher the lab people that are
- NOTE Confidence: 0.943943513333333

 $00:53:07.777 \rightarrow 00:53:10.108$ doing they really the work with the

NOTE Confidence: 0.943943513333333

 $00:53:10.180 \longrightarrow 00:53:12.469$ animals we are to do the sequencing

NOTE Confidence: 0.943943513333333

 $00{:}53{:}12{.}470 \dashrightarrow 00{:}53{:}14{.}654$ and of course these are all as you

NOTE Confidence: 0.943943513333333

 $00:53:14.654 \rightarrow 00:53:16.793$ can see young they come they get

NOTE Confidence: 0.943943513333333

 $00:53:16.793 \rightarrow 00:53:19.495$ trained in my lab they and after they

NOTE Confidence: 0.943943513333333

 $00:53:19.495 \rightarrow 00:53:21.703$ leave to get their faculty position.

NOTE Confidence: 0.943943513333333

00:53:21.710 -> 00:53:25.306 So here is the Fania belona Dr.

NOTE Confidence: 0.943943513333333

 $00{:}53{:}25{.}310 \dashrightarrow 00{:}53{:}29{.}198$ Belona the lab chief of for that has been

NOTE Confidence: 0.943943513333333

 $00{:}53{:}29{.}198 \dashrightarrow 00{:}53{:}31{.}840$ working with me for the last 25 years.

NOTE Confidence: 0.943943513333333

 $00:53:31.840 \longrightarrow 00:53:33.562$ And of course she's of she's the

NOTE Confidence: 0.943943513333333

 $00{:}53{:}33{.}562 \dashrightarrow 00{:}53{:}35{.}476$ person that I have to thank the most

NOTE Confidence: 0.943943513333333

 $00:53:35.476 \rightarrow 00:53:37.443$ for the work that I just show you

NOTE Confidence: 0.943943513333333

 $00{:}53{:}37{.}443 \dashrightarrow 00{:}53{:}39{.}277$ and I thank you for your attention.

NOTE Confidence: 0.941371755555556

 $00:53:49.200 \rightarrow 00:53:50.919$ If there is any question of course we're here

NOTE Confidence: 0.9352219

 $00:53:56.200 \rightarrow 00:53:58.712$ SD trial are you looking at the level

NOTE Confidence: 0.9352219

 $00:53:58.712 \rightarrow 00:54:01.355$ of troop tube or was it just all comers

 $00:54:01.355 \rightarrow 00:54:03.200$ and if you can repeat the question.

NOTE Confidence: 0.9352219

 $00:54:03.200 \longrightarrow 00:54:06.984$ Yeah. So Dr. Reem is asking if the.

NOTE Confidence: 0.9352219

00:54:06.990 --> 00:54:09.066 The Trudelvi trial that I'm going

NOTE Confidence: 0.9352219

 $00{:}54{:}09{.}066 \dashrightarrow 00{:}54{:}11{.}220$ to presented ask whether we have

NOTE Confidence: 0.9352219

00:54:11.220 --> 00:54:13.810 look into the expression of the drop

NOTE Confidence: 0.9352219

 $00{:}54{:}13.881 \dashrightarrow 00{:}54{:}16.563$ two that is a target or not and the

NOTE Confidence: 0.9352219

 $00{:}54{:}16{.}563 \dashrightarrow 00{:}54{:}19{.}789$ answer David is yes in the phase one.

NOTE Confidence: 0.9352219

 $00{:}54{:}19.790 \dashrightarrow 00{:}54{:}22.910$ So the stage one of this trial we

NOTE Confidence: 0.9352219

 $00{:}54{:}22{.}910 \dashrightarrow 00{:}54{:}25{.}419$ specifically look to drop to expression

NOTE Confidence: 0.9352219

 $00{:}54{:}25{.}419 \dashrightarrow 00{:}54{:}28{.}159$ and only patient with 50 with expression

NOTE Confidence: 0.9352219

00:54:28.159 --> 00:54:31.090 or drop to any expression 1 + 2

NOTE Confidence: 0.9352219

 $00{:}54{:}31{.}090 \dashrightarrow 00{:}54{:}33{.}578$ plus or three plus in at least 50%.

NOTE Confidence: 0.9352219

 $00:54:33.580 \rightarrow 00:54:35.372$ Of the tumor cell we are eligible

NOTE Confidence: 0.9352219

 $00{:}54{:}35{.}372 \dashrightarrow 00{:}54{:}36{.}420$ for the stage one.

NOTE Confidence: 0.938815971428571

 $00{:}55{:}00{.}830 \dashrightarrow 00{:}55{:}04{.}180$ This is a good question again the so how

 $00{:}55{:}04{.}180 \dashrightarrow 00{:}55{:}06{.}105$ many of these tumor with recurring disease

NOTE Confidence: 0.938815971428571

 $00{:}55{:}06.105 \dashrightarrow 00{:}55{:}08.429$ they have estrogen or progesterone receptor.

NOTE Confidence: 0.938815971428571

 $00{:}55{:}08{.}430 \dashrightarrow 00{:}55{:}11{.}823$ So let me answer to your question this way.

NOTE Confidence: 0.938815971428571

 $00:55:11.830 \rightarrow 00:55:14.734$ UTRAN cancer is a very heterogeneous

NOTE Confidence: 0.938815971428571

 $00:55:14.734 \longrightarrow 00:55:17.360$ type of tumor, so you have.

NOTE Confidence: 0.938815971428571

 $00:55:17.360 \longrightarrow 00:55:19.302$ From one side, the endometrioid,

NOTE Confidence: 0.938815971428571

 $00:55:19.302 \rightarrow 00:55:22.004$ so the one that are really resembling

NOTE Confidence: 0.938815971428571

 $00:55:22.004 \rightarrow 00:55:24.039$ very closely the normal normal

NOTE Confidence: 0.938815971428571

 $00{:}55{:}24.039 \dashrightarrow 00{:}55{:}26.009$ endometrium that can be well,

NOTE Confidence: 0.938815971428571

 $00:55:26.010 \rightarrow 00:55:27.526$ moderately or poorly differentiated.

NOTE Confidence: 0.938815971428571

 $00{:}55{:}27{.}526 \dashrightarrow 00{:}55{:}29{.}800$ And on the other side of

NOTE Confidence: 0.938815971428571

00:55:29.871 - > 00:55:31.486 the spectrum you have the,

NOTE Confidence: 0.938815971428571

 $00:55:31.490 \longrightarrow 00:55:34.129$ what we call still called type 2,

NOTE Confidence: 0.938815971428571

 $00:55:34.130 \rightarrow 00:55:36.370$ right, the uterine serous carcinoma,

NOTE Confidence: 0.938815971428571

 $00:55:36.370 \dashrightarrow 00:55:39.610$ the carcinosarcoma, the clear cell,

NOTE Confidence: 0.938815971428571

 $00:55:39.610 \longrightarrow 00:55:41.878$ you know uterine cancer,

 $00:55:41.878 \longrightarrow 00:55:43.579$ so in general.

NOTE Confidence: 0.938815971428571

 $00:55:43.580 \rightarrow 00:55:46.058$ If you want you to endometrioid they

NOTE Confidence: 0.938815971428571

 $00{:}55{:}46.058 \dashrightarrow 00{:}55{:}48.339$ over express a strogen receptor even when

NOTE Confidence: 0.938815971428571

 $00:55:48.339 \rightarrow 00:55:50.655$ they come back with recurring disease.

NOTE Confidence: 0.938815971428571

 $00{:}55{:}50.660 \dashrightarrow 00{:}55{:}52.484$ But those are only a minority

NOTE Confidence: 0.938815971428571

 $00{:}55{:}52{.}484 \dashrightarrow 00{:}55{:}54{.}046$ of the recurring patient that

NOTE Confidence: 0.938815971428571

 $00:55:54.046 \longrightarrow 00:55:55.738$ we are treating in our clinic.

NOTE Confidence: 0.938815971428571

 $00:55:55.740 \longrightarrow 00:55:57.170$ Majority of the recurrences as

NOTE Confidence: 0.938815971428571

 $00{:}55{:}57{.}170 \dashrightarrow 00{:}55{:}59{.}311$ I show you they take place in

NOTE Confidence: 0.938815971428571

00:55:59.311 -> 00:56:00.619 the uterine serious group,

NOTE Confidence: 0.938815971428571

 $00{:}56{:}00{.}620 \dashrightarrow 00{:}56{:}03{.}910$ the carcinoso coma group that are the

NOTE Confidence: 0.938815971428571

 $00{:}56{:}03{.}910 \dashrightarrow 00{:}56{:}07{.}406$ one that usually they do not express

NOTE Confidence: 0.938815971428571

 $00{:}56{:}07{.}406 \dashrightarrow 00{:}56{:}09{.}872$ as trogen receptor or the minority

NOTE Confidence: 0.938815971428571

 $00{:}56{:}09{.}872 \dashrightarrow 00{:}56{:}13{.}308$ even when they do I must tell you.

NOTE Confidence: 0.938815971428571

 $00{:}56{:}13.308 \dashrightarrow 00{:}56{:}14.796$ By 20 year experience,

 $00:56:14.800 \longrightarrow 00:56:17.176$ they do not respond to anti

NOTE Confidence: 0.938815971428571

 $00:56:17.176 \longrightarrow 00:56:18.760$ estrogen treatment because most

NOTE Confidence: 0.938815971428571

 $00:56:18.834 \rightarrow 00:56:21.120$ likely the receptor is not active.

NOTE Confidence: 0.938815971428571

 $00:56:21.120 \rightarrow 00:56:22.536$ It's present on the surface but

NOTE Confidence: 0.938815971428571

 $00:56:22.536 \rightarrow 00:56:24.199$ the pathway is not really working

NOTE Confidence: 0.9402536

 $00{:}56{:}29{.}910 \dashrightarrow 00{:}56{:}30{.}230$ sure.

NOTE Confidence: 0.912391133333333

 $00{:}56{:}45{.}460 \dashrightarrow 00{:}56{:}47{.}773$ So what is what are the side effect of

NOTE Confidence: 0.912391133333333

00:56:47.773 --> 00:56:49.609 the trudel
vides assitu Zumagovita can

NOTE Confidence: 0.912391133333333

 $00:56:49.609 \rightarrow 00:56:52.308$ trial that is using this top poisoner

NOTE Confidence: 0.912391133333333

 $00:56:52.308 \rightarrow 00:56:54.456$ is inhibitor as a toxic payload.

NOTE Confidence: 0.912391133333333

 $00:56:54.460 \longrightarrow 00:56:56.860$ So the answer is this is a very

NOTE Confidence: 0.912391133333333

 $00{:}56{:}56{.}860 \dashrightarrow 00{:}56{:}58{.}979$ well tolerated drug and why am I

NOTE Confidence: 0.912391133333333

 $00:56:58.979 \rightarrow 00:57:01.598$ saying that is because it does not

NOTE Confidence: 0.912391133333333

 $00:57:01.598 \rightarrow 00:57:04.191$ cause the bone marrow toxicity

NOTE Confidence: 0.912391133333333

 $00:57:04.191 \longrightarrow 00:57:06.453$ that you know they can cause.

NOTE Confidence: 0.912391133333333

 $00:57:06.460 \longrightarrow 00:57:08.567$ And the other thing that we have

 $00:57:08.567 \rightarrow 00:57:10.795$ noticed is that even the GI toxicity

NOTE Confidence: 0.912391133333333

 $00:57:10.795 \longrightarrow 00:57:12.679$ that is typical of this drug.

NOTE Confidence: 0.912391133333333

00:57:12.680 --> 00:57:14.948 Is much, much smaller and this

NOTE Confidence: 0.912391133333333

 $00:57:14.948 \longrightarrow 00:57:17.792$ is once again is related to the

NOTE Confidence: 0.912391133333333

00:57:17.792 --> 00:57:19.980 targeted type of approach, right.

NOTE Confidence: 0.912391133333333

 $00:57:19.980 \longrightarrow 00:57:23.270$ So are only the tumor cell they

NOTE Confidence: 0.912391133333333

 $00{:}57{:}23.270 \dashrightarrow 00{:}57{:}26.034$ over express drop two that they get that

NOTE Confidence: 0.912391133333333

 $00:57:26.034 \rightarrow 00:57:29.240$ they get this toxic payload internalized.

NOTE Confidence: 0.912391133333333

 $00{:}57{:}29{.}240$ --> $00{:}57{:}31{.}816$ 1 important thing that I also am going NOTE Confidence: 0.912391133333333

 $00:57:31.816 \rightarrow 00:57:35.930$ to stress to ASCO is that the Cornell

NOTE Confidence: 0.912391133333333

 $00:57:35.930 \rightarrow 00:57:38.625$ toxicity for instance that we are seeing

NOTE Confidence: 0.912391133333333

 $00:57:38.625 \dashrightarrow 00:57:41.310$ with many antibody that are conjugate.

NOTE Confidence: 0.912391133333333

 $00{:}57{:}41{.}310 \dashrightarrow 00{:}57{:}43{.}626$ Is not present with this antibody.

NOTE Confidence: 0.912391133333333

 $00{:}57{:}43.630 \dashrightarrow 00{:}57{:}45.191$ So that is another great thing because

NOTE Confidence: 0.912391133333333

00:57:45.191 --> 00:57:47.312 as you know we're getting more and more NOTE Confidence: 0.912391133333333

 $00:57:47.312 \rightarrow 00:57:48.717$ approval in terrible treatment with

NOTE Confidence: 0.912391133333333

 $00:57:48.764 \rightarrow 00:57:50.546$ antibody that are conjugate our patient.

NOTE Confidence: 0.912391133333333

00:57:50.550 --> 00:57:52.602 But we must have now enough

NOTE Confidence: 0.912391133333333

 $00:57:52.602 \rightarrow 00:57:54.262$ ophthalmology before even starting to

NOTE Confidence: 0.912391133333333

 $00{:}57{:}54{.}262 \dashrightarrow 00{:}57{:}56{.}198$ look to the corner for this specific

NOTE Confidence: 0.912391133333333

 $00:57:56.198 \longrightarrow 00:57:58.550$ type of a DC is not necessary.

NOTE Confidence: 0.912391133333333

 $00:57:58.550 \longrightarrow 00:57:59.126$ We don't,

NOTE Confidence: 0.912391133333333

 $00:57:59.126 \rightarrow 00:58:01.142$ we don't really see that toxicity those.

NOTE Confidence: 0.912391133333333

 $00{:}58{:}01{.}150 \dashrightarrow 00{:}58{:}02{.}670$ So it's overall well tolerated.

NOTE Confidence: 0.912391133333333

 $00:58:02.670 \longrightarrow 00:58:06.126$ The only patient that are really

NOTE Confidence: 0.912391133333333

 $00:58:06.126 \rightarrow 00:58:08.430$ experiencing some significant toxicity

NOTE Confidence: 0.912391133333333

 $00{:}58{:}08{.}430 \dashrightarrow 00{:}58{:}10{.}957$ are the patient that have a polymorphism.

NOTE Confidence: 0.912391133333333

00:58:10.960 --> 00:58:16.078 In EU GT2B7 that is again this

NOTE Confidence: 0.912391133333333

 $00{:}58{:}16.080 \dashrightarrow 00{:}58{:}18.030$ is this mechanism right that we

NOTE Confidence: 0.912391133333333

 $00{:}58{:}18{.}030 \dashrightarrow 00{:}58{:}20{.}494$ have in our liver to catabolize at

NOTE Confidence: 0.912391133333333

 $00:58:20.494 \longrightarrow 00:58:22.319$ the top of isomerase about,

00:58:22.320 --> 00:58:26.515 I'll say 1-2 out of 10 May

NOTE Confidence: 0.912391133333333

00:58:26.515 --> 00:58:28.000 have this polymorphism.

NOTE Confidence: 0.912391133333333

 $00:58:28.000 \rightarrow 00:58:29.519$ They are more sensitive to the drug,

NOTE Confidence: 0.912391133333333

 $00:58:29.520 \rightarrow 00:58:31.716$ they usually respond better their cancer,

NOTE Confidence: 0.912391133333333

 $00{:}58{:}31.720 \dashrightarrow 00{:}58{:}33.592$ but there we have to do a dose reduction.

NOTE Confidence: 0.912391133333333

 $00:58:33.600 \rightarrow 00:58:35.760$ We go down from 10 usually to 7.5.

NOTE Confidence: 0.906658214

 $00:58:40.760 \longrightarrow 00:58:42.120$ We have two more answers

NOTE Confidence: 0.80246687

 $00{:}58{:}55{.}480 \dashrightarrow 00{:}58{:}55{.}600$ that

NOTE Confidence: 0.92767435

 $00{:}58{:}59{.}120 \dashrightarrow 00{:}59{:}04{.}160$ yes, so we I show you here our

NOTE Confidence: 0.92767435

 $00{:}59{:}04.160 \dashrightarrow 00{:}59{:}06.043$ initial work with trust to Zuma and

NOTE Confidence: 0.92767435

 $00:59:06.043 \dashrightarrow 00:59:08.010$ we have shown again for the first

NOTE Confidence: 0.92767435

 $00{:}59{:}08.010 \dashrightarrow 00{:}59{:}09.660$ time that trust to Zuma added.

NOTE Confidence: 0.92767435

 $00:59:09.660 \longrightarrow 00:59:12.444$ To chemotherapy helps but in the

NOTE Confidence: 0.92767435

 $00{:}59{:}12{.}444 \dashrightarrow 00{:}59{:}14{.}940$ recurrent setting we have multiple

NOTE Confidence: 0.92767435

 $00:59:14.940 \rightarrow 00:59:18.300$ trial using antibody that are conjugate.

 $00:59:18.300 \longrightarrow 00:59:24.220$ So we have used some one that

NOTE Confidence: 0.92767435

 $00{:}59{:}24{.}220 \dashrightarrow 00{:}59{:}28{.}640$ is called symptom 985 that is a

NOTE Confidence: 0.92767435

00:59:28.640 --> 00:59:30.600 I don't know if you're familiar

NOTE Confidence: 0.92767435

00:59:30.600 --> 00:59:33.884 but again is a is the backbone is NOTE Confidence: 0.92767435

 $00:59:33.884 \rightarrow 00:59:36.064$ throstozoma the linker is clivable.

NOTE Confidence: 0.92767435

 $00{:}59{:}36{.}064$ --> $00{:}59{:}38{.}530$ The toxic payload is an alkylating NOTE Confidence: 0.92767435

 $00:59:38.606 \rightarrow 00:59:40.790$ semi synthetic calculating agent

NOTE Confidence: 0.92767435

 $00:59:40.790 \rightarrow 00:59:43.520$ duocarmising extremely potent but also

NOTE Confidence: 0.92767435

 $00{:}59{:}43{.}520 \dashrightarrow 00{:}59{:}46{.}272$ toxic and so we have seen dramatic

NOTE Confidence: 0.92767435

 $00{:}59{:}46{.}272 \dashrightarrow 00{:}59{:}48{.}302$ response in the recurrence setting

NOTE Confidence: 0.92767435

00:59:48.302 --> 00:59:50.254 intrastuzumab resistant uterine 0

NOTE Confidence: 0.92767435

 $00{:}59{:}50{.}254 \dashrightarrow 00{:}59{:}53{.}030$ carcinoma over expressing up to new. NOTE Confidence: 0.92767435

 $00:59:53.030 \rightarrow 00:59:54.842$ But the corneltosis it was significant

NOTE Confidence: 0.92767435

 $00:59:54.842 \longrightarrow 00:59:57.335$ in many of this patient we have to

NOTE Confidence: 0.92767435

 $00{:}59{:}57{.}335$ --> $00{:}59{:}59{.}556$ stop treatment for for several weeks to NOTE Confidence: 0.92767435

 $00:59:59.556 \rightarrow 01:00:02.740$ allow recovery and that of course.

- NOTE Confidence: 0.92767435
- $01{:}00{:}02.740 \dashrightarrow 01{:}00{:}04.756$ Was not good because during those

01:00:04.756 --> 01:00:06.100 68 sometime more weeks,

NOTE Confidence: 0.92767435

 $01:00:06.100 \longrightarrow 01:00:07.336$ the tumor will start growing again.

NOTE Confidence: 0.92767435

 $01:00:07.340 \longrightarrow 01:00:07.460$ Yes,

NOTE Confidence: 0.856499678571429

 $01:00:22.210 \longrightarrow 01:00:23.568$ we do in a very special way.

NOTE Confidence: 0.856499678571429

 $01:00:23.570 \longrightarrow 01:00:24.848$ There's different ways to do it.

NOTE Confidence: 0.856499678571429

 $01:00:24.850 \longrightarrow 01:00:27.018$ She's doing it.

NOTE Confidence: 0.856499678571429

 $01:00:27.018 \longrightarrow 01:00:29.290$ So we only do it in the newest.

NOTE Confidence: 0.856499678571429

 $01:00:29.290 \longrightarrow 01:00:33.030$ So we already have the diagnosis.

NOTE Confidence: 0.856499678571429

 $01:00:33.030 \rightarrow 01:00:34.210$ We give 3 cycles. We

NOTE Confidence: 0.950317

 $01:00:36.490 \longrightarrow 01:00:36.890$ confirm the.

NOTE Confidence: 0.9251585

01:00:39.440 --> 01:00:40.280 But if it's

NOTE Confidence: 0.9587031666666667

 $01{:}00{:}42.360 \dashrightarrow 01{:}00{:}44.960$ done different ways, there's some

NOTE Confidence: 0.9587031666666667

 $01{:}00{:}44.960 \dashrightarrow 01{:}00{:}47.440$ studies that have it up front.

NOTE Confidence: 0.9402536

 $01:00:51.640 \rightarrow 01:00:52.040$ Thank you.

01:00:53.560 --> 01:00:56.540 Thank you so much to our NOTE Confidence: 0.9352219 01:00:56.540 --> 01:00:58.000 wonderful 2 speakers. NOTE Confidence: 0.9352219 01:00:58.000 --> 01:00:59.236 Have a wonderful day. Thank you.