## WEBVTT

 $00:00:00.000 \longrightarrow 00:00:03.186$  Funding for Yale Cancer Answers is

NOTE Confidence: 0.9380176

00:00:03.186 --> 00:00:06.200 provided by Smilow Cancer Hospital.

NOTE Confidence: 0.9380176

 $00:00:06.200 \longrightarrow 00:00:08.400$  Welcome to Yale Cancer Answers

NOTE Confidence: 0.9380176

 $00:00:08.400 \longrightarrow 00:00:10.160$  with Doctor Anees Chappar.

NOTE Confidence: 0.9380176

00:00:10.160 --> 00:00:11.656 Yale Cancer Answers features

NOTE Confidence: 0.9380176

00:00:11.656 --> 00:00:13.526 the latest information on cancer

NOTE Confidence: 0.9380176

00:00:13.526 --> 00:00:15.430 care by welcoming oncologists and

NOTE Confidence: 0.9380176

 $00:00:15.430 \longrightarrow 00:00:17.638$  specialists who are on the forefront

NOTE Confidence: 0.9380176

 $00:00:17.699 \longrightarrow 00:00:19.397$  of the battle to fight cancer.

NOTE Confidence: 0.9380176

 $00:00:19.400 \dashrightarrow 00:00:21.752$  This week it's a conversation about new

NOTE Confidence: 0.9380176

00:00:21.752 --> 00:00:24.026 research into the early detection of

NOTE Confidence: 0.9380176

 $00:00:24.026 \longrightarrow 00:00:26.426$  ovarian cancer with Doctor Stacy Malaker.

NOTE Confidence: 0.9380176

 $00:00:26.430 \longrightarrow 00:00:28.434$  Dr. Malaker is an assistant professor

NOTE Confidence: 0.9380176

 $00:00:28.434 \longrightarrow 00:00:30.238$  in the Department of Chemistry

NOTE Confidence: 0.9380176

 $00:00:30.238 \longrightarrow 00:00:32.084$  at Yale University, and Dr.

00:00:32.084 --> 00:00:33.926 Chagpar is a professor of Surgical

NOTE Confidence: 0.9380176

 $00:00:33.926 \dashrightarrow 00:00:36.188$  on cology at the Yale School of Medicine.

NOTE Confidence: 0.92948914

 $00:00:37.390 \longrightarrow 00:00:39.084$  So, Stacy, maybe we can start off

NOTE Confidence: 0.92948914

 $00:00:39.084 \longrightarrow 00:00:41.048$  by you telling us a little bit more

NOTE Confidence: 0.92948914

 $00:00:41.048 \longrightarrow 00:00:42.789$  about yourself and what it is you do.

NOTE Confidence: 0.92948914

 $00{:}00{:}43.870 \dashrightarrow 00{:}00{:}46.685$  I got my PhD at the University of Virginia

NOTE Confidence: 0.92948914

 $00:00:46.685 \longrightarrow 00:00:49.793$  where I was in the lab of Professor

NOTE Confidence: 0.92948914

00:00:49.793 --> 00:00:53.032 Donald Hunt and he is one of

NOTE Confidence: 0.92948914

 $00:00:53.032 \longrightarrow 00:00:55.078$  the founding fathers of biological mass

NOTE Confidence: 0.92948914

00:00:55.078 --> 00:00:57.766 spectrometry and mass spec is kind

NOTE Confidence: 0.92948914

 $00:00:57.766 \longrightarrow 00:01:00.415$  of what I do or what I'm known for.

NOTE Confidence: 0.92948914

 $00:01:00.420 \longrightarrow 00:01:03.280$  And then I did my postdoc in the

NOTE Confidence: 0.92948914

 $00:01:03.361 \longrightarrow 00:01:06.396$  the lab of Carolyn Bertozzi, who just

NOTE Confidence: 0.92948914

00:01:06.396 --> 00:01:09.812 recently won the Nobel Prize in Chemistry.

NOTE Confidence: 0.92948914

 $00:01:09.820 \longrightarrow 00:01:13.008$  And there I got really interested in

NOTE Confidence: 0.92948914

 $00:01:13.008 \longrightarrow 00:01:16.580$  a class of of proteins called mucins

 $00:01:16.580 \longrightarrow 00:01:19.408$  which have tons and tons of sugar

NOTE Confidence: 0.92948914

 $00:01:19.408 \longrightarrow 00:01:20.620$  units on them.

NOTE Confidence: 0.92948914

 $00:01:20.620 \longrightarrow 00:01:22.336$  And so I spent

NOTE Confidence: 0.92948914

 $00:01:22.340 \longrightarrow 00:01:24.180$  five years there researching those.

NOTE Confidence: 0.92948914

00:01:24.180 --> 00:01:27.260 And so now in my own laboratory,

NOTE Confidence: 0.92948914

 $00:01:27.260 \longrightarrow 00:01:30.152$  I combine the expertise of the

NOTE Confidence: 0.92948914

00:01:30.152 --> 00:01:32.636 instrumentation or the mass spec

NOTE Confidence: 0.92948914

 $00{:}01{:}32.636 \to 00{:}01{:}35.414$  and the sugars or glycobiology

NOTE Confidence: 0.92948914

 $00:01:35.420 \longrightarrow 00:01:37.436$  and we do something

NOTE Confidence: 0.92948914

00:01:37.436 --> 00:01:38.300 that's called glycoproteomics,

NOTE Confidence: 0.92948914

 $00:01:38.300 \longrightarrow 00:01:40.036$  which is studying sugars

NOTE Confidence: 0.92948914

 $00:01:40.036 \longrightarrow 00:01:41.338$  that modify proteins.

NOTE Confidence: 0.9306428

 $00{:}01{:}41.580 \dashrightarrow 00{:}01{:}43.848$  So now every body wants to know, what

NOTE Confidence: 0.9306428

 $00:01:43.848 \longrightarrow 00:01:46.815$  does any of this have to do with cancer?

NOTE Confidence: 0.9366184

 $00:01:47.920 \longrightarrow 00:01:50.110$  Sure. So sugars are altered in

00:01:50.110 --> 00:01:51.990 pretty much every disease that's

NOTE Confidence: 0.9366184

 $00:01:51.990 \longrightarrow 00:01:54.078$  ever been studied and

NOTE Confidence: 0.9366184

 $00:01:54.080 \longrightarrow 00:01:55.436$  primarily in cancer,

NOTE Confidence: 0.9366184

 $00:01:55.436 \longrightarrow 00:01:57.696$  but also other diseases like

NOTE Confidence: 0.9366184

 $00:01:57.696 \longrightarrow 00:01:59.826$  inflammatory bowel disease or cystic

NOTE Confidence: 0.9366184

 $00:01:59.826 \longrightarrow 00:02:01.788$  fibrosis or even heart disease.

NOTE Confidence: 0.9366184

 $00:02:01.788 \longrightarrow 00:02:04.164$  And so we try to monopolize

NOTE Confidence: 0.9366184

 $00:02:04.164 \longrightarrow 00:02:07.115$  on those changes in the sugar

NOTE Confidence: 0.9366184

00:02:07.115 --> 00:02:09.224 structures to identify

NOTE Confidence: 0.9366184

 $00:02:09.224 \longrightarrow 00:02:10.472$  new biomarkers or potential

NOTE Confidence: 0.9366184

 $00{:}02{:}10.472 --> 00{:}02{:}11.720 \ {\rm the rapeutics}.$ 

NOTE Confidence: 0.9366184

 $00:02:12.960 \longrightarrow 00:02:15.935$  Tell us more about your

NOTE Confidence: 0.9366184

 $00:02:15.935 \longrightarrow 00:02:18.086$  research in particular, what are you

NOTE Confidence: 0.9366184

 $00:02:18.086 \longrightarrow 00:02:20.130$  looking at and how might this make

NOTE Confidence: 0.9366184

 $00:02:20.194 \longrightarrow 00:02:22.366$  a difference to people with cancer?

NOTE Confidence: 0.934648

 $00:02:23.490 \longrightarrow 00:02:26.850$  Sure, this project in

 $00{:}02{:}26.850 \dashrightarrow 00{:}02{:}29.090$  particular regarding ovarian cancer,

NOTE Confidence: 0.934648

 $00:02:29.090 \longrightarrow 00:02:32.420$  right now more than 70% of women

NOTE Confidence: 0.934648

 $00:02:32.420 \longrightarrow 00:02:34.120$  are diagnosed with ovarian

NOTE Confidence: 0.934648

 $00:02:34.120 \longrightarrow 00:02:35.804$  cancer in the late stages,

NOTE Confidence: 0.934648

 $00:02:35.804 \longrightarrow 00:02:38.558$  so stage 3 or stage 4 and the five

NOTE Confidence: 0.934648

 $00:02:38.558 \longrightarrow 00:02:40.778$  year survival rate for women diagnosed

NOTE Confidence: 0.934648

 $00:02:40.778 \longrightarrow 00:02:43.049$  in those stages is really poor.

NOTE Confidence: 0.934648

 $00:02:43.050 \longrightarrow 00:02:45.170$  It's less than 20%.

NOTE Confidence: 0.934648

00:02:45.170 --> 00:02:47.685 Now if ovarian cancer is caught in

NOTE Confidence: 0.934648

00:02:47.685 --> 00:02:49.590 early stages like stage 1 or two,

NOTE Confidence: 0.934648

 $00:02:49.590 \dashrightarrow 00:02:52.026$  that five year survival rate goes up to 95%.

NOTE Confidence: 0.934648

 $00:02:52.026 \longrightarrow 00:02:53.958$  But the problem is that we don't

NOTE Confidence: 0.934648

 $00{:}02{:}53.958 \dashrightarrow 00{:}02{:}55.784$  have a really good biomarker

NOTE Confidence: 0.934648

 $00:02:55.784 \longrightarrow 00:02:57.789$  for ovarian cancer right now.

NOTE Confidence: 0.934648

00:02:57.790 --> 00:03:00.238 Right now what is currently used

 $00:03:00.238 \longrightarrow 00:03:02.251$  is something that's called CA-125

NOTE Confidence: 0.934648

 $00{:}03{:}02.251 \dashrightarrow 00{:}03{:}04.918$  and CA-125 happens to be one of

NOTE Confidence: 0.934648

00:03:04.918 --> 00:03:07.107 those mucin type proteins that

NOTE Confidence: 0.934648

00:03:07.107 --> 00:03:09.347 I was talking about earlier.

NOTE Confidence: 0.934648

 $00:03:09.350 \longrightarrow 00:03:11.343$  And so it's this really,

NOTE Confidence: 0.934648

00:03:11.343 --> 00:03:12.908 really huge protein that's decorated

NOTE Confidence: 0.934648

 $00:03:12.908 \longrightarrow 00:03:15.190$  by tons and tons and tons of sugars.

NOTE Confidence: 0.934648

 $00:03:15.190 \longrightarrow 00:03:18.700$  And so 80% of its mass is actually sugar

NOTE Confidence: 0.934648

 $00{:}03{:}18.700 \dashrightarrow 00{:}03{:}22.484$  units as opposed to the protein backbone.

NOTE Confidence: 0.934648

 $00:03:22.490 \longrightarrow 00:03:26.362$  Again, the sugar units

NOTE Confidence: 0.934648

 $00{:}03{:}26.362 {\:\dashrightarrow\:} 00{:}03{:}29.322$  are perpetually disordered in cancer

NOTE Confidence: 0.934648

00:03:29.322 --> 00:03:33.210 yet when doctors are detecting the CA-125,

NOTE Confidence: 0.934648

 $00:03:33.210 \longrightarrow 00:03:35.360$  they're usually only detecting the

NOTE Confidence: 0.934648

 $00:03:35.360 \longrightarrow 00:03:37.510$  unmodified regions of the protein.

NOTE Confidence: 0.934648

00:03:37.510 --> 00:03:40.331 And so we want to identify altered

NOTE Confidence: 0.934648

 $00:03:40.331 \longrightarrow 00:03:43.030$  sugar units on this huge protein

 $00:03:43.030 \longrightarrow 00:03:45.870$  to ideally detect cancer earlier.

NOTE Confidence: 0.934648

 $00:03:45.870 \longrightarrow 00:03:48.318$  So that if we can do that and identify

NOTE Confidence: 0.934648

 $00:03:48.318 \longrightarrow 00:03:49.658$  something that's changed early

NOTE Confidence: 0.934648

 $00:03:49.658 \longrightarrow 00:03:51.662$  on in the progression of cancer,

NOTE Confidence: 0.934648

 $00:03:51.670 \longrightarrow 00:03:54.166$  then we could ostensibly develop a

NOTE Confidence: 0.934648

 $00:03:54.166 \longrightarrow 00:03:56.989$  better biomarker and early stage detection.

NOTE Confidence: 0.9350412

 $00:03:58.070 \longrightarrow 00:04:01.146$  Yeah, I think

NOTE Confidence: 0.9350412

 $00:04:01.146 \longrightarrow 00:04:03.210$  the problem though is

NOTE Confidence: 0.9350412

 $00:04:03.210 \longrightarrow 00:04:05.274$  that for ovarian cancer,

NOTE Confidence: 0.9350412

 $00:04:05.280 \longrightarrow 00:04:07.720$  it's not incredibly common.

NOTE Confidence: 0.9350412

 $00:04:07.720 \longrightarrow 00:04:09.757$  You're quite right, when it is diagnosed,

NOTE Confidence: 0.9350412

 $00:04:09.760 \longrightarrow 00:04:11.460$  it's diagnosed late because we

NOTE Confidence: 0.9350412

 $00:04:11.460 \longrightarrow 00:04:13.160$  don't have a screening test.

NOTE Confidence: 0.9350412

00:04:13.160 --> 00:04:16.198 But one of the questions always is,

NOTE Confidence: 0.9350412

 $00:04:16.200 \longrightarrow 00:04:18.380$  you know, are there blood

 $00:04:18.380 \longrightarrow 00:04:20.560$  tests for detection of cancer?

NOTE Confidence: 0.9350412

 $00:04:20.560 \longrightarrow 00:04:22.756$  Are there blood tests for screening?

NOTE Confidence: 0.9350412

 $00:04:22.760 \longrightarrow 00:04:25.476$  And while CA-125 is a biomarker that

NOTE Confidence: 0.9350412

00:04:25.476 --> 00:04:29.128 might be used to help doctors in terms

NOTE Confidence: 0.9350412

00:04:29.128 --> 00:04:31.533 of monitoring progression of disease,

NOTE Confidence: 0.9350412

 $00{:}04{:}31.540 \dashrightarrow 00{:}04{:}34.340$  it's really not a widespread

NOTE Confidence: 0.9350412

00:04:34.340 --> 00:04:37.140 screening tool like for example,

NOTE Confidence: 0.9350412

 $00:04:37.140 \longrightarrow 00:04:39.036$  a colaquard would be or

NOTE Confidence: 0.9350412

00:04:39.036 --> 00:04:40.300 a mammogram would be.

NOTE Confidence: 0.9350412

00:04:40.300 --> 00:04:42.939 So is your research trying to look

NOTE Confidence: 0.9350412

 $00{:}04{:}42.939 \dashrightarrow 00{:}04{:}45.260$  at these altered sugar moieties,

NOTE Confidence: 0.9350412

00:04:45.260 --> 00:04:47.780 really trying to find a screening modality?

NOTE Confidence: 0.9350412

00:04:47.780 --> 00:04:48.896 And if so,

NOTE Confidence: 0.9350412

 $00:04:48.896 \longrightarrow 00:04:50.756$  would that be administered on

NOTE Confidence: 0.9350412

 $00:04:50.756 \longrightarrow 00:04:52.760$  a population basis like to all

NOTE Confidence: 0.9350412

00:04:52.760 --> 00:04:55.007 women or would it be for women

00:04:55.007 --> 00:04:57.419 who are particularly at high risk?

NOTE Confidence: 0.9268508

 $00{:}04{:}58.540 \dashrightarrow 00{:}05{:}00.654$  So that's a great question and I

NOTE Confidence: 0.9268508

 $00:05:00.654 \longrightarrow 00:05:02.820$  think that as a basic scientist,

NOTE Confidence: 0.9268508

 $00:05:02.820 \longrightarrow 00:05:05.660$  I can only say that I'm

NOTE Confidence: 0.9268508

 $00:05:05.660 \longrightarrow 00:05:07.346$  hopeful that we'll be able

NOTE Confidence: 0.9268508

 $00:05:07.346 \longrightarrow 00:05:08.782$  to identify something that has

NOTE Confidence: 0.9268508

 $00:05:08.782 \longrightarrow 00:05:10.057$  changed early on in cancer.

NOTE Confidence: 0.9268508

00:05:10.060 --> 00:05:13.580 So we're using serum from high risk patients,

NOTE Confidence: 0.9268508

 $00{:}05{:}13.580 \dashrightarrow 00{:}05{:}17.048$  some of whom developed ovarian cancer.

NOTE Confidence: 0.9268508

 $00:05:17.050 \longrightarrow 00:05:18.986$  And so the idea would be that we

NOTE Confidence: 0.9268508

 $00{:}05{:}18.986 \dashrightarrow 00{:}05{:}20.502$  do identify something that could

NOTE Confidence: 0.9268508

 $00:05:20.502 \longrightarrow 00:05:22.446$  be used as a screening modality,

NOTE Confidence: 0.9268508

 $00{:}05{:}22.450 \dashrightarrow 00{:}05{:}24.442$  but I don't want to make any early

NOTE Confidence: 0.9268508

 $00{:}05{:}24.442 \dashrightarrow 00{:}05{:}26.010$  promises since we haven't actually,

NOTE Confidence: 0.9268508

 $00:05:26.010 \longrightarrow 00:05:27.930$  you know, identified anything quite yet.

 $00:05:28.210 \longrightarrow 00:05:29.715$  Tell us a little bit

NOTE Confidence: 0.93441564

00:05:29.715 --> 00:05:30.810 more about your project.

NOTE Confidence: 0.93441564

00:05:30.810 --> 00:05:32.736 I mean, when you say you're

NOTE Confidence: 0.93441564

 $00:05:32.736 \longrightarrow 00:05:34.490$  looking at high risk women,

NOTE Confidence: 0.93441564

 $00:05:34.490 \longrightarrow 00:05:37.208$  you tell us more about who those women are.

NOTE Confidence: 0.93441564

 $00{:}05{:}37.210 \dashrightarrow 00{:}05{:}39.964$  And the concept that you kind of laid out,

NOTE Confidence: 0.93441564

 $00:05:39.970 \longrightarrow 00:05:41.490$  if I've understood it correctly,

NOTE Confidence: 0.93441564

 $00:05:41.490 \longrightarrow 00:05:42.890$  is that you're looking

NOTE Confidence: 0.93441564

 $00{:}05{:}42.890 \dashrightarrow 00{:}05{:}44.290$  at these high-risk women.

NOTE Confidence: 0.93441564

00:05:44.290 --> 00:05:47.220 You're taking blood samples from

NOTE Confidence: 0.93441564

 $00:05:47.220 \longrightarrow 00:05:49.615$  them and comparing those of them

NOTE Confidence: 0.93441564

 $00:05:49.615 \longrightarrow 00:05:51.958$  who went on to truly develop

NOTE Confidence: 0.93441564

 $00:05:51.958 \longrightarrow 00:05:54.730$  ovarian cancer to those who didn't?

NOTE Confidence: 0.93441564

 $00:05:54.730 \longrightarrow 00:05:56.370$  Is that right?

NOTE Confidence: 0.928521

 $00:05:56.370 \longrightarrow 00:05:57.930$  That's basically correct.

NOTE Confidence: 0.928521

 $00:05:57.930 \longrightarrow 00:06:01.570$  So we have access to approximately 4000

 $00:06:01.650 \longrightarrow 00:06:04.370$  serum samples from high-risk women.

NOTE Confidence: 0.928521

 $00:06:04.370 \longrightarrow 00:06:05.855$  These are women that have

NOTE Confidence: 0.928521

 $00:06:05.855 \longrightarrow 00:06:07.496$  been diagnosed with the BRCA,

NOTE Confidence: 0.928521

 $00:06:07.496 \longrightarrow 00:06:09.280$  one or two mutations.

NOTE Confidence: 0.928521

 $00:06:09.280 \dashrightarrow 00:06:12.836$  So from the point of genetic diagnosis,

NOTE Confidence: 0.928521

 $00:06:12.840 \longrightarrow 00:06:15.759$  you know throughout the years many,

NOTE Confidence: 0.928521

 $00:06:15.760 \longrightarrow 00:06:17.315$  many samples have been collected

NOTE Confidence: 0.928521

 $00:06:17.315 \longrightarrow 00:06:18.559$  from these various women.

NOTE Confidence: 0.928521

 $00:06:18.560 \longrightarrow 00:06:21.185$  And so to kind of develop our

NOTE Confidence: 0.928521

 $00:06:21.185 \longrightarrow 00:06:23.164$  technology we're using women that

NOTE Confidence: 0.928521

 $00:06:23.164 \longrightarrow 00:06:25.109$  have not actually been diagnosed

NOTE Confidence: 0.928521

 $00:06:25.109 \longrightarrow 00:06:27.800$  just to be able to identify the

NOTE Confidence: 0.928521

 $00{:}06{:}27.800 \dashrightarrow 00{:}06{:}29.905$  CA-125 modifications or sugar units

NOTE Confidence: 0.928521

 $00{:}06{:}29.905 \dashrightarrow 00{:}06{:}32.493$  and then we'd basically be given

NOTE Confidence: 0.928521

 $00:06:32.493 \longrightarrow 00:06:34.533$  a blinded sample and hopefully

00:06:34.533 --> 00:06:36.596 identify those biomarkers

NOTE Confidence: 0.928521

 $00:06:36.596 \longrightarrow 00:06:39.476$  or what have you that could indicate

NOTE Confidence: 0.928521

 $00{:}06{:}39.476 \dashrightarrow 00{:}06{:}42.010$  cancer versus non cancerous samples.

NOTE Confidence: 0.91922235

 $00:06:42.770 \longrightarrow 00:06:46.970$  And so that sounds really interesting

NOTE Confidence: 0.91922235

 $00:06:46.970 \longrightarrow 00:06:49.730$  when we think about BRC A1 and two

NOTE Confidence: 0.91922235

 $00:06:49.730 \longrightarrow 00:06:52.255$  often times we think not only of

NOTE Confidence: 0.91922235

 $00:06:52.255 \longrightarrow 00:06:54.520$  ovarian cancer but also of breast

NOTE Confidence: 0.91922235

 $00:06:54.520 \longrightarrow 00:06:57.125$  cancer and one of the questions that

NOTE Confidence: 0.91922235

 $00{:}06{:}57.125 --> 00{:}07{:}00.565$  is often asked is, is there a

NOTE Confidence: 0.91922235

 $00:07:00.565 \longrightarrow 00:07:03.369$  blood test for breast cancer as well.

NOTE Confidence: 0.91922235

 $00{:}07{:}03.370 \longrightarrow 00{:}07{:}05.520$  You mentioned earlier that the

NOTE Confidence: 0.91922235

 $00{:}07{:}05.520 \longrightarrow 00{:}07{:}08.454$  sugar moieties tend to be, you know,

NOTE Confidence: 0.91922235

 $00:07:08.454 \longrightarrow 00:07:10.414$  involved or disrupted or altered

NOTE Confidence: 0.91922235

 $00{:}07{:}10.414 \dashrightarrow 00{:}07{:}12.589$  in a variety of processes.

NOTE Confidence: 0.91922235

 $00:07:12.590 \longrightarrow 00:07:14.510$  Do you think that your technology

NOTE Confidence: 0.91922235

 $00{:}07{:}14.510 \dashrightarrow 00{:}07{:}17.141$  might have a role to play in breast

 $00:07:17.141 \longrightarrow 00:07:19.061$  cancer as well as ovarian cancer?

NOTE Confidence: 0.91922235

 $00{:}07{:}19.070 \dashrightarrow 00{:}07{:}21.632$  Or is it really something specific about

NOTE Confidence: 0.91922235

00:07:21.632 --> 00:07:23.828 ovarian cancer that you're looking at?

NOTE Confidence: 0.93097055

00:07:24.510 --> 00:07:27.149 It's pretty much any epithelial cancer,

NOTE Confidence: 0.93097055

 $00:07:27.150 \longrightarrow 00:07:29.035$  you know, has these altered

NOTE Confidence: 0.93097055

 $00:07:29.035 \longrightarrow 00:07:30.994$  muc in structures and so

NOTE Confidence: 0.93097055

00:07:30.994 --> 00:07:35.390 CA-125 is known as Mucin 16 or Mach 16.

NOTE Confidence: 0.93097055

 $00:07:35.390 \longrightarrow 00:07:38.742$  Mucin one or Mach one is dysregulated or

NOTE Confidence: 0.93097055

 $00{:}07{:}38.742 \dashrightarrow 00{:}07{:}41.557$  upregulated in over 90% of breast carcinomas.

NOTE Confidence: 0.93097055

 $00{:}07{:}41.557 \dashrightarrow 00{:}07{:}44.131$  So this could ostensibly be extended

NOTE Confidence: 0.93097055

 $00:07:44.131 \longrightarrow 00:07:46.158$  to other cancers.

NOTE Confidence: 0.93097055

 $00:07:46.158 \longrightarrow 00:07:48.224$  Pancreatic cancer is another one that

NOTE Confidence: 0.93097055

 $00:07:48.224 \longrightarrow 00:07:50.065$  would be really interesting to look at.

NOTE Confidence: 0.93097055

 $00:07:50.070 \longrightarrow 00:07:52.326$  Pretty much any epithelial cancer is

NOTE Confidence: 0.93097055

 $00:07:52.326 \longrightarrow 00:07:53.830$  associated with dysregulated mucins.

 $00:07:54.830 \longrightarrow 00:07:58.350$  And so presumably in this population

NOTE Confidence: 0.93529564

 $00{:}07{:}58.350 \dashrightarrow 00{:}08{:}02.287$  of BRCA one and two gene mutation carriers,

NOTE Confidence: 0.93529564

 $00:08:02.290 \longrightarrow 00:08:05.134$  you'd be able to see not only the

NOTE Confidence: 0.93529564

 $00:08:05.134 \longrightarrow 00:08:07.102$  comparison between those who developed

NOTE Confidence: 0.93529564

 $00:08:07.102 \longrightarrow 00:08:09.810$  ovarian cancer and those who did not,

NOTE Confidence: 0.93529564

 $00:08:09.810 \longrightarrow 00:08:11.987$  but also those who developed breast cancer

NOTE Confidence: 0.93529564

 $00:08:11.987 \longrightarrow 00:08:14.114$  or in fact pancreatic cancer because

NOTE Confidence: 0.93529564

 $00:08:14.114 \longrightarrow 00:08:16.746$  that's another cancer that tends to be

NOTE Confidence: 0.93529564

 $00{:}08{:}16.808 \to 00{:}08{:}18.970$  associated with those mutations, right?

NOTE Confidence: 0.93121064

00:08:19.410 --> 00:08:21.622 Yeah, absolutely. I would have to talk

NOTE Confidence: 0.93121064

 $00{:}08{:}21.622 \dashrightarrow 00{:}08{:}23.740$  to my collaborators to see how many

NOTE Confidence: 0.93121064

 $00:08:23.740 \longrightarrow 00:08:25.408$  of these women actually did develop

NOTE Confidence: 0.93121064

 $00{:}08{:}25.465 \dashrightarrow 00{:}08{:}27.210$  breast and or pancreatic cancer.

NOTE Confidence: 0.93121064

 $00:08:27.210 \longrightarrow 00:08:28.645$  But that could be done.

NOTE Confidence: 0.93182003

 $00:08:29.600 \longrightarrow 00:08:32.048$  So you know one of the things when we

NOTE Confidence: 0.93182003

00:08:32.048 --> 00:08:34.477 think about that kind of an experiment,

 $00:08:34.480 \longrightarrow 00:08:35.775$  one would think that time

NOTE Confidence: 0.93182003

00:08:35.775 --> 00:08:37.280 has something to do with it,

NOTE Confidence: 0.93182003

 $00:08:37.280 \longrightarrow 00:08:41.072$  right that it takes time to develop

NOTE Confidence: 0.93182003

00:08:41.072 --> 00:08:43.837 these alterations in the protein

NOTE Confidence: 0.93182003

 $00{:}08{:}43.837 \dashrightarrow 00{:}08{:}46.435$  structure or in the sugar structure

NOTE Confidence: 0.93182003

 $00{:}08{:}46.440 \dashrightarrow 00{:}08{:}48.358$  and it takes time to develop cancer.

NOTE Confidence: 0.93182003

 $00:08:48.360 \longrightarrow 00:08:53.328$  So have you found any correlation

NOTE Confidence: 0.93182003

 $00{:}08{:}53.328 \rightarrow 00{:}08{:}56.520$  between the the timing of things,

NOTE Confidence: 0.93182003

 $00{:}08{:}56.520 \dashrightarrow 00{:}08{:}59.131$  I mean presumably if somebody just gets

NOTE Confidence: 0.93182003

 $00{:}08{:}59.131 \dashrightarrow 00{:}09{:}02.856$ a blood sample today and you know

NOTE Confidence: 0.93182003

 $00:09:02.856 \dashrightarrow 00:09:05.187$  and then isn't followed for very long,

NOTE Confidence: 0.93182003

 $00:09:05.190 \longrightarrow 00:09:07.188$  you may not find an association.

NOTE Confidence: 0.9339164

00:09:08.350 --> 00:09:10.445 Yeah, that's a really great point.

NOTE Confidence: 0.9339164

00:09:10.445 --> 00:09:12.790 And you know this is we're very,

NOTE Confidence: 0.9339164

 $00:09:12.790 \longrightarrow 00:09:14.750$  very, very early on in this project.

 $00:09:14.750 \longrightarrow 00:09:17.190$  It was just awarded a few months ago.

NOTE Confidence: 0.9339164

 $00{:}09{:}17.190 \dashrightarrow 00{:}09{:}19.254$  And so I anticipate we will

NOTE Confidence: 0.9339164

 $00{:}09{:}19.254 \dashrightarrow 00{:}09{:}21.030$  actually see changes over time.

NOTE Confidence: 0.9339164

 $00:09:21.030 \longrightarrow 00:09:22.986$  But because again

NOTE Confidence: 0.9339164

 $00:09:22.986 \longrightarrow 00:09:24.677$  we haven't actually done much

NOTE Confidence: 0.9339164

00:09:24.677 --> 00:09:26.267 of the research quite yet,

NOTE Confidence: 0.9339164

 $00:09:26.270 \longrightarrow 00:09:27.827$  I can't give you a straight answer to that.

NOTE Confidence: 0.9314148

 $00:09:28.310 \longrightarrow 00:09:32.896$  But of these 4000 women,

NOTE Confidence: 0.9314148

 $00{:}09{:}32.896 \longrightarrow 00{:}09{:}36.794$  are you kind of looking at these women

NOTE Confidence: 0.9314148

 $00:09:36.794 \longrightarrow 00:09:40.224$  going forward as well or is this kind of

NOTE Confidence: 0.9314148

 $00:09:40.224 \longrightarrow 00:09:42.606$  a deidentified mass sample that you've

NOTE Confidence: 0.9314148

 $00:09:42.606 \longrightarrow 00:09:45.602$  got where you've got some clinical

NOTE Confidence: 0.9314148

 $00{:}09{:}45.602 \dashrightarrow 00{:}09{:}48.085$  correlation data and would have to

NOTE Confidence: 0.9314148

 $00:09:48.085 \longrightarrow 00:09:50.762$  use covariates to see whether a

NOTE Confidence: 0.9314148

 $00:09:50.762 \longrightarrow 00:09:52.389$  relationship existed. Fo example,

NOTE Confidence: 0.9314148

 $00:09:52.389 \longrightarrow 00:09:54.147$  looking at age as a surrogate.

00:09:55.610 --> 00:09:57.050 OK. So just to clarify,

NOTE Confidence: 0.9352205

00:09:57.050 --> 00:09:58.494 it's not 4000 women,

NOTE Confidence: 0.9352205

 $00:09:58.494 \longrightarrow 00:10:00.660$  it's 4000 samples that have been

NOTE Confidence: 0.9352205

 $00:10:00.729 \longrightarrow 00:10:02.969$  collected from I think 50 to 100

NOTE Confidence: 0.9352205

 $00:10:02.969 \dashrightarrow 00:10:05.089$  women over the course of their life.

NOTE Confidence: 0.9352205

 $00:10:07.890 \longrightarrow 00:10:09.265$  I see, so then you're comparing samples

NOTE Confidence: 0.9352205

 $00:10:09.265 \longrightarrow 00:10:10.970$  as you go along in time.

NOTE Confidence: 0.9352205

 $00:10:10.970 \longrightarrow 00:10:14.102$  So there might be out of the

NOTE Confidence: 0.9352205

00:10:14.102 --> 00:10:17.690 4000, say 100 people,

NOTE Confidence: 0.9352205

 $00:10:17.690 \longrightarrow 00:10:19.888$  then that would be like 40 time

NOTE Confidence: 0.9352205

 $00{:}10{:}19.888 \mathrel{\text{--}}{>} 00{:}10{:}21.609$  points per person on average,

NOTE Confidence: 0.9352205

 $00:10:21.850 \longrightarrow 00:10:22.770$  something like that.

NOTE Confidence: 0.94081473

00:10:24.060 --> 00:10:25.544 So then that's very cool, right,

NOTE Confidence: 0.94081473

 $00:10:25.544 \longrightarrow 00:10:27.488$  because then you could see whether

NOTE Confidence: 0.94081473

 $00:10:27.488 \longrightarrow 00:10:29.720$  these people are

00:10:29.720 --> 00:10:32.380 acquiring these mutations.

NOTE Confidence: 0.94081473

 $00:10:32.380 \longrightarrow 00:10:35.430$  Exactly, exactly.

NOTE Confidence: 0.94081473

 $00:10:35.430 \longrightarrow 00:10:37.970$  So now that makes a lot more

NOTE Confidence: 0.94081473

00:10:37.970 --> 00:10:40.455 sense because now you can actually see,

NOTE Confidence: 0.94081473

00:10:40.460 --> 00:10:43.228 you know, how long does it take for

NOTE Confidence: 0.94081473

 $00{:}10{:}43.228 \operatorname{--}{>} 00{:}10{:}45.933$  people to develop these alterations and

NOTE Confidence: 0.94081473

 $00:10:45.933 \longrightarrow 00:10:49.380$  do these alterations once they do occur,

NOTE Confidence: 0.94081473

00:10:49.380 --> 00:10:53.324 how quickly or not do people develop cancer?

NOTE Confidence: 0.94081473

 $00:10:53.330 \longrightarrow 00:10:54.968$  Is that kind of the idea?

NOTE Confidence: 0.94081473

00:10:54.970 --> 00:10:56.410 Yes, precisely.

NOTE Confidence: 0.94081473

 $00:10:56.410 \longrightarrow 00:10:57.874$  Yeah, that's very cool.

NOTE Confidence: 0.94081473

 $00:10:57.874 \longrightarrow 00:10:59.610$  So tell us a little bit more.

NOTE Confidence: 0.94081473

 $00:10:59.610 \longrightarrow 00:11:01.170$  I realized that this is

NOTE Confidence: 0.94081473

 $00{:}11{:}01.170 \dashrightarrow 00{:}11{:}03.170$ a fresh project,

NOTE Confidence: 0.94081473

 $00:11:03.170 \longrightarrow 00:11:05.690$  hot off the presses, just awarded.

NOTE Confidence: 0.94081473

 $00:11:05.690 \longrightarrow 00:11:07.573$  Tell us about some of the research

 $00:11:07.573 \longrightarrow 00:11:09.650$  that kind of led up to this award.

NOTE Confidence: 0.94081473

 $00:11:09.650 \longrightarrow 00:11:12.368$  What have you found in your

NOTE Confidence: 0.94081473

00:11:12.368 --> 00:11:13.727 more earlier studies?

NOTE Confidence: 0.924282204

 $00:11:15.690 \longrightarrow 00:11:18.510$  When I was a post doc,

NOTE Confidence: 0.924282204

 $00{:}11{:}18.510 \dashrightarrow 00{:}11{:}21.079$  when we do mass spectrometry we

NOTE Confidence: 0.924282204

00:11:21.079 --> 00:11:24.161 usually take a protein and we digest

NOTE Confidence: 0.924282204

00:11:24.161 --> 00:11:26.903 it using enzymes into short peptides

NOTE Confidence: 0.924282204

00:11:26.903 --> 00:11:30.065 and then you know we basically blast

NOTE Confidence: 0.924282204

 $00:11:30.065 \longrightarrow 00:11:32.825$  those apart by bombarding them

NOTE Confidence: 0.924282204

 $00:11:32.825 \longrightarrow 00:11:35.444$  with gas molecules and/or radical

NOTE Confidence: 0.924282204

00:11:35.444 --> 00:11:37.796 anions and by the way that they

NOTE Confidence: 0.924282204

 $00:11:37.796 \longrightarrow 00:11:40.216$  fall apart we can kind of piece back

NOTE Confidence: 0.924282204

 $00{:}11{:}40.216 \dashrightarrow 00{:}11{:}42.210$  what was present there previously.

NOTE Confidence: 0.924282204

 $00:11:42.210 \longrightarrow 00:11:43.930$  But one of the problems with these really,

NOTE Confidence: 0.924282204

00:11:43.930 --> 00:11:45.830 really densely like oscillated proteins

00:11:45.830 --> 00:11:48.074 or you know sugar modified proteins

NOTE Confidence: 0.924282204

 $00:11:48.074 \longrightarrow 00:11:50.290$  is that they can't be chopped up by

NOTE Confidence: 0.924282204

 $00:11:50.290 \longrightarrow 00:11:52.527$  the normal enzymes that we would use.

NOTE Confidence: 0.924282204

 $00:11:52.530 \longrightarrow 00:11:55.806$  And so when I was in my postdoc I

NOTE Confidence: 0.924282204

 $00:11:55.806 \longrightarrow 00:11:57.810$  characterized a series of enzymes that

NOTE Confidence: 0.924282204

 $00:11:57.810 \longrightarrow 00:12:00.248$  we call mucineases that are actually able

NOTE Confidence: 0.924282204

 $00{:}12{:}00.248 \dashrightarrow 00{:}12{:}02.810$  to create short segments of the protein

NOTE Confidence: 0.924282204

 $00:12:02.810 \longrightarrow 00:12:05.448$  that are amenable to mass spec analysis.

NOTE Confidence: 0.924282204

 $00:12:05.450 \longrightarrow 00:12:07.515$  So before we couldn't look at these

NOTE Confidence: 0.924282204

00:12:07.515 --> 00:12:10.170 at all by my instrumentation method,

NOTE Confidence: 0.924282204

 $00:12:10.170 \longrightarrow 00:12:12.242$  but now we can actually get pieces and

NOTE Confidence: 0.924282204

 $00:12:12.242 \longrightarrow 00:12:14.580$  see them in the in the mass spectrometer.

NOTE Confidence: 0.93528324

 $00:12:14.740 \longrightarrow 00:12:17.511$  So why is that important?

NOTE Confidence: 0.93528324

 $00{:}12{:}17.511 \dashrightarrow 00{:}12{:}20.430$  Why is looking at these with mass

NOTE Confidence: 0.93528324

 $00:12:20.529 \longrightarrow 00:12:22.415$  spec so important as opposed to

NOTE Confidence: 0.93528324

 $00:12:22.415 \longrightarrow 00:12:24.620$  looking at them with other techniques?

 $00:12:24.620 \longrightarrow 00:12:26.040$  Or are there no other

NOTE Confidence: 0.93528324

 $00:12:26.040 \longrightarrow 00:12:27.460$  techniques to look at them?

NOTE Confidence: 0.93528324

00:12:28.380 --> 00:12:32.068 I mean, you could potentially

NOTE Confidence: 0.93528324

00:12:32.068 --> 00:12:36.450 use staining techniques, or NOTE Confidence:

0.93528324

 $00:12:36.450 \longrightarrow 00:12:38.430$  certain other techniques.

NOTE Confidence: 0.93528324

 $00:12:38.430 \longrightarrow 00:12:39.470$  I'm not saying that mass

NOTE Confidence: 0.93528324

 $00:12:39.470 \longrightarrow 00:12:40.510$  spec is the only technique.

NOTE Confidence: 0.93528324

00:12:40.510 --> 00:12:43.667 However, in my opinion,

NOTE Confidence: 0.93528324

 $00:12:43.670 \longrightarrow 00:12:45.070$  and of course I'm biased,

NOTE Confidence: 0.93528324

00:12:45.070 --> 00:12:47.401 it's the best way of actually digging

NOTE Confidence: 0.93528324

 $00:12:47.401 \longrightarrow 00:12:49.711$  into what sugar structures are modifying

NOTE Confidence: 0.93528324

 $00:12:49.711 \longrightarrow 00:12:52.189$  what amino acids in what patterns.

NOTE Confidence: 0.93528324

 $00:12:52.190 \longrightarrow 00:12:54.006$  And you're not going to get that molecular

NOTE Confidence: 0.93528324

00:12:54.006 --> 00:12:55.707 level of detail using other methods.

NOTE Confidence: 0.9232246

 $00:12:57.280 \longrightarrow 00:13:00.112$  So one of the things

00:13:00.112 --> 00:13:02.513 that you did before embarking on

NOTE Confidence: 0.9232246

 $00:13:02.513 \longrightarrow 00:13:05.384$  this was to figure out how you

NOTE Confidence: 0.9232246

 $00:13:05.384 \longrightarrow 00:13:07.604$  could actually use mass spec to

NOTE Confidence: 0.9232246

 $00:13:07.604 \longrightarrow 00:13:10.025$  look at at these sugar moieties

NOTE Confidence: 0.9232246

00:13:10.025 --> 00:13:12.200 in these proteins going forward.

NOTE Confidence: 0.9232246

00:13:12.760 --> 00:13:14.596 Precisely, yes. And so my lab,

NOTE Confidence: 0.9232246

 $00:13:14.600 \longrightarrow 00:13:16.796$  you know, I have kind of two arms in

NOTE Confidence: 0.9232246

00:13:16.796 --> 00:13:19.872 my laboratory, one being, you know,

NOTE Confidence: 0.9232246

 $00{:}13{:}19.872 \dashrightarrow 00{:}13{:}21.426$  instrumentation development and

NOTE Confidence: 0.9232246

00:13:21.426 --> 00:13:24.038 method development so that we can

NOTE Confidence: 0.9232246

 $00{:}13{:}24.038 \dashrightarrow 00{:}13{:}25.966$  better see these altered sugar

NOTE Confidence: 0.9232246

 $00:13:25.966 \longrightarrow 00:13:27.858$  structures and various diseases.

NOTE Confidence: 0.9232246

 $00:13:27.860 \longrightarrow 00:13:30.276$  And then another arm where we study

NOTE Confidence: 0.9232246

 $00:13:30.276 \longrightarrow 00:13:32.060$  the biological role of the altered,

NOTE Confidence: 0.9232246

00:13:32.060 --> 00:13:34.220 glycosylation patterns in

NOTE Confidence: 0.9232246

 $00:13:34.220 \longrightarrow 00:13:35.540$  cellular systems.

 $00:13:36.220 \longrightarrow 00:13:37.966$  Fantastic. Well, we're going to take

NOTE Confidence: 0.9379816

 $00:13:37.966 \longrightarrow 00:13:40.099$  a short break for a medical minute,

NOTE Confidence: 0.9379816

 $00:13:40.100 \longrightarrow 00:13:41.840$  but please stay tuned to learn

NOTE Confidence: 0.9379816

 $00:13:41.840 \longrightarrow 00:13:43.407$  more about the early detection

NOTE Confidence: 0.9379816

00:13:43.407 --> 00:13:45.459 of ovarian cancer with my guest,

NOTE Confidence: 0.9379816

 $00:13:45.460 \longrightarrow 00:13:46.900$  Doctor Stacy Malaker.

NOTE Confidence: 0.9379816

00:13:47.500 --> 00:13:49.545 Funding for Yale Cancer Answers

NOTE Confidence: 0.9379816

 $00{:}13{:}49.545 \dashrightarrow 00{:}13{:}51.590$  comes from Smilow Cancer Hospital,

NOTE Confidence: 0.9379816

 $00:13:51.590 \longrightarrow 00:13:53.782$  where their Oncodermatology program

NOTE Confidence: 0.9379816

 $00:13:53.782 \longrightarrow 00:13:55.426$  treats dermatologic concerns,

NOTE Confidence: 0.9379816

00:13:55.430 --> 00:13:57.435 including very dry skin, itching,

NOTE Confidence: 0.9379816

 $00{:}13{:}57.435 \dashrightarrow 00{:}13{:}59.440$  and skin changes that arise as

NOTE Confidence: 0.9379816

 $00{:}13{:}59.514 \dashrightarrow 00{:}14{:}01.590$  side effects from chemotherapy.

NOTE Confidence: 0.9379816

 $00:14:01.590 \longrightarrow 00:14:05.590$  Smilowcancerhospital.org.

NOTE Confidence: 0.9379816

00:14:05.590 --> 00:14:07.486 The American Cancer Society

 $00:14:07.486 \longrightarrow 00:14:09.590$  estimates that over 200,000 cases

NOTE Confidence: 0.9379816

 $00:14:09.590 \longrightarrow 00:14:11.290$  of Melanoma will be diagnosed

NOTE Confidence: 0.9379816

00:14:11.290 --> 00:14:13.389 in the United States this year,

NOTE Confidence: 0.9379816

 $00:14:13.390 \longrightarrow 00:14:16.659$  with over 1000 patients in Connecticut alone.

NOTE Confidence: 0.9379816

00:14:16.660 --> 00:14:18.830 While Melanoma accounts for only

NOTE Confidence: 0.9379816

 $00:14:18.830 \longrightarrow 00:14:21.308$  about 1% of skin cancer cases,

NOTE Confidence: 0.9379816

00:14:21.308 --> 00:14:24.340 it causes the most skin cancer deaths,

NOTE Confidence: 0.9379816

 $00:14:24.340 \longrightarrow 00:14:25.744$  but when detected early,

NOTE Confidence: 0.9379816

 $00:14:25.744 \longrightarrow 00:14:28.420$  it is easily treated and highly curable.

NOTE Confidence: 0.9379816

00:14:28.420 --> 00:14:30.448 Clinical trials are currently

NOTE Confidence: 0.9379816

 $00{:}14{:}30.448 \dashrightarrow 00{:}14{:}32.476$  underway at federally designated

NOTE Confidence: 0.9379816

 $00:14:32.476 \longrightarrow 00:14:34.226$  comprehensive Cancer centers such

NOTE Confidence: 0.9379816

 $00{:}14{:}34.226 \dashrightarrow 00{:}14{:}36.578$  as Yale Cancer Center and Smilow

NOTE Confidence: 0.9379816

00:14:36.578 --> 00:14:38.685 Cancer Hospital to test innovative

NOTE Confidence: 0.9379816

00:14:38.685 --> 00:14:40.393 new treatments for Melanoma.

NOTE Confidence: 0.9379816

 $00:14:40.400 \longrightarrow 00:14:42.560$  The goal of the Specialized Programs

 $00:14:42.560 \longrightarrow 00:14:44.709$  of Research Excellence in Skin Cancer

NOTE Confidence: 0.9379816

 $00{:}14{:}44.709 \dashrightarrow 00{:}14{:}46.659$  grant is to better understand the

NOTE Confidence: 0.9379816

00:14:46.659 --> 00:14:49.024 biology of skin cancer with a focus

NOTE Confidence: 0.9379816

 $00:14:49.024 \longrightarrow 00:14:51.166$  on discovering targets that will lead

NOTE Confidence: 0.9379816

 $00:14:51.166 \longrightarrow 00:14:53.596$  to improve diagnosis and treatment.

NOTE Confidence: 0.9379816

00:14:53.600 --> 00:14:56.032 More information is available

NOTE Confidence: 0.9379816

 $00:14:56.032 \longrightarrow 00:14:57.062$  at yalecancercenter.org.

NOTE Confidence: 0.9379816

 $00{:}14{:}57.062 \dashrightarrow 00{:}14{:}59.594$  You're listening to Connecticut Public Radio.

NOTE Confidence: 0.93090713

 $00:15:01.760 \longrightarrow 00:15:03.960$  Welcome back to Yale Cancer Answers.

NOTE Confidence: 0.93090713

 $00:15:03.960 \longrightarrow 00:15:05.560$  This is Dr. Anees Chagpar,

NOTE Confidence: 0.93090713

 $00:15:05.560 \longrightarrow 00:15:07.582$  and I'm joined tonight by my

NOTE Confidence: 0.93090713

 $00{:}15{:}07.582 \dashrightarrow 00{:}15{:}09.054$  guest doctor Stacy Malaker.

NOTE Confidence: 0.93090713

 $00{:}15{:}09.054 \dashrightarrow 00{:}15{:}10.944$  We're talking about the early

NOTE Confidence: 0.93090713

 $00:15:10.944 \longrightarrow 00:15:12.640$  detection of ovarian cancer.

NOTE Confidence: 0.93090713

00:15:12.640 --> 00:15:13.960 As all of you know,

 $00:15:13.960 \longrightarrow 00:15:15.910$  this has been widely talked

NOTE Confidence: 0.93090713

 $00:15:15.910 \longrightarrow 00:15:18.332$  about as the silent cancer and

NOTE Confidence: 0.93090713

 $00:15:18.332 \longrightarrow 00:15:20.317$  the cancer that whispers.

NOTE Confidence: 0.93090713

 $00:15:20.320 \longrightarrow 00:15:22.992$  And Stacy in her lab is trying to

NOTE Confidence: 0.93090713

 $00:15:22.992 \longrightarrow 00:15:25.557$  figure out whether we can actually,

NOTE Confidence: 0.93090713

00:15:25.560 --> 00:15:27.680 well, make ovarian cancer speak

NOTE Confidence: 0.93090713

 $00:15:27.680 \longrightarrow 00:15:30.295$  a little bit more loudly by

NOTE Confidence: 0.93090713

 $00:15:30.295 \longrightarrow 00:15:32.770$  looking at sugar molecules and

NOTE Confidence: 0.93090713

 $00{:}15{:}32.770 \dashrightarrow 00{:}15{:}35.520$  how they're disrupted or altered.

NOTE Confidence: 0.93090713

00:15:35.520 --> 00:15:37.116 And Stacy, right before the break,

NOTE Confidence: 0.93090713

 $00:15:37.120 \longrightarrow 00:15:39.451$  one of the things that you were

NOTE Confidence: 0.93090713

 $00:15:39.451 \longrightarrow 00:15:41.916$  talking about is that in the work

NOTE Confidence: 0.93090713

00:15:41.916 --> 00:15:44.148 up to your current project which

NOTE Confidence: 0.93090713

 $00:15:44.148 \longrightarrow 00:15:46.480$  is looking at how these alterations

NOTE Confidence: 0.93090713

 $00:15:46.480 \longrightarrow 00:15:49.176$  over time are changing and how that

NOTE Confidence: 0.93090713

 $00:15:49.176 \longrightarrow 00:15:51.352$  might affect people with a BRCA 1 or 2

00:15:51.352 --> 00:15:54.390 mutation both in the

NOTE Confidence: 0.93090713

 $00{:}15{:}54.390 \dashrightarrow 00{:}15{:}55.710$  development of ovarian cancer

NOTE Confidence: 0.93090713

 $00:15:55.710 \longrightarrow 00:15:57.280$  your primary of interest,

NOTE Confidence: 0.93090713

 $00:15:57.280 \longrightarrow 00:15:58.944$  but also other cancers.

NOTE Confidence: 0.93090713

 $00:15:58.944 \longrightarrow 00:16:01.861$  One of the things that your lab

NOTE Confidence: 0.93090713

 $00:16:01.861 \longrightarrow 00:16:04.470$  did was to really look at how

NOTE Confidence: 0.93090713

 $00:16:04.470 \longrightarrow 00:16:07.370$  you can use mass spectrometry

NOTE Confidence: 0.93090713

 $00:16:07.370 \longrightarrow 00:16:10.690$  to look at these alterations,

NOTE Confidence: 0.93090713

 $00:16:10.690 \longrightarrow 00:16:13.810$  which is something that you really

NOTE Confidence: 0.93090713

 $00{:}16{:}13.810 \dashrightarrow 00{:}16{:}17.538$  couldn't do otherwise and you couldn't

NOTE Confidence: 0.93090713

 $00:16:17.538 \longrightarrow 00:16:20.610$  do and look at at the molecular

NOTE Confidence: 0.93090713

 $00:16:20.610 \longrightarrow 00:16:22.210$  level with mass spectrometry.

NOTE Confidence: 0.93090713

 $00{:}16{:}22.210 \dashrightarrow 00{:}16{:}25.730$  So I guess the other question that I have is,

 $00:16:26.002 \longrightarrow 00:16:27.906$  can you tell us a little bit

NOTE Confidence: 0.93090713

 $00:16:27.906 \longrightarrow 00:16:29.529$  more about this technology?

NOTE Confidence: 0.93090713

00:16:29.530 --> 00:16:32.029 I mean presumably if you can now

 $00:16:32.029 \longrightarrow 00:16:34.885$  look at the sugar moieties and as

NOTE Confidence: 0.93090713

 $00{:}16{:}34.885 \dashrightarrow 00{:}16{:}37.868$  you said before the break that these

NOTE Confidence: 0.93090713

 $00{:}16{:}37.868 \operatorname{--}{>} 00{:}16{:}40.442$  alterations are seen in not just

NOTE Confidence: 0.93090713

 $00:16:40.442 \longrightarrow 00:16:43.608$  cancer but a variety of other diseases.

NOTE Confidence: 0.93090713

00:16:43.610 --> 00:16:47.486 How is this being utilized now

NOTE Confidence: 0.93090713

 $00:16:47.490 \longrightarrow 00:16:50.535$  in terms of of looking at other

NOTE Confidence: 0.93090713

 $00:16:50.535 \longrightarrow 00:16:52.730$  cancers and other diseases?

NOTE Confidence: 0.93090713

00:16:52.730 --> 00:16:54.926 I mean, how do you see this moving forward?

NOTE Confidence: 0.92526495

00:16:56.330 --> 00:16:58.346 Yeah, I mean, the world

NOTE Confidence: 0.92526495

 $00:16:58.346 \longrightarrow 00:16:59.690$  is our oyster really.

NOTE Confidence: 0.92526495

 $00:16:59.690 \longrightarrow 00:17:01.524$  We have this is 1 project of

NOTE Confidence: 0.92526495

 $00:17:01.524 \longrightarrow 00:17:03.529$  of many in my lab right now.

NOTE Confidence: 0.92526495

 $00{:}17{:}03.530 \dashrightarrow 00{:}17{:}06.754$  We're looking at cardiovascular disease.

NOTE Confidence: 0.92526495

 $00:17:06.754 \longrightarrow 00:17:08.170$  We're looking at

NOTE Confidence: 0.92526495

 $00:17:08.170 \longrightarrow 00:17:09.490$  breast cancer,

 $00:17:09.490 \longrightarrow 00:17:12.400$  but in a different fashion.

NOTE Confidence: 0.92526495

 $00:17:12.400 \longrightarrow 00:17:14.885$  And we also look at changes in

NOTE Confidence: 0.92526495

 $00{:}17{:}14.885 \dashrightarrow 00{:}17{:}16.692$  intestinal linings and stress and

NOTE Confidence: 0.92526495

00:17:16.692 --> 00:17:19.037 depression and so on and so forth.

NOTE Confidence: 0.92526495

 $00:17:19.040 \longrightarrow 00:17:21.158$  And so we're really trying to

NOTE Confidence: 0.92526495

 $00{:}17{:}21.158 \dashrightarrow 00{:}17{:}22.570$  monopolize on these developments

NOTE Confidence: 0.92526495

00:17:22.631 --> 00:17:24.941 that we've made in order to study

NOTE Confidence: 0.92526495

 $00:17:24.941 \longrightarrow 00:17:26.612$  altered sugar structures in a

NOTE Confidence: 0.92526495

00:17:26.612 --> 00:17:28.157 whole host of different diseases.

NOTE Confidence: 0.93757594

00:17:29.520 --> 00:17:32.184 And so tell us a little bit more about,

NOTE Confidence: 0.93757594

 $00{:}17{:}32.190 \dashrightarrow 00{:}17{:}34.190$  you know, these sugar moieties.

NOTE Confidence: 0.93757594

 $00{:}17{:}34.190 \to 00{:}17{:}36.734$  I mean, I know that you became very

NOTE Confidence: 0.93757594

 $00:17:36.734 \longrightarrow 00:17:38.963$  interested in these during your postdoc

NOTE Confidence: 0.93757594

 $00:17:38.963 \longrightarrow 00:17:41.255$  working with a Nobel Prize winner

NOTE Confidence: 0.93757594

 $00:17:41.328 \longrightarrow 00:17:43.470$  whose lab really looked at these,

NOTE Confidence: 0.93757594

 $00:17:43.470 \longrightarrow 00:17:46.070$  these molecules. But you know,

 $00:17:46.070 \longrightarrow 00:17:49.490$  these days I think a lot of people think

NOTE Confidence: 0.93757594

 $00{:}17{:}49.490 {\:{\circ}{\circ}{\circ}}>00{:}17{:}52.436$  about cancer from the perspective of

NOTE Confidence: 0.93757594

 $00:17:52.436 \longrightarrow 00:17:56.733$  genetics and they think about it from the

NOTE Confidence: 0.93757594

 $00:17:56.733 \longrightarrow 00:17:59.045$  perspective of environmental factors.

NOTE Confidence: 0.93757594

 $00:17:59.050 \longrightarrow 00:18:01.703$  But we really don't think about how

NOTE Confidence: 0.93757594

 $00:18:01.703 \longrightarrow 00:18:04.528$  these two things affect sugars.

NOTE Confidence: 0.93757594

 $00:18:04.530 \longrightarrow 00:18:06.778$  So can you tell us a little bit

NOTE Confidence: 0.93757594

 $00{:}18{:}06.778 \dashrightarrow 00{:}18{:}08.253$  more about those interactions

NOTE Confidence: 0.93757594

 $00:18:08.253 \longrightarrow 00:18:10.767$  and how prevalent they are?

NOTE Confidence: 0.93757594

 $00{:}18{:}10.770 \dashrightarrow 00{:}18{:}13.087$  I mean, do you really think that

NOTE Confidence: 0.93757594

 $00:18:13.087 \longrightarrow 00:18:15.517$  by looking at these sugar muleides

NOTE Confidence: 0.93757594

00:18:15.517 --> 00:18:18.186 that we might actually, you know,

NOTE Confidence: 0.93757594

 $00{:}18{:}18.186 \dashrightarrow 00{:}18{:}21.182$  kind of unlock a portion of cancer

NOTE Confidence: 0.93757594

 $00:18:21.182 \longrightarrow 00:18:23.805$  biology that had heretofore been

NOTE Confidence: 0.93757594

00:18:23.805 --> 00:18:26.967 largely well overlooked to some degree?

 $00:18:28.090 \longrightarrow 00:18:28.890$  Yeah, absolutely.

NOTE Confidence: 0.9343778

 $00{:}18{:}28.890 \dashrightarrow 00{:}18{:}31.874$  I think that sugar structures,

NOTE Confidence: 0.9343778

00:18:31.874 --> 00:18:33.626 sugar structures, excuse me,

NOTE Confidence: 0.9343778

00:18:33.626 --> 00:18:35.766 are extremely difficult to study.

NOTE Confidence: 0.9343778

 $00:18:35.770 \longrightarrow 00:18:37.978$  One of the issues is that

NOTE Confidence: 0.9343778

00:18:37.978 --> 00:18:39.450 you just mentioned genetics,

NOTE Confidence: 0.9343778

 $00:18:39.450 \longrightarrow 00:18:41.090$  glycobiology or the sugar

NOTE Confidence: 0.9343778

00:18:41.090 --> 00:18:42.730 structures are not templated,

NOTE Confidence: 0.9343778

 $00:18:42.730 \longrightarrow 00:18:45.298$  meaning that there are 200 different

NOTE Confidence: 0.9343778

 $00:18:45.298 \longrightarrow 00:18:47.648$  enzymes that build these sugar

NOTE Confidence: 0.9343778

 $00:18:47.648 \longrightarrow 00:18:49.848$  structures on the surface of our cells.

NOTE Confidence: 0.9343778

 $00:18:49.850 \longrightarrow 00:18:51.600$  And so you can't necessarily

NOTE Confidence: 0.9343778

00:18:51.600 --> 00:18:53.760 look at changes in those enzyme

NOTE Confidence: 0.9343778

 $00:18:53.760 \longrightarrow 00:18:55.818$  levels via genetics in order to

NOTE Confidence: 0.9343778

00:18:55.818 --> 00:18:57.623 build back up what's possibly

NOTE Confidence: 0.9343778

00:18:57.623 --> 00:19:00.087 going to be on the cell surface.

 $00:19:00.090 \longrightarrow 00:19:01.608$  And so because of that it's

00:19:02.404 --> 00:19:04.786 much more difficult to study and

NOTE Confidence: 0.9343778

00:19:04.786 --> 00:19:07.688 so it's lagged behind in you know,

NOTE Confidence: 0.9343778

 $00:19:07.690 \longrightarrow 00:19:11.365$  in comparison to more general fields like

NOTE Confidence: 0.9343778

 $00:19:11.365 \longrightarrow 00:19:14.089$  genomics or transcriptomics or proteomics.

NOTE Confidence: 0.9343778

 $00:19:14.090 \longrightarrow 00:19:15.286$  And so

NOTE Confidence: 0.9343778

 $00:19:15.286 \longrightarrow 00:19:17.424$  we really want to monopolize on these

NOTE Confidence: 0.9343778

 $00:19:17.424 \longrightarrow 00:19:19.391$  changes in order to break open a

NOTE Confidence: 0.9343778

 $00:19:19.391 \longrightarrow 00:19:21.368$  whole new area of cancer biology.

NOTE Confidence: 0.9391263

 $00:19:22.130 \longrightarrow 00:19:25.522$  I mean, do you think that there's an

NOTE Confidence: 0.9391263

 $00{:}19{:}25.522 \dashrightarrow 00{:}19{:}28.509$  interplay between genomics and

NOTE Confidence: 0.9391263

 $00:19:28.509 \longrightarrow 00:19:30.648$  these sugar structures?

NOTE Confidence: 0.9391263

00:19:30.650 --> 00:19:33.426 Or do you think that these are two

NOTE Confidence: 0.9391263

 $00:19:33.426 \longrightarrow 00:19:35.924$  separate issues that they cause or

NOTE Confidence: 0.9391263

 $00:19:35.924 \longrightarrow 00:19:38.084$  are affected by cancer independently?

NOTE Confidence: 0.9391263

 $00:19:38.090 \longrightarrow 00:19:38.970$  In other words, I mean,

 $00:19:38.970 \longrightarrow 00:19:40.512$  do you think that these two

NOTE Confidence: 0.9391263

00:19:40.512 --> 00:19:41.970 play together or not really?

NOTE Confidence: 0.933372370000001

 $00:19:42.450 \longrightarrow 00:19:43.570$  Oh, they definitely do.

NOTE Confidence: 0.933372370000001

 $00:19:43.570 \longrightarrow 00:19:45.942$  It's just that you can't look at enzyme

NOTE Confidence: 0.933372370000001

 $00{:}19{:}45.942 \dashrightarrow 00{:}19{:}47.844$  changes and then immediately know how

NOTE Confidence: 0.933372370000001

 $00:19:47.844 \longrightarrow 00:19:49.912$  that's going to change the sugar

NOTE Confidence: 0.933372370000001

 $00:19:49.912 \longrightarrow 00:19:52.066$  structures on the outside of the cell.

NOTE Confidence: 0.933372370000001

 $00:19:52.066 \dashrightarrow 00:19:55.042$  But you can kind of gain hypothesis by

NOTE Confidence: 0.933372370000001

 $00{:}19{:}55.042 \dashrightarrow 00{:}19{:}57.450$  looking at changes in the enzyme levels.

NOTE Confidence: 0.933372370000001

 $00:19:57.450 \longrightarrow 00:20:00.450$  So if for instance,

NOTE Confidence: 0.933372370000001

 $00{:}20{:}00.450 \dashrightarrow 00{:}20{:}02.005$  there's a capping structure called

NOTE Confidence: 0.933372370000001

 $00{:}20{:}02.005 \dashrightarrow 00{:}20{:}04.674$  sialic acid and you can look at the sial

NOTE Confidence: 0.933372370000001

 $00{:}20{:}04.674 \dashrightarrow 00{:}20{:}06.740$  transfer ases and if those are up or down

NOTE Confidence: 0.933372370000001

 $00{:}20{:}06.740 \dashrightarrow 00{:}20{:}08.498$  you could then gather that your

NOTE Confidence: 0.933372370000001

 $00:20:08.498 \longrightarrow 00:20:10.593$  structures will have more or less of a

00:20:10.593 --> 00:20:12.460 certain type of that sugar structure,

NOTE Confidence: 0.933372370000001

00:20:12.460 --> 00:20:14.272 but it won't tell you exactly

NOTE Confidence: 0.933372370000001

 $00:20:14.272 \longrightarrow 00:20:15.178$  what it's modifying.

NOTE Confidence: 0.933372370000001

 $00:20:15.180 \longrightarrow 00:20:16.979$  So what protein it's on or

NOTE Confidence: 0.933372370000001

 $00:20:16.979 \longrightarrow 00:20:18.684$  it won't tell you exactly what

NOTE Confidence: 0.933372370000001

 $00:20:18.684 \longrightarrow 00:20:20.430$  type of sugar structure it's

NOTE Confidence: 0.933372370000001

 $00:20:20.430 \longrightarrow 00:20:22.099$  on and so on and so forth.

NOTE Confidence: 0.92736816

 $00:20:22.660 \longrightarrow 00:20:25.188$  And so going back to

NOTE Confidence: 0.92736816

 $00{:}20{:}25.188 \dashrightarrow 00{:}20{:}27.570$  the project for which you were just

NOTE Confidence: 0.92736816

00:20:27.570 --> 00:20:29.708 awarded a grant where you're looking

NOTE Confidence: 0.92736816

00:20:29.708 --> 00:20:32.392 at these BRCA mutation carriers,

NOTE Confidence: 0.92736816

00:20:32.392 --> 00:20:36.090 is it possible that BRCA in and of itself,

NOTE Confidence: 0.92736816

00:20:36.090 --> 00:20:41.090 I mean we know BRCA as being a gene which

NOTE Confidence: 0.92736816

 $00:20:41.090 \longrightarrow 00:20:44.210$  is largely responsible for DNA repair.

NOTE Confidence: 0.92736816

00:20:44.210 --> 00:20:47.207 And so when you get a mutation in that,

NOTE Confidence: 0.92736816

 $00:20:47.210 \longrightarrow 00:20:50.275$  it's difficult to correct those

00:20:50.275 --> 00:20:52.702 mistakes that your DNA may have and

NOTE Confidence: 0.92736816

 $00:20:52.702 \longrightarrow 00:20:54.940$  the thinking is that

NOTE Confidence: 0.92736816

 $00:20:54.940 \longrightarrow 00:20:57.187$  really leads to the higher risk of

NOTE Confidence: 0.92736816

 $00:20:57.260 \longrightarrow 00:20:59.690$  developing a variety of malignancies.

NOTE Confidence: 0.92736816

 $00:20:59.690 \longrightarrow 00:21:03.452$  So if genetics and these altered

NOTE Confidence: 0.92736816

 $00:21:03.452 \longrightarrow 00:21:05.960$  sugar structures are related,

NOTE Confidence: 0.92736816

 $00:21:05.960 \longrightarrow 00:21:08.046$  do you think that

NOTE Confidence: 0.92736816

 $00{:}21{:}08.046 \dashrightarrow 00{:}21{:}10.790$  BRCA might be doing something to the sugar

NOTE Confidence: 0.92736816

00:21:10.863 --> 00:21:13.439 structures and are you looking at that?

NOTE Confidence: 0.92736816

00:21:13.440 --> 00:21:14.416 For example,

NOTE Confidence: 0.92736816

00:21:14.416 --> 00:21:17.506 are you comparing BRCA carriers to

NOTE Confidence: 0.92736816

 $00:21:17.506 \longrightarrow 00:21:20.396$  people who are not BRCA carriers

NOTE Confidence: 0.92736816

 $00{:}21{:}20.396 \dashrightarrow 00{:}21{:}22.186$  and seeing whether there's a

NOTE Confidence: 0.92736816

 $00:21:22.186 \longrightarrow 00:21:23.870$  difference in terms of

NOTE Confidence: 0.92736816

00:21:23.870 --> 00:21:25.590 these sugar structures between

 $00:21:25.590 \longrightarrow 00:21:26.880$  these two populations?

NOTE Confidence: 0.92099065

 $00:21:27.760 \longrightarrow 00:21:29.080$  That's not something that we're currently

NOTE Confidence: 0.92099065

 $00:21:29.080 \longrightarrow 00:21:31.880$  looking at simply because we

NOTE Confidence: 0.92099065

 $00:21:31.880 \longrightarrow 00:21:34.896$  only have access to these BRCA1 and 2 samples.

00:21:37.372 --> 00:21:40.276 But we could ostensibly look at healthy,

NOTE Confidence: 0.92099065

 $00:21:40.280 \longrightarrow 00:21:42.080$  you know, healthy samples or healthy

NOTE Confidence: 0.92099065

00:21:42.080 --> 00:21:44.160 patient serum in order to compare them.

NOTE Confidence: 0.92099065

00:21:44.160 --> 00:21:45.678 So definitely something we could do,

NOTE Confidence: 0.92099065

 $00:21:45.680 \longrightarrow 00:21:46.600$  but not something that's

NOTE Confidence: 0.92099065

 $00:21:46.600 \longrightarrow 00:21:49.480$  currently on our docket.

NOTE Confidence: 0.92099065

 $00{:}21{:}49.480 \dashrightarrow 00{:}21{:}51.209$  And then the other thing that I

NOTE Confidence: 0.92099065

 $00:21:51.209 \longrightarrow 00:21:53.291$  kind of wonder about is one of the

NOTE Confidence: 0.92099065

00:21:53.291 --> 00:21:54.800 questions I always get asked is,

NOTE Confidence: 0.92099065

 $00:21:54.800 \longrightarrow 00:21:57.596$  well, why did I get cancer?

NOTE Confidence: 0.92099065

00:21:57.600 --> 00:22:00.016 Can you tell us a little bit more

NOTE Confidence: 0.92099065

 $00:22:00.016 \longrightarrow 00:22:02.112$  about whether you think that

 $00:22:02.112 \longrightarrow 00:22:04.352$  having these altered sugar moides

NOTE Confidence: 0.92099065

 $00{:}22{:}04.352 \dashrightarrow 00{:}22{:}06.678$  might have something to do with

NOTE Confidence: 0.92099065

00:22:06.680 --> 00:22:09.000 people's risk of developing cancer?

NOTE Confidence: 0.92099065

 $00:22:09.000 \longrightarrow 00:22:10.720$  And secondary to that,

NOTE Confidence: 0.92099065

 $00:22:10.720 \longrightarrow 00:22:13.300$  why do people have these alterations

NOTE Confidence: 0.92099065

 $00:22:13.373 \longrightarrow 00:22:15.558$  in these sugar moides anyways?

NOTE Confidence: 0.92099065

 $00:22:15.560 \longrightarrow 00:22:17.440$  I mean what causes that?

NOTE Confidence: 0.8961784

 $00:22:17.680 \longrightarrow 00:22:19.399$  Again, that's a very,

NOTE Confidence: 0.8961784

00:22:19.399 --> 00:22:21.118 very loaded question.

NOTE Confidence: 0.8961784

 $00:22:21.120 \longrightarrow 00:22:22.560$  So what was the first part

NOTE Confidence: 0.8961784

 $00:22:22.560 \longrightarrow 00:22:23.600$  of the question?

NOTE Confidence: 0.8961784

 $00{:}22{:}23.640 \dashrightarrow 00{:}22{:}26.270$  Could these altered sugar Moides

NOTE Confidence: 0.8961784

 $00:22:26.270 \longrightarrow 00:22:28.692$  be part of the explanation of why

NOTE Confidence: 0.8961784

 $00{:}22{:}28.692 \dashrightarrow 00{:}22{:}30.645$ some people develop cancer even

NOTE Confidence: 0.8961784

00:22:30.645 --> 00:22:32.745 though they did everything right?

NOTE Confidence: 0.934395

 $00:22:33.030 \longrightarrow 00:22:36.960$  Sure. So I mean there are many,

00:22:36.960 --> 00:22:38.670 many possible answers to that question,

NOTE Confidence: 0.934395

 $00:22:38.670 \longrightarrow 00:22:40.080$  but I'll probably lean into the

NOTE Confidence: 0.934395

 $00:22:40.080 \longrightarrow 00:22:41.750$  one that I'm most familiar with.

NOTE Confidence: 0.934395

 $00:22:41.750 \longrightarrow 00:22:43.755$  So you know, cancer immunotherapies

NOTE Confidence: 0.934395

 $00:22:43.755 \longrightarrow 00:22:46.541$  are the the new pillar of

NOTE Confidence: 0.934395

00:22:46.541 --> 00:22:48.665 treatment as I'm sure you're aware.

NOTE Confidence: 0.934395

 $00:22:48.670 \longrightarrow 00:22:50.325$  And so altered sugar structures

NOTE Confidence: 0.934395

 $00{:}22{:}50.325 \dashrightarrow 00{:}22{:}52.763$  are a way that cancer cells can

NOTE Confidence: 0.934395

 $00:22:52.763 \longrightarrow 00:22:54.899$  actually avoid the immune system and

NOTE Confidence: 0.934395

 $00:22:54.899 \longrightarrow 00:22:57.212$  the immune system is really key in

NOTE Confidence: 0.934395

00:22:57.212 --> 00:22:59.640 getting rid of cells that have become

NOTE Confidence: 0.934395

 $00{:}22{:}59.640 \dashrightarrow 00{:}23{:}02.310$  transformed or cancerous.

NOTE Confidence: 0.934395

 $00{:}23{:}02.310 \dashrightarrow 00{:}23{:}04.326$  And so there's this really fine-tuned

NOTE Confidence: 0.934395

 $00:23:04.326 \longrightarrow 00:23:06.308$  balance there where you want your

NOTE Confidence: 0.934395

 $00:23:06.308 \longrightarrow 00:23:08.030$  immune system to be active and

 $00:23:08.030 \longrightarrow 00:23:09.708$  killing off these cancer cells.

NOTE Confidence: 0.934395

 $00{:}23{:}09.710 \dashrightarrow 00{:}23{:}11.822$  Now the sugar moieties can actually

NOTE Confidence: 0.934395

 $00:23:11.822 \longrightarrow 00:23:14.356$  act as a mechanism to shield the

NOTE Confidence: 0.934395

 $00:23:14.356 \longrightarrow 00:23:16.116$  cancer cell from immune cells

NOTE Confidence: 0.934395

 $00:23:16.116 \longrightarrow 00:23:18.257$  that would normally kill it off.

NOTE Confidence: 0.934395

 $00:23:18.260 \longrightarrow 00:23:19.252$  For instance,

NOTE Confidence: 0.934395

 $00:23:19.252 \longrightarrow 00:23:22.948$  my lab studies what's called a checkpoint

NOTE Confidence: 0.934395

 $00:23:22.948 \longrightarrow 00:23:24.932$  inhibitor where when that

NOTE Confidence: 0.934395

 $00{:}23{:}24.932 \dashrightarrow 00{:}23{:}27.080$  checkpoint inhibitor is bound to one

NOTE Confidence: 0.934395

00:23:27.148 --> 00:23:29.578 of its ligands through sugar structures,

NOTE Confidence: 0.934395

 $00:23:29.580 \longrightarrow 00:23:31.980$  it shuts down T cell function.

NOTE Confidence: 0.934395

 $00:23:31.980 \longrightarrow 00:23:35.244$  And it's so important that antibodies

NOTE Confidence: 0.934395

 $00:23:35.244 \longrightarrow 00:23:37.788$  that block that interaction are currently

NOTE Confidence: 0.934395

 $00:23:37.788 \longrightarrow 00:23:39.749$  being investigated in the clinic.

NOTE Confidence: 0.934395

00:23:39.750 --> 00:23:41.822 And so we're trying to again monopolize

NOTE Confidence: 0.934395

 $00{:}23{:}41.822 \dashrightarrow 00{:}23{:}43.473$  on the altered sugar structures

 $00:23:43.473 \longrightarrow 00:23:45.228$  in order to potentially develop

NOTE Confidence: 0.934395

 $00{:}23{:}45.228 \dashrightarrow 00{:}23{:}47.110$  a better cancer immunotherapy.

NOTE Confidence: 0.934395

00:23:47.110 --> 00:23:49.162 But basically kind of summarizing that

NOTE Confidence: 0.934395

 $00:23:49.162 \longrightarrow 00:23:51.567$  is that these sugar moieties can serve

NOTE Confidence: 0.934395

 $00:23:51.567 \longrightarrow 00:23:53.835$  to shut down various types of immune

NOTE Confidence: 0.934395

 $00:23:53.903 \longrightarrow 00:23:56.255$  cells which then allow the tumor cells

NOTE Confidence: 0.934395

 $00:23:56.255 \longrightarrow 00:23:58.503$  to proliferate and become

NOTE Confidence: 0.934395

00:23:58.503 --> 00:24:01.389 a solid tumor or various cancers.

NOTE Confidence: 0.9340521

 $00:24:02.430 \longrightarrow 00:24:06.502$  So why do some people get these

NOTE Confidence: 0.9340521

 $00{:}24{:}06.502 \dashrightarrow 00{:}24{:}08.757$  altered sugar moieties that can

NOTE Confidence: 0.9340521

00:24:08.757 --> 00:24:11.049 essentially shut down your immune

NOTE Confidence: 0.9340521

00:24:11.049 --> 00:24:13.748 system or at least its ability to

NOTE Confidence: 0.9340521

00:24:13.748 --> 00:24:16.319 detect cancer and other people don't?

NOTE Confidence: 0.9340521

 $00:24:16.320 \longrightarrow 00:24:19.600$  I mean, are there factors that drive that?

NOTE Confidence: 0.9340521

00:24:19.600 --> 00:24:21.196 You know, some people might be wondering,

 $00:24:21.200 \longrightarrow 00:24:22.943$  is it the sugar that I'm eating

NOTE Confidence: 0.9340521

00:24:22.943 --> 00:24:24.998 or is it how I metabolize it?

NOTE Confidence: 0.9340521

 $00:24:25.000 \longrightarrow 00:24:26.780$  Or is it, you know, diabetes?

NOTE Confidence: 0.9340521

 $00:24:26.780 \longrightarrow 00:24:30.200$  Or is it something to do with my genetics?

NOTE Confidence: 0.9360402

 $00:24:31.210 \longrightarrow 00:24:32.554$  Yeah. So I mean that's a great question

NOTE Confidence: 0.9360402

 $00:24:32.554 \longrightarrow 00:24:33.890$  that I don't have the answer for.

NOTE Confidence: 0.9360402

 $00:24:33.890 \longrightarrow 00:24:35.530$  I will specify that the

NOTE Confidence: 0.9360402

 $00:24:35.530 \longrightarrow 00:24:37.364$  sugars that you're eating are very,

NOTE Confidence: 0.9360402

00:24:37.370 --> 00:24:38.970 very, very different than the

NOTE Confidence: 0.9360402

 $00:24:38.970 \longrightarrow 00:24:40.250$  sugars I'm talking about.

NOTE Confidence: 0.9360402

 $00{:}24{:}40.250 \dashrightarrow 00{:}24{:}42.140$  I mean, essentially they can get

NOTE Confidence: 0.9360402

 $00:24:42.140 \longrightarrow 00:24:44.043$  metabolized and turned into the sugar

NOTE Confidence: 0.9360402

 $00{:}24{:}44.043 \dashrightarrow 00{:}24{:}46.087$  structures that are on the cell surface.

NOTE Confidence: 0.9360402

 $00:24:46.090 \longrightarrow 00:24:47.776$  But I'm not looking at glucose

NOTE Confidence: 0.9360402

 $00:24:47.776 \longrightarrow 00:24:49.489$  or sucrose or anything like that.

NOTE Confidence: 0.9360402

 $00{:}24{:}49.490 \dashrightarrow 00{:}24{:}51.330$  These are very different structures.

 $00:24:52.530 \longrightarrow 00:24:53.698$  And so, you know,

NOTE Confidence: 0.9360402

00:24:53.698 --> 00:24:56.529 I think a lot of people may be asking,

NOTE Confidence: 0.9360402

 $00:24:56.530 \longrightarrow 00:25:00.606$  especially now that the WHO is

NOTE Confidence: 0.9360402

 $00:25:00.606 \longrightarrow 00:25:02.971$  coming out with their statement

NOTE Confidence: 0.9360402

 $00{:}25{:}02.971 \dashrightarrow 00{:}25{:}05.495$  against some artificial sweeteners of

NOTE Confidence: 0.9360402

 $00:25:05.495 \longrightarrow 00:25:08.573$  thinking that they may be carcinogenic.

NOTE Confidence: 0.9360402

 $00:25:08.580 \longrightarrow 00:25:11.928$  Do do those have anything to do with the

NOTE Confidence: 0.9360402

00:25:11.928 --> 00:25:15.216 sugar moieties that you're talking about?

NOTE Confidence: 0.9360402

00:25:15.220 --> 00:25:18.739 I don't know, my understanding

NOTE Confidence: 0.9337291

 $00:25:18.740 \longrightarrow 00:25:20.284$  for those those altered

NOTE Confidence: 0.9337291

 $00:25:20.284 \longrightarrow 00:25:22.214$  sugar moieties that are in,

NOTE Confidence: 0.9337291

 $00{:}25{:}22.220 \dashrightarrow 00{:}25{:}23.920$  you know artificial sweeteners and

NOTE Confidence: 0.9337291

 $00{:}25{:}23.920 \dashrightarrow 00{:}25{:}25.938$  so on is that they can't be broken

NOTE Confidence: 0.9337291

00:25:25.938 --> 00:25:27.329 down or metabolized in the same

NOTE Confidence: 0.9337291

 $00:25:27.329 \longrightarrow 00:25:28.673$  way that normal sugars would be.

 $00:25:28.680 \longrightarrow 00:25:31.277$  But that is just what I understand.

NOTE Confidence: 0.9337291

 $00{:}25{:}31.280 \dashrightarrow 00{:}25{:}34.080$  I have not studied up on that too much.

NOTE Confidence: 0.92916805

 $00:25:35.120 \longrightarrow 00:25:39.440$  So for the alterations

NOTE Confidence: 0.92916805

 $00:25:39.440 \longrightarrow 00:25:42.518$  of sugar moties, I mean the

NOTE Confidence: 0.92916805

 $00:25:42.518 \longrightarrow 00:25:44.354$  the truth of the matter is,

NOTE Confidence: 0.92916805

 $00:25:44.360 \longrightarrow 00:25:46.166$  that at least the research

NOTE Confidence: 0.92916805

 $00:25:46.166 \longrightarrow 00:25:47.680$  that you've done so far,

NOTE Confidence: 0.92916805

 $00:25:47.680 \longrightarrow 00:25:52.108$  your hypothesis is that these alterations

NOTE Confidence: 0.92916805

 $00:25:52.110 \longrightarrow 00:25:54.504$  have a role to play in cancer,

NOTE Confidence: 0.92916805

00:25:54.510 --> 00:25:57.385 whether it's the immune system

NOTE Confidence: 0.92916805

00:25:57.385 --> 00:26:00.264 evading cancers or you know,

NOTE Confidence: 0.92916805

 $00{:}26{:}00.264 \dashrightarrow 00{:}26{:}03.149$  increasing risk or whatever.

NOTE Confidence: 0.92916805

 $00:26:03.150 \longrightarrow 00:26:05.257$  Do we know of any risk factors

NOTE Confidence: 0.92916805

 $00{:}26{:}05.257 \dashrightarrow 00{:}26{:}07.140$  that make people more susceptible

NOTE Confidence: 0.92916805

00:26:07.140 --> 00:26:09.425 to having altered sugar moieties,

NOTE Confidence: 0.92916805

 $00:26:09.430 \longrightarrow 00:26:10.750$  the ones that you're studying?

 $00:26:12.310 \longrightarrow 00:26:15.390$  I mean not that I'm aware of.

NOTE Confidence: 0.9343979

 $00{:}26{:}15.390 --> 00{:}26{:}17.118$  I think that if you did

NOTE Confidence: 0.9343979

00:26:17.118 --> 00:26:17.982 genetic studies again,

NOTE Confidence: 0.9343979

00:26:17.990 --> 00:26:20.395 you could probably create hypothesis

NOTE Confidence: 0.9343979

 $00{:}26{:}20.395 \dashrightarrow 00{:}26{:}22.319$  and individuals regarding different

NOTE Confidence: 0.9343979

 $00:26:22.319 \longrightarrow 00:26:24.665$  enzymes that are up or down regulated.

NOTE Confidence: 0.9343979

 $00:26:24.670 \longrightarrow 00:26:26.506$  But as far as I'm aware,

NOTE Confidence: 0.9343979

00:26:26.510 --> 00:26:29.102 there's not anything like a

NOTE Confidence: 0.9343979

00:26:29.102 --> 00:26:30.582 BRCA1 that would definitely

NOTE Confidence: 0.9343979

 $00:26:30.582 \longrightarrow 00:26:32.548$  indicate that you're going to have

NOTE Confidence: 0.9343979

 $00:26:32.548 \longrightarrow 00:26:33.908$  these altered sugar structures.

NOTE Confidence: 0.93575597

00:26:34.990 --> 00:26:37.926 And my perception is from your

NOTE Confidence: 0.93575597

 $00{:}26{:}37.926 {\:{\circ}{\circ}{\circ}}>00{:}26{:}40.029$  description of your earlier study,

NOTE Confidence: 0.93575597

 $00:26:40.030 \longrightarrow 00:26:42.214$  is that it's not like you're born

NOTE Confidence: 0.93575597

 $00:26:42.214 \longrightarrow 00:26:44.110$  with these altered sugar moieties,

 $00:26:44.110 \longrightarrow 00:26:46.306$  it's that they develop over time.

NOTE Confidence: 0.93575597

 $00:26:46.310 \longrightarrow 00:26:47.818$  Is that right?

NOTE Confidence: 0.9361285

00:26:47.820 --> 00:26:49.737 I mean it would be kind of similar to,

NOTE Confidence: 0.9361285

 $00:26:49.740 \longrightarrow 00:26:52.035$  you know, genetic mutations that

NOTE Confidence: 0.9361285

 $00:26:52.035 \longrightarrow 00:26:55.420$  accumulate over time in cancer cells.

NOTE Confidence: 0.9361285

 $00:26:55.420 \longrightarrow 00:26:57.742$  And again,

NOTE Confidence: 0.9361285

 $00{:}26{:}57.742 \longrightarrow 00{:}26{:}59.308$  you were asking if genetics and

NOTE Confidence: 0.9361285

 $00:26:59.308 \longrightarrow 00:27:00.980$  altered sugar structures are related.

NOTE Confidence: 0.9361285

00:27:00.980 --> 00:27:02.780 If you acquire many,

NOTE Confidence: 0.9361285

00:27:02.780 --> 00:27:05.300 many genetic mutations over time,

NOTE Confidence: 0.9361285

 $00:27:05.300 \longrightarrow 00:27:07.340$  you tend to develop cancer.

NOTE Confidence: 0.9361285

00:27:07.340 --> 00:27:09.734 Similarly, you would also

NOTE Confidence: 0.9361285

 $00:27:09.734 \longrightarrow 00:27:11.734$  mutate these various glycan structures

NOTE Confidence: 0.9361285

 $00:27:11.734 \longrightarrow 00:27:14.100$  on the surface of cells.

NOTE Confidence: 0.9316189

 $00{:}27{:}15.270 \dashrightarrow 00{:}27{:}18.086$  And so it sounds like there's a lot

NOTE Confidence: 0.9316189

 $00:27:18.086 \longrightarrow 00:27:20.757$  going on in your laboratory both

 $00:27:20.757 \longrightarrow 00:27:23.886$  on the kind of developing the

NOTE Confidence: 0.9316189

 $00{:}27{:}23.886 \dashrightarrow 00{:}27{:}26.402$  methodologies as well as in terms of

NOTE Confidence: 0.9316189

 $00:27:26.402 \longrightarrow 00:27:28.346$  looking at the actual clinical impact

NOTE Confidence: 0.9316189

 $00:27:28.346 \longrightarrow 00:27:30.547$  of these altered sugar moieties.

NOTE Confidence: 0.9316189

 $00:27:30.550 \longrightarrow 00:27:33.022$  Looking forward, what projects are you

NOTE Confidence: 0.9316189

 $00:27:33.022 \longrightarrow 00:27:35.658$  most excited about and what do you

NOTE Confidence: 0.9316189

00:27:35.658 --> 00:27:38.115 think we can expect to hear about in

NOTE Confidence: 0.9316189

 $00:27:38.115 \longrightarrow 00:27:40.428$  the next year or two or five or 10?

NOTE Confidence: 0.93945575

00:27:42.030 --> 00:27:45.140 Oh gosh, my students listen to this and

NOTE Confidence: 0.93945575

 $00{:}27{:}45.140 \dashrightarrow 00{:}27{:}46.989$  I won't say their individual projects.

NOTE Confidence: 0.93945575

00:27:46.990 --> 00:27:48.826 I don't want to pick favorites.

NOTE Confidence: 0.93945575

 $00{:}27{:}48.830 \dashrightarrow 00{:}27{:}50.710$  Obviously I'm very excited about

NOTE Confidence: 0.93945575

00:27:50.710 --> 00:27:52.590 this ovarian cancer project simply

NOTE Confidence: 0.93945575

00:27:52.650 --> 00:27:54.460 because I think that, you know,

NOTE Confidence: 0.93945575

 $00:27:54.460 \longrightarrow 00:27:56.980$  CA-125 is really a black box of information

 $00:27:57.044 \longrightarrow 00:27:59.276$  that I think we can monopolize on to

NOTE Confidence: 0.93945575

 $00{:}27{:}59.276 \longrightarrow 00{:}28{:}01.390$  develop an improved diagnostic tool.

NOTE Confidence: 0.93945575

 $00:28:01.390 \longrightarrow 00:28:03.855$  And it's a somewhat selfish project

NOTE Confidence: 0.93945575

 $00:28:03.855 \longrightarrow 00:28:07.349$  because I am a BRCA 2 carrier.

NOTE Confidence: 0.93945575

 $00{:}28{:}07.350 \dashrightarrow 00{:}28{:}09.996$  So I would like to identify ovarian

NOTE Confidence: 0.93945575

 $00:28:09.996 \longrightarrow 00:28:12.364$  cancer earlier for my own self

NOTE Confidence: 0.93945575

 $00:28:12.364 \longrightarrow 00:28:14.885$  and family in in addition to all

NOTE Confidence: 0.93945575

 $00:28:14.885 \longrightarrow 00:28:17.630$  of the women that are at risk.

NOTE Confidence: 0.93945575

00:28:17.630 --> 00:28:19.150 But I also, you know,

NOTE Confidence: 0.93945575

00:28:19.150 --> 00:28:21.790 I love all of my projects equally in my lab,

NOTE Confidence: 0.93945575

 $00:28:21.790 \longrightarrow 00:28:23.794$  and I'm really excited about the

NOTE Confidence: 0.93945575

 $00:28:23.794 \longrightarrow 00:28:25.430$  instrumentation developments that we have,

NOTE Confidence: 0.93945575

00:28:25.430 --> 00:28:27.908 as well as really cracking open

NOTE Confidence: 0.93945575

 $00:28:27.910 \longrightarrow 00:28:30.270$  all of the biological underlying

NOTE Confidence: 0.93945575

 $00:28:30.270 \longrightarrow 00:28:31.749$  of altered glycosylation.

NOTE Confidence: 0.9326976

 $00{:}28{:}32.590 \dashrightarrow 00{:}28{:}34.882$  Doctor Stacy Malaker is an assistant

 $00:28:34.882 \longrightarrow 00:28:36.845$  professor in the Department of

NOTE Confidence: 0.9326976

 $00:28:36.845 \longrightarrow 00:28:38.349$  Chemistry at Yale University.

NOTE Confidence: 0.9326976

 $00:28:38.350 \longrightarrow 00:28:40.528$  If you have questions, the address

NOTE Confidence: 0.9326976

 $00{:}28{:}40.528 \dashrightarrow 00{:}28{:}42.909$  is Cancer Answers at Yale dot Edu.

NOTE Confidence: 0.9326976

 $00{:}28{:}42.910 \dashrightarrow 00{:}28{:}45.334$  And past editions of the program

NOTE Confidence: 0.9326976

 $00{:}28{:}45.334 \dashrightarrow 00{:}28{:}47.730$  are available in audio and written

NOTE Confidence: 0.9326976

 $00:28:47.730 \longrightarrow 00:28:48.746$  form at yalecancercenter.org.

NOTE Confidence: 0.9326976

 $00{:}28{:}48.746 \dashrightarrow 00{:}28{:}51.274$  We hope you'll join us next week to

NOTE Confidence: 0.9326976

 $00:28:51.274 \longrightarrow 00:28:53.199$  learn more about the fight against

NOTE Confidence: 0.9326976

 $00{:}28{:}53.199 \dashrightarrow 00{:}28{:}55.110$  cancer here on Connecticut Public Radio.

NOTE Confidence: 0.9326976

 $00:28:55.110 \longrightarrow 00:28:57.618$  Funding for Yale Cancer Answers is

NOTE Confidence: 0.9326976

 $00:28:57.618 \longrightarrow 00:29:00.000$  provided by Smilow Cancer Hospital.