Welcome again to Cancer Center.

Grand rounds, good afternoon and as is often the case, we have some really exciting data and progress and science and clinical care to share with you in this.

The theme for today, which is a really interesting one and different because we have multiple speakers, is in hematology where we’ve made enormous progress.

And we’re both biology translational work, population science and clinical care in
the major hematologic latency is leukemia and myeloid malignancy’s lymphoma. Multiple myeloma where I think we have really talented people working on these problems and really translating science into progress for our patients and looking forward to all four presentations. But I will actually turn to our Essentially, our leader and host for the session to make the introductions, so I’ll start simply by introducing Doctor Stephanie Allyne. As you know, Stephanie is the chief of hematology at Yale, as well as an associate.
00:01:20.200 --> 00:01:21.550 professor of medicine.
00:01:21.550 --> 00:01:22.374 In addition,
00:01:22.374 --> 00:01:24.434 Stephanie Overseas the DeLuca Center
00:01:24.434 --> 00:01:26.949 for Innovation and Hematology Research,
00:01:26.950 --> 00:01:28.950 which is really an exciting
00:01:28.950 --> 00:01:31.900 addition to our work in hematology,
00:01:31.900 --> 00:01:34.498 with the promise that not only
00:01:34.498 --> 00:01:35.797 translating our discoveries
00:01:35.797 --> 00:01:38.030 into clinical care innovation.
00:01:38.030 --> 00:01:41.144 But also in furthering the support
00:01:41.144 --> 00:01:44.200 of our faculty and trainees.
00:01:44.200 --> 00:01:47.528 Stephanie joined the faculty in Yale in 2006.
00:01:47.530 --> 00:01:50.738 An we were so pleased that she was
00:01:50.738 --> 00:01:53.462 selected as our new Chief in 2020
00:01:53.462 --> 00:01:56.771 and is doing a great job and looking
00:01:56.771 --> 00:01:59.543 forward to her an our faculty
sharing the exciting work at Yale.

So Stephanie I turn to you.

Let me share my slides.

OK, can you see the slides well?

Yes, OK, Alright,

let’s tally Saturday will preside,

And there will be four of us,

so I will be the one driving the slide.

So any hiccups blame me.

Um, so I’ll be.

I’ll be giving a brief introduction

to hematology and then tomorrow

probably will present on behalf
00:02:33.671 --> 00:02:36.161 of the myeloid malignancies team
NOTE Confidence: 0.87615436
00:02:36.161 --> 00:02:37.655 and traumatizing associate,
NOTE Confidence: 0.87615436
00:02:37.660 --> 00:02:38.974 professor of medicine,
NOTE Confidence: 0.87615436
00:02:38.974 --> 00:02:39.412 hematology,
NOTE Confidence: 0.87615436
00:02:39.412 --> 00:02:42.511 and he’s a director of the MLP
NOTE Confidence: 0.87615436
00:02:42.511 --> 00:02:44.091 malignancies disease team and
NOTE Confidence: 0.87615436
00:02:44.091 --> 00:02:46.859 the firm chief of the data firm,
NOTE Confidence: 0.87615436
00:02:46.860 --> 00:02:49.050 and everybody knows just how
NOTE Confidence: 0.87615436
00:02:49.050 --> 00:02:51.702 much moving Anan changes had to
NOTE Confidence: 0.87615436
00:02:51.702 --> 00:02:54.006 happen over the past years to
NOTE Confidence: 0.87615436
00:02:54.006 --> 00:02:56.389 keep our clinical services alive.
NOTE Confidence: 0.87615436
00:02:56.390 --> 00:02:58.754 And to my actually completed his
NOTE Confidence: 0.87615436
00:02:58.754 --> 00:03:01.840 doctor rent in medical hematology oncology.
NOTE Confidence: 0.87615436
00:03:01.840 --> 00:03:04.150 Myung then joined the Paoli
NOTE Confidence: 0.87615436
00:03:04.150 --> 00:03:05.998 Calmettes’s Institute in Marseille
NOTE Confidence: 0.87615436
00:03:05.998 --> 00:03:08.715 and did a full ride scholarship
at Johns Hopkins University, and I think that’s how eventually he landed with us. With Steve Gore recruiting him. After tomorrow, Doctor Francine Foss, Professor of Medicine and hematology and Rheumatology, would present on behalf of the former team and. Francine really is the expert in T cell lymphoma, internationally, nationally, renowned, and she’s a research director of an informal disease team and obviously director of the Tucson Informer program.
and you will hear exciting news

what is happening in the format to

selling Thelma and last but not least,

doesn’t Italian infrared,

so will present on behalf of the

multiple myeloma disease team.

And she’s the research director

of the much more disease team

and an assistant professor in.

Medicine medical oncology.

She actually obtained her medical degree

in Tennessee and the country of Georgia,

then did postdoctoral fellowships at Emory

Northwestern and Lucky for us a deal.

She did her residency and fellowship.

Odicle refill Oscar left us,
but we were able to recruit her back to be part of the team.

So um, before I introduce much about hematology, I think it’s absolutely time that we all say thank you to Charlie. So Charlie, you took over as our interim section Chief in January 2018 and has been a just wonderful, wonderful journey since then. So you can see just how big the group is in hematology and growing. And I just want to read a few of the quotes from our faculty.
And that is so Charlie,
you recognize the strength,
resilience and enthusiasm in
the new teologi faculty, staff,
advanced practice providers,
nurses entry needs,
and you supported our teams in so many ways.
Thank you for the leadership and
commitment you provided as interim Chief.
Your strong support and advocacy
for our section is not appreciated.
Thank you for being an outstanding
leader and role model for trainees
through your dedication to
excellence in patient care,
teaching and innovation.
It has been an honor to learn from you and the excellent faculty of your Cancer Center and thank you for everything you've done for the Cancer Center and was a section of hematology over the past several years. You have been a wonderful resource and a guide from my academic growth, and I'm extremely grateful to you. So thank you, Charlie. Thank you in the entire division. It's a real point of pride to me to have been a member and now along of this very August center of excellence. So thank you.
And we have a little bit more in store, so the thing we really wanted to thank you for is how you have supported all our people and recruitment of all these people highlighted in yellow. You know Murrayshall is to my like my name is this team chilling costar Intercine safely to the lymphoma team. Barbona is our director of classic hematology, Sabrina Browning as a member of our myeloma and come up as a team. And as you can see, also of our billing team. And then in addition to that, you have recruited hematology focused faculty to the to the Smiler.
network and the care centers, and we’re so fortunate to work with all our colleagues. And of course Mike’s mission as a director of the Center for Molecular and Cellular. I always say hematology and oncology in there were so excited about, you know, all our new people and teams and what we can achieve together. So, um, so now in my introduction to hematology. So what you see here is kind of our. These are hematology core teams. I say core because these are the people located in New Haven,
North Haven.

And these are the teams that are working hard to bring the best patient care to all patients in Connecticut and beyond through their research that clinical care and collaboration was all the teams we have smile. Oh yeah, even health and University. So we have five teams in my life. malignancies team, an informer team, the stem cell transplant, the malignancies team, an informer team, the stem cell transplant, and South Therapies team. We call it classical hematology because as we all know. Behind him, disorders are not that denied. Our patients are quite ill and then our
minds from my Loma and commodities team.

So as I mentioned before,

we're not alone.

We're not just a small,

tiny little point on the map in New Haven,

but through our smaller network we

have reached across the entire state

and it is just wonderful to know that

we can bring the excellent signs and

treatment to all patients in Connecticut.

Then again,

here I'm highlighting our

hematology focused faculty,

which is also new over the last.

Several years and it’s just wonderful.
How much team building can we do?

There are many discussions already going on between our disease team leaders and faculty and tremble in North Haven, Guilford, Hardford and all the other places.

So it’s wonderful opportunity. Um, we’re not alone and we’re not doing this in isolation, so we’re fortunate to be at a place like here where there are many different centers there.

We have our Decker Jews tallied in the clinical Trials Office. Again, Marcus leading that CMC L Michaels
NOTE Confidence: 0.7637411
00:08:39.889 --> 00:08:41.490 bosenberg leading yell center
NOTE Confidence: 0.7637411
00:08:41.490 --> 00:08:43.138 female oncologix Mark Lemon,
NOTE Confidence: 0.7637411
00:08:43.140 --> 00:08:45.105 leading the year Cancer Biology
NOTE Confidence: 0.7637411
00:08:45.105 --> 00:08:46.284 Institute kinda Neugebauer,
NOTE Confidence: 0.7637411
00:08:46.290 --> 00:08:47.954 leading the Larne Center.
NOTE Confidence: 0.7637411
00:08:47.954 --> 00:08:51.777 And then Diane Krause in the my C Age
NOTE Confidence: 0.7637411
00:08:51.777 --> 00:08:54.689 and then cheers of the different departments.
NOTE Confidence: 0.7637411
00:08:54.690 --> 00:08:55.779 Chengdu for pathology.
NOTE Confidence: 0.7637411
00:08:55.779 --> 00:08:57.231 David charts now Department
NOTE Confidence: 0.7637411
00:08:57.231 --> 00:08:58.320 we know immunobiology.
NOTE Confidence: 0.7818945
00:08:58.320 --> 00:08:59.768 We have collaborators in
NOTE Confidence: 0.7818945
00:08:59.768 --> 00:09:01.216 the stem cell center.
NOTE Confidence: 0.7818945
00:09:01.220 --> 00:09:03.768 We have collaborators in the Copper center.
NOTE Confidence: 0.7818945
00:09:03.770 --> 00:09:05.666 Marcella Nunez Smith just gave a
NOTE Confidence: 0.7818945
00:09:05.666 --> 00:09:08.237 talk to the leaders this morning
NOTE Confidence: 0.7818945
and Antonio riders and many,
NOTE Confidence: 0.7818945

many more that I can’t even
NOTE Confidence: 0.7818945

get on this slide.
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So, um, I just wanted to give a brief
NOTE Confidence: 0.84497094

update on the Delucas Center for
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Innovation and Hematology Research,
NOTE Confidence: 0.84497094

another amazing initiative that Charlie
NOTE Confidence: 0.84497094

made happen and we are now we’ve just
NOTE Confidence: 0.84497094

riding our progress report for the past
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two years and just to give you an update.
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So we have a very active hematology
NOTE Confidence: 0.84497094

tissue bank with samples from patients
NOTE Confidence: 0.84497094

with all human to logic disorders.
NOTE Confidence: 0.84497094

We have over 4000 samples from over 2000.
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Nations, we have a bone marrows prefer blood.
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We’re working **** ** also collecting lymph
node tissue and other tissue biopsies.

And we also offer specialized processing for clinical trials in hematology.

And this is again a huge.

Team effort so you see our core people on the right and Michelle Patel, rounder, Feli, Padma Mamillapalli and our latest recruit Jennifer from donor Hoban.

But everybody in hematology is contributing to this.

All our aisps an MP7 in North Haven at Trumbull.

Another care centers and we hope to leverage this tissue bank to better understand.
these users and offering treatments.

We have awarded 9 pilot grants or $50,000 each.

Our goal is to have these being through these be collaborative between collisions and basic scientists.

We have one career development award E chaining cathari.

These are two year awards and the our base posted for new applicants.

We have many many applications publications that I could not list here.

We have installed the freezer works by repository database and we’re very close to populating it.

This data that eventually will allow
NOTE Confidence: 0.84497094
00:10:58.196 --> 00:11:00.190 clinician scientists in a deidentified
NOTE Confidence: 0.84497094
00:11:00.190 --> 00:11:02.745 manner screen what we have in the
database so they can come up with ideas
NOTE Confidence: 0.84497094
00:11:02.745 --> 00:11:05.173 and let us know how we can help them.
NOTE Confidence: 0.84497094
00:11:05.173 --> 00:11:10.076 We're providing access to novel technologies.
NOTE Confidence: 0.84497094
00:11:10.080 --> 00:11:11.860 Jennifer Amisha Jazz performed single
NOTE Confidence: 0.84497094
00:11:11.860 --> 00:11:13.640 site DNA sequencing for clinical
NOTE Confidence: 0.84497094
00:11:13.700 --> 00:11:15.280 trial samples for to Maccabean.
NOTE Confidence: 0.84497094
00:11:15.280 --> 00:11:23.620 We have many more ideas and we are also
NOTE Confidence: 0.84497094
00:11:17.836 --> 00:11:20.236 seeking to provide Technical Support for
correlative studies and data analysis.
NOTE Confidence: 0.84497094
00:11:20.236 --> 00:11:27.130 So let me now go over to the disease teams,
00:11:27.130 --> 00:11:29.461 so this these are the members of the stem
NOTE Confidence: 0.84497094
00:11:29.461 --> 00:11:32.039 cell transplantation cellular therapies team,
NOTE Confidence: 0.84497094
00:11:32.040 --> 00:11:34.092 but they will not present today
NOTE Confidence: 0.84497094
because you should stay tuned and log into the Young Cancer Center grounds.

I think it’s March 23rd when Doctor Supinder for her with her present on cellular therapies.

You will also not here today from our classical hematology team, because Doctor Boner just presented not so long ago in grand rounds.

But we’re also very hopeful to getting a slot in the summer when we will have, hopefully exciting new recruits to add to the team.

And we can take a full grand rounds for our classic hematology.

So the team you will be hearing from today.
The first one is on myeloid malignancy's team and Doctor Priebe will present on behalf of this group and so now I'm heading over to Doctor Kirby for the first presentation. OK, thank you. Thanks a lot Stephanie, within blessed to have already Nikolai in myself. Within single, fewer a few weeks ago on this ground round and I guess that you know probably most of the members of the team already. That on our side and that any color without self on universe recruit Rory share is.
Doctor Brokaw from San Francisco and up to it from Trimble that collaboration is basically something that is extremely important for us. And on the translational abside Stephanie for sure that has been instrumental in the group as well as Manoj Pillai. Next slide, please.

So as you may know, in MLP malignancy daughter asked basically 10 years we had a real paradigm shift thanks to the revolution of our understanding of the genomics of the disease. That’s been true in acute myeloid leukemia, so that’s really fast.
Through in the acute myeloid leukemia.

Then in my dysplastic syndrome, but is basically representative of what we have seen also in our knowledge of my wife, activities, orders, or bone marrow failures, for example, and this revolution, translated more recently. And we can switch style.

Um, in a boarding shift also in the management of this disease and since 2017 we add more approval than over the last 2025 years that goes with basically drug at R agnostic of any.
Potential genomic amenities such as the CP351 for example or more recently than 8:00 o’clock, but also drug that are no standard of care for basically mutated diseases, such as flat 3 mutated AML or, potentially IDH mutated diseases. Interestingly, this revolution that started with the access my leukemia, we’re starting to see speaking up. Also, in multiple packages, orders with the recent approval of Fedratinib as well as with my dysplastic syndrome with recently the approach the TGF beta inhibitor spider step
as well at the oral decide to be in. Stop all of that is extremely important and allow us to basically.

Have a real improvement for or patient regarding prognostic forgetting option of treatment. But we still in situation where a cute smile, leukemias and melodies plastic syndromes are how to treat disease. If you standardize the incidence rate of looking at my leukemia, that’s still in the top five of the most deadliest cancer that we have to face in the US and one of the goals that we had.
In the group has been to really work at different aspect to try to improve the result of the different treatment and we can switch to the next slide. We really tried to work from single cell to population research and population outcome and that’s basically to have a stun. But that’s really what is the MLP malignancy group going from. Stephanie was mentioning our first result in a single cell sequencing in acute myeloid leukemia.
As the researcher Stephanie Ann manner. In the labs around uh. Mouse model of. Testing syndromes RNA for montage to the collaboration that we have with the Cobra Group, led by AMR Nikolai and Rory on outcome research. From a pure clinical research, sense points or group has been instrumental in several key studies that led to approval of these drugs over the last few years, but or portfolio and our goal is really focused on early development of a new agent. Right now click portfolio is.
00:17:21.790 --> 00:17:26.568 roughly 40% of phase one trials.
NOTE Confidence: 0.8108151
00:17:26.570 --> 00:17:28.580 With.
NOTE Confidence: 0.8108151
00:17:28.580 --> 00:17:31.247 Basically a good #6 out of 19
NOTE Confidence: 0.8108151
00:17:31.247 --> 00:17:33.530 active trial that our investigator
NOTE Confidence: 0.8108151
00:17:33.530 --> 00:17:36.692 initiated trial and thanks to all
NOTE Confidence: 0.8108151
00:17:36.692 --> 00:17:39.618 the commitment of the research staff.
NOTE Confidence: 0.8108151
00:17:39.620 --> 00:17:42.626 Thanks to all of the commitment
NOTE Confidence: 0.8108151
00:17:42.626 --> 00:17:45.444 of the different members of Group
NOTE Confidence: 0.8108151
00:17:45.444 --> 00:17:48.308 we right now at a rate of index
NOTE Confidence: 0.8108151
00:17:48.402 --> 00:17:50.907 cases included in clinical trials
NOTE Confidence: 0.8108151
00:17:50.907 --> 00:17:55.301 on main campus that is 15 to 20%.
NOTE Confidence: 0.8108151
00:17:55.301 --> 00:17:57.809 Which is pretty remarkable.
NOTE Confidence: 0.83947045
00:17:57.810 --> 00:18:01.394 We definitely aim to do better and
NOTE Confidence: 0.83947045
00:18:01.394 --> 00:18:04.996 we definitely aim to be able to
NOTE Confidence: 0.83947045
00:18:04.996 --> 00:18:07.948 export the research we’re doing from
NOTE Confidence: 0.83947045
00:18:08.050 --> 00:18:11.122 the main campus to different side
00:18:11.122 --> 00:18:14.974 of the network and be able to add
better access for our patients in the
network as mentioned by Stephanie,
we have pretty efficient and pretty large.
Think annotate by bank and I think
we can give kudos to Stephanie
to each of the exports in 2011,
so we’re in a decade in right now.
We should definitely celebrate about that.
And that gives us a lot of leverage,
potentially to be able to develop some
translation research.
As well as potentially bringing
some collaboration from the outside
with pharmaceutical companies as
well as from academic centers and
go to the next next slide.
Just wanted to highlight basically two
Basically going on the first one are the
Remember first discussion years ago
Remember first discussion years ago
when we were in Hopkins together
around potentially all this knew,
around potentially all this knew,
around the additional,
uh, the 1st generation, but also right now the second
geneneration of even the energy agent
to conventional therapy.
We're right now activating.
Very broad induction chemotherapy plus volume
study under the umbrella of a sitter.
With the first patient that we hope
to include in the next few weeks,
we just have finished the,
cool on the map plus entinostat NDS
trial for the group of patients with
mild agent Majestik syndrome
00:20:19.915 --> 00:20:21.935 failed by accommodating agent,
NOTE Confidence: 0.83947045
00:20:21.940 --> 00:20:24.694 and we are looking forward for
NOTE Confidence: 0.83947045
00:20:24.694 --> 00:20:27.748 this data that has been basically
NOTE Confidence: 0.83947045
00:20:27.748 --> 00:20:31.120 possible to to develop thanks to.
NOTE Confidence: 0.83947045
00:20:31.120 --> 00:20:33.785 Elaboration with Yale Science and
NOTE Confidence: 0.83947045
00:20:33.785 --> 00:20:36.960 that’s work with initially TK came.
NOTE Confidence: 0.83947045
00:20:36.960 --> 00:20:40.770 One of us still here and we have also a
NOTE Confidence: 0.83947045
00:20:40.869 --> 00:20:43.894 chronic my leukemia studies sponsored
NOTE Confidence: 0.83947045
00:20:43.894 --> 00:20:48.198 by equal on the addition of bambrough
NOTE Confidence: 0.83947045
00:20:48.198 --> 00:20:51.714 on tyrosine kinase inhibitor for a
NOTE Confidence: 0.83947045
00:20:51.714 --> 00:20:54.942 patient with chronic migraine leukemias,
NOTE Confidence: 0.83947045
00:20:54.942 --> 00:20:57.678 an Axia Resul disease.
NOTE Confidence: 0.83947045
00:20:57.680 --> 00:20:59.135 Beside this trial,
NOTE Confidence: 0.83947045
00:20:59.135 --> 00:21:01.560 we also have been implicated.
NOTE Confidence: 0.83947045
00:21:01.560 --> 00:21:04.000 Basically thanks to our leadership
NOTE Confidence: 0.83947045
00:21:04.000 --> 00:21:07.464 on some of the development of the
newest generation of Elon College.

Agents such as the T in three inhibitor and we hope to have more results to show that in the near future next slide.

So the idea hmm DS program? That’s something that I already. Presented basically a few weeks ago during the ground rounds, but I definitely like this program as it’s really a homegrown program thanks to the work and interaction with Stephanie and Ranjit. So that’s really a good example of what we can achieve at yelling working together. We have been able to show how
we can exploit the weaknesses of IDH mutated AML in NDS. As you can see, when we have an idea of mutation and the presence of two HG in the media that creates some braknis phenotype that can be potentially targeted by using popped inhibitor on the upper right side. Can see that treatment with olaparib in a primary samples in grafted in mice can potentially significantly reduced the disease burden in. Lisa grafted with acute myeloid leukemia.
the resistance to IDH inhibitors in the clinic and that led to the step trial. Brentley ongoing with right now. Six patient included, including four at. At here with basically the use of elaborate the parking meter. For a male and MBS patient with IDH mutation. Interestingly, for the discussion with the tap, we’ve been able to have a patient that were not yet exposed to IDH limit, or potentially integrating the trial with an early evaluation that would also potentially to switch to ID age limit,
or there was no some clearcut benefit after a few weeks on treatment, so.

That’s two of the main program that we have right now.

I’m not going to basically talk too much about the outcome.

Nicole, I would review of what was going on a few a few weeks ago, but as we were discussing we really want to be able to to bridge the most basic science aspect of it. What we’re doing in the in the labs to this outcome. Research go go ahead.

We still have a lot of people
00:24:19.992 --> 00:24:22.160 in the results clinical issue.

00:24:22.160 --> 00:24:23.628 Then that's the topic.

00:24:23.628 --> 00:24:28.406 We're not the addressing all of them, but some of them are definitely on the focus of the group.

00:24:28.406 --> 00:24:35.520 The management of high risk disease in AML and MD S complex karyotype.

00:24:35.520 --> 00:24:38.033 NLL mutated or rearrange disease are one of the focus of development of the group.

00:24:38.033 --> 00:24:41.076 And I specially Rory is developing some different concepts around.

00:24:41.080 --> 00:24:44.044 We talked about that one of the challenges, the optimization of targeted therapies and email steropes.
basically that we’re still working with the disease that is not as common as breast or colon cancer, and we end up with more combination and sequence of agent then patient. And that’s potentially where the interaction with the lab and the work that we’re doing with Stephanie is extreme and mineral. Extremely important. Targeting of leukemia stem cell looking initiating cells and management. Measurable residual disease at the molecular level is obviously something that is on our mind, and that’s a lot of interaction.
Also with the transplant group in South Therapy Group on our potentially to work around this concept. And last but not least, before getting to the point where we have active disease, working on predisposition and potentially climb. This is something that is extremely interesting right now. In collaboration we will have. With the cardiology group and the Generation project,
Feel that we’re pretty blessed in the group to have already’s. Pretty solid foundation, but we wanted to do better and that’s gonna be by leveraging the excellence that we have it here the molecular biology aspect with basically right now are some development in single cell sequencing using multi omics approaches that can potentially be used in my life but not only in my right mind agencies. Potential collaboration and avenues of collaboration with the yellow. Center of immuno oncology. And with a group of Marcus mention also to potentially work on resistance disease,
Immuno oncology in my redeemer agencies and the really fruitful and really exciting collaboration we have with the Copper Group carry gross schalmei. With a lot of publication and lots more that will come in the next. We want to expand our collaboration. That’s going to be more collaboration and the basic Science Group partners like June Liu Shan Quinn Go are definitely people that will be more on the stage with us in more portable computers as well as more collaboration on the clinical research side.
Obviously with the BMT and cell Therapy Group, but also without regard for the adolescent young adults, and I think that’s all I have.

Thank you so much tomorrow. This is really awesome, so we’ll go straight into our next presentation by Doctor Falls on behalf of Mylan Malignancy’s team.

Hi, this is Francine and I’d like to thank Stephanie and also Charlie for all of your work to bring us to where we are today with this faculty on the development that we’ve had in the lymphoma program.

So lymphoma program consists of myself,
Scott Huntington, ERISA, Soupy, Shalin, Kothari, Toshin Sethi, Francesco Montanari, both of whom have recently joined us. Francesca is in Greenwich.

This is the landscape of lymphoma. So lymphoma really is many different diseases, and that’s a challenge for us as a group with limited resources. So it turns out that out of the 21,000 cases of cancer in Connecticut per year, there are about 900 plus or minus cases.
00:28:40.277 --> 00:28:42.499 cases of non Hodgkin’s lymphoma.
NOTE Confidence: 0.752058699999999
00:28:42.500 --> 00:28:44.535 So that’s really our denominator
NOTE Confidence: 0.752058699999999
00:28:44.535 --> 00:28:46.942 population in our efforts are to
NOTE Confidence: 0.752058699999999
00:28:46.942 --> 00:28:49.063 try to get many of these patients.
NOTE Confidence: 0.752058699999999
00:28:49.070 --> 00:28:50.454 Want to clinical trials?
NOTE Confidence: 0.752058699999999
00:28:50.454 --> 00:28:53.258 If you look at the curve at the
NOTE Confidence: 0.752058699999999
00:28:53.258 --> 00:28:55.533 bottom you can see the frequency of
NOTE Confidence: 0.752058699999999
00:28:55.533 --> 00:28:58.123 the different types of non Hodgkin’s
NOTE Confidence: 0.752058699999999
00:28:58.123 --> 00:29:01.126 lymphoma with diffuse large B cell lymphoma,
NOTE Confidence: 0.752058699999999
00:29:01.126 --> 00:29:01.932 follicular lymphoma,
NOTE Confidence: 0.752058699999999
00:29:01.932 --> 00:29:04.350 the low grade marginal zone lymphoma
NOTE Confidence: 0.752058699999999
00:29:04.413 --> 00:29:06.315 is being the most common subtypes,
NOTE Confidence: 0.752058699999999
00:29:06.320 --> 00:29:07.457 and T cell,
NOTE Confidence: 0.752058699999999
00:29:07.457 --> 00:29:09.352 lymphoma and mantle and mantle
NOTE Confidence: 0.752058699999999
00:29:09.352 --> 00:29:11.200 cell lymphoma and burqas are.
NOTE Confidence: 0.752058699999999
00:29:11.200 --> 00:29:13.264 You can see are rare compared
to these other common subtypes.

Next slide.

Um, so the unresolved political issues, of course, are how we make sense of 90 different subtypes of lymphoma and divide those into general themes.

For clinical trials, I’ll talk a little bit about that in a minute, but you know, we do have diseases where the modern regimes pretty much are OK in terms of producing favorable outcomes for these patients,
but there are within these more favorable subgroups of patients. Those patients with high risk molecular features that continue to fail conventional therapy and would be subjects for novel clinical trials and those include double hit diffuse, large B cell lymphoma and one of the things that we. Done with that disease is to combine dose adjusted epoch with Lenalidomide. In addition, we have some patients with low risk lymphoma where we could actually test D escalation of therapy and a good example of that is classical.
Hodgkin’s disease and I’m going to show you another example. For diffuse large B cell lymphoma we also have histologies where the modern regiments are in ineffective, pretty much inadequate for these aggressive patients such as the T cell lymphoma’s. The piece of distributing mantle cell lymphoma’s and the post transplant lymphoproliferative disorders so in these cases where are trying to incorporate novel agents into the frontline but also trying to figure out how to incorporate.
allogeneic stem cell transplant and the newer car T cell therapies, would love to use molecular and genetic profiling in these diseases to select patients rationally for pathway directed clinical trials, thanks to the work of Stephanie. To get our tissue bank up and running, Sorry sorry, OK next slide. So, um, just touching on how we’re deescalating therapy. This is a really great example of a trial that Scott has opened for relapsed aggressive B cell lymphoma’s where we have a non chemotherapy approach.
So basically this study involves oral agents of BTK inhibitor and mtor inhibitor, an image, and it turns out that when you combine these three oral therapies, this is a very effective strategy for patients with aggressive lymphoma who have failed conventional chemotherapy. Some of them have failed transplant as well. On the next slide, Stephanie, you can see an example of one of our patients. It turns out that yell actually has involved the majority of patients that involved the majority of patients that patients to date in this trial and responses have been observed among.
a different subtype of patients.

You can see with DL, BCL with low grade lymphoma, mantle cell lymphoma etc and patients with very dramatic responses with large masses and extensive disease. So we were really excited about this approach. And again, we’d all like to start thinking more about oral rather than Ivy therapies for patients that are going to be chronically needing therapy, such as many of our lymphoma patients. Unfortunately, next slide would highlight another approach,
which is to take an active drug, look at its resistance mechanisms and come up with. Similar drugs that might be active in resistant mutational settings. In this case. BTK inhibitors are one of our most active newer agents in B cell lymphoma, but most of the failures to these agents are related to the development of specific mutations and now we have these new non-covalent BTK inhibitors that actually are able to subvert those mutations and induced responses in patients and this is an
agent that we’re working with now.

00:33:01.360 --> 00:33:02.214 Locks 0305.

00:33:02.214 --> 00:33:04.349 Falls into that category that’s highly active.

00:33:04.349 --> 00:33:05.203 We are currently embarking upon a number of phase two studies that we hope to make available broadly across our care centers looking at a number of different B cell lymphoma is with these novel agents.

00:33:07.346 --> 00:33:09.611 Next slide.

00:33:09.611 --> 00:33:11.909 Um, this is some interesting work and translational research done by shelling pathari in our group.

00:33:11.909 --> 00:33:13.800 So Charlyn is looking at ways of.
Dealing with the development of resistance to active agents such as phonetic, LAX, and hear a set of experiments where he has shown that proteasome inhibition is synergistic with phonetic lacks in patients with B cell lymphoma an. In these clinical models, so you can see the bot is mid carve.

Is Mebane exacerbated by themselves, which are proteasome inhibitors have some activity to induce a pop ptosis? As does phonetic LAX? But when you combine these agents you can see synergistic activity.
And based on this in vitro work that Charlene has done, he’s embarking on a seat app sponsored Phase 1, two clinical trial where he’s combining genetic LAX with these proteasome inhibitors and we’re very excited for Shaolin to get that trial up and running on the next slide, you can see the correlative science that’s developed around this clinical trial, including a single celled Association to look at these specific BCL, two family members. He’s also doing exome sequencing,
RNA seek and trying to obtain tissues from these patients for later PDX development. So that’s all very exciting and we’re really proud of shelling for that. In additional, he’s doing what other groups are doing to embark on looking for a circulating tumor DNA in these patients, and hopefully some of us will be able to incorporate that into our disease group studies as well next slide. So and then the last thing I want to mention is exciting work that Charlene is doing with the Cats lab to use a different approach called protest back to target antiapoptotic.
protein proteins potentially, but their critic benefit soap Protex basically is a hetero bifunctional small molecule that consists of a linker and two warheads. One of them binding to the target protein such as the BCL two and the other recruiting the E3 ligase to basically leads to the degradation of that. Protein, so we're looking forward to this work coming to fruition in the clinic in the future. Just highlights some of the work that Scott has done working with the Copper Group to look at at
outcomes in patients with lymphoma.

And so Scott has done a number of studies including these two, one of which shows that patients with who are older than 80 have a higher frequency of discontinuing active therapies like a brute, not within the first 180 days. So obviously one has to go back and look at why that is and ways that we can. Alter that therapy so those patients can continue to be treated and then also Scott has done a lot of work with cost effectiveness and this is just one of his studies looking.
at different treatments for CLL.

Getting a brute nip upfront versus getting it later on.

So again, Scott has presented some of this work at the national meetings.

Then there are a number of studies ongoing with copper.

Next slide will then segue us into T cell lymphoma.

Is overall heterogeneous as arviso bonas and tend not to do as well with conventional therapies as you can see in the outcomes curve.

progression free survival in median survival is poor for many of these.
Aggressive T cell subtypes.

The next slide.

Highlights one of the things that we've been trying to do so patients with T cell lymphoma get chop based chemotherapy upfront and only a small percentage of them actually are cured.

With this approach, patients would go on to an autologous stem cell transplant if they have a complete remission, but it turns out that when you look at registry studies, one of which we conducted here, only about 25% of patients
upfront ever make a transplant. The reason being that many of those patients don’t have a good remission. And so this is a study that I worked on with tar Sheen and this study is hopefully going to get started very soon where we combined Mogamulizumab with upfront chemotherapy. In this case, epoch for patients with aggressive T cell lymphoma. The idea being that Mogamulizumab is a CCR four monoclonal antibody that targets both tumor cells at but also more importantly T regs in the micro environment, so we’re hoping that there’s going to be an interaction.
Both in the micro environment as well as potential synergy with the tumor cells. When this is combined with chemotherapy, she has also developed some very nice correlative studies to go along with this. The next slide. Will take you into the world of relapsed T cell lymphoma and just to demonstrate to you the work of our group to look at a number of different agents targeting a number of different pathways that are relevant and some of these studies. Yale has been the top one or two in terms of accrual for these studies.
nationwide and these are novel agents and I'll touch on a couple of them in the next slide or two.

Um, go back and this is this is next slide.

OK, this is tipifarnib which is a farnesyltransferase inhibitor, but it turns out that it also down regulates CX CL 12 which is in the micro environment patients who have expression of CX CL 12 in the micro environment don't do as well as you can see on this survival curve and we've had some incredibly dramatic responses using this single oral agent and some of our patients
such as this gentleman who has failed multiple therapies in autologous transplant that this is really salvage. A lot of people were hoping to initiate some IIs with this molecule in the next couple of months. Next slide shows you another approach with micro RNA, so this is the first in man study of this micro RNA which targets a number of different types of lymphoma. We put a number of patients with cutaneous lymphoma on this trial and you could see responses in pretty much all of the patients that were treated.
based on these waterfall plots and
One of the most important findings.
This micro RNA, which is its activity in HTLV one
associated adult T cell leukemia,
and again we put the majority of
patients in this cohort on this
trial and we also initiated the
correlative studies that you see below,
showing that the molecule directly
inhibits the proliferation of ATL cells.
Modulates the expression of various
activation markers and you can see
the changes in proliferation index
with this molecule and this is.
All samples from our patients that were drawn at different time points, so we're very excited to get this data published.
some of the efforts that Francesca has looking at in a couple of different areas. In the context of aggressive lymphomas, this is her global T cell consortium study, which she conducted at Columbia and is now brought to Yale, and this is now a randomized phase two study where she’s looking at 5:00 or oral is incited gene in Rome and Epson compared to investigators choice. This is a multicenter study.

And we’re really excited that Francesca has brought this to us and the next slide is another effort in an area that we have not explored. Which is PTLD so post transplant
lymphoproliferative disorders.

In this study, Francesca is using sequential treatment for patients that are CD 20 and CD 30 positive and you can see the scheme for this trial here. This is also an IIT that’s being conducted in collaboration with the Mayo Clinic.

So I think we have some really nice work from some of our younger investigators and we’re very excited about that.
00:42:19.443 --> 00:42:21.950 to go in the lymphoid malignancy’s.
NOTE Confidence: 0.8136023

00:42:21.950 --> 00:42:24.074 So clearly we will benefit from
NOTE Confidence: 0.8136023

00:42:24.074 --> 00:42:25.947 the annotated database that you’ve
NOTE Confidence: 0.8136023

00:42:25.947 --> 00:42:28.287 heard about and hopefully will be
NOTE Confidence: 0.8136023

00:42:28.287 --> 00:42:30.159 contributing our samples in an
NOTE Confidence: 0.8136023

00:42:30.159 --> 00:42:31.744 ongoing fashion to the biobank.
NOTE Confidence: 0.8136023

00:42:31.750 --> 00:42:34.158 We also need to develop a sequencing
NOTE Confidence: 0.8136023

00:42:34.158 --> 00:42:36.018 platform which we really don’t
NOTE Confidence: 0.8136023

00:42:36.018 --> 00:42:37.928 have here at the institution.
NOTE Confidence: 0.8136023

00:42:37.930 --> 00:42:39.378 For lymphoma we’re currently
NOTE Confidence: 0.8136023

00:42:39.378 --> 00:42:40.826 sending our samples out,
NOTE Confidence: 0.8136023

00:42:40.830 --> 00:42:43.002 but we certainly would like to
NOTE Confidence: 0.8136023

00:42:43.002 --> 00:42:45.500 do that in the near future.
NOTE Confidence: 0.8136023

00:42:45.500 --> 00:42:47.474 I talked about the biobank and
NOTE Confidence: 0.8136023

00:42:47.474 --> 00:42:48.790 how important that is,
NOTE Confidence: 0.8136023

00:42:48.790 --> 00:42:51.454 but also we’re now starting to talk about
thematic direction for the program, and one of the areas that we’re focusing on at least in T cell lymphoma, as you’ve seen, is on the micro environment, and you know immunomodulatory strategies that can be used with correlative science in these two cell lymphoma studies, and then finally, I just want to acknowledge the translational science collaborators and this by no means is an exhaustive list of folks, but there have been a number of Studies that have been funded by both DeLuca.
00:43:19.260 --> 00:43:22.137 as well as other mechanisms for funding,
NOTE Confidence: 0.8136023
00:43:22.140 --> 00:43:24.426 and those include studies done with
NOTE Confidence: 0.8136023
00:43:24.426 --> 00:43:27.028 the Cats lab with Marcus is lab,
NOTE Confidence: 0.8136023
00:43:27.030 --> 00:43:28.910 which is now being engaged
NOTE Confidence: 0.8136023
00:43:28.910 --> 00:43:30.038 with these studies.
NOTE Confidence: 0.8136023
00:43:30.040 --> 00:43:32.302 The Lola slab in pharmacology and
NOTE Confidence: 0.8136023
00:43:32.302 --> 00:43:35.013 even the imaging labs with a nuclear
NOTE Confidence: 0.8136023
00:43:35.013 --> 00:43:37.275 medicine with Doctor Chi and others
NOTE Confidence: 0.8136023
00:43:37.275 --> 00:43:39.440 that I didn’t mention as well.
NOTE Confidence: 0.8136023
00:43:39.440 --> 00:43:42.437 So I think we have a very bright future
NOTE Confidence: 0.8136023
00:43:42.437 --> 00:43:45.445 ahead of us in the lymphoid malignancies.
NOTE Confidence: 0.8136023
00:43:45.450 --> 00:43:46.202 Thank you,
NOTE Confidence: 0.8136023
00:43:46.202 --> 00:43:46.578 Stephanie.
NOTE Confidence: 0.8164225
00:43:48.050 --> 00:43:49.630 Thank you for I’ve jumped.
NOTE Confidence: 0.8164225
00:43:49.630 --> 00:43:51.590 Thank you for seeing that is after
NOTE Confidence: 0.8164225
00:43:51.590 --> 00:43:53.816 you tour Tour de force and lymphoma
00:43:53.816 --> 00:43:55.808 is so excited about everything that
00:43:55.872 --> 00:43:58.112 is to come and very excited about
00:43:58.112 --> 00:43:59.710 our next presentation by Doctor
00:43:59.710 --> 00:44:01.600 Natalia never eats a on multiple
00:44:01.600 --> 00:44:02.617 myeloma in commodities.
00:44:02.617 --> 00:44:04.990 Stephanie, thank you so much for the
00:44:05.052 --> 00:44:06.816 opportunity to let me present today
00:44:06.816 --> 00:44:09.546 and I would like to echo some of your
00:44:09.546 --> 00:44:11.680 comments and thank thank Charlie for his
00:44:11.680 --> 00:44:13.570 outstanding leadership at the Cancer Center.
00:44:13.570 --> 00:44:15.495 So today I’m happy to present on
00:44:15.495 --> 00:44:17.469 behalf of myeloma team with brief
00:44:17.469 --> 00:44:18.917 clinical and research updates.
00:44:18.920 --> 00:44:20.810 Here’s our team myself. Terry Parker.
00:44:20.810 --> 00:44:22.470 Know far Bar and Sabrina.
00:44:22.470 --> 00:44:25.000
Browning as well as Elon Gorshin at Guilford.

Next slide please.

I’ll start with brief Brecht background about the disease.

As you all know, multiple myeloma is the clonal plasma cell neoplasm originating in the bone marrow. The diagnosis rests on the bone marrow biopsy and some of the key immunohistochemical stains and protein studies on the blood in the urine. The ETL pathogenesis of this disorder remains largely obscure.

We know of certain associations, such as perhaps antigenic stimulation.
Pathogenic microbes, lipid antigens and other associations such as metabolic syndromes, diabetes and such, but in large majority of patients we don’t know the cause and much remains to be elucidated in this research. Next slide, please.

So as you know, the diseases continuum and we know that myeloma every case is preceded by the precursor state called M Gus monoclonal gammopathy of undetermined significance and after years of its presence it progress.
00:45:26.760 --> 00:45:29.240 is to the early phase myeloma,
NOTE Confidence: 0.816422500000001
00:45:29.240 --> 00:45:31.110 commonly referred to as small
NOTE Confidence: 0.816422500000001
00:45:31.110 --> 00:45:32.606 during or asymptomatic myeloma.
NOTE Confidence: 0.816422500000001
00:45:32.610 --> 00:45:35.148 But a full blown clinical disease
NOTE Confidence: 0.816422500000001
00:45:35.148 --> 00:45:37.244 which requires active therapy is
NOTE Confidence: 0.816422500000001
00:45:37.244 --> 00:45:39.547 the disease which leads to end organ
NOTE Confidence: 0.816422500000001
00:45:39.547 --> 00:45:41.958 damage in the form of high calcium,
NOTE Confidence: 0.816422500000001
00:45:41.960 --> 00:45:44.270 renal failure, anemia and bone lesions.
NOTE Confidence: 0.816422500000001
00:45:44.270 --> 00:45:46.657 And in the old days we would
NOTE Confidence: 0.816422500000001
00:45:46.657 --> 00:45:48.299 resort to plane X Rays.
NOTE Confidence: 0.816422500000001
00:45:48.300 --> 00:45:48.708 However,
NOTE Confidence: 0.816422500000001
00:45:48.708 --> 00:45:51.156 this has been largely replaced by
NOTE Confidence: 0.816422500000001
00:45:51.156 --> 00:45:52.943 advanced imaging modalities such as
NOTE Confidence: 0.816422500000001
00:45:52.943 --> 00:45:55.360 whole body MRI or Whole Body PET CT scans.
NOTE Confidence: 0.816422500000001
00:45:55.360 --> 00:45:57.340 Next please.
NOTE Confidence: 0.816422500000001
00:45:57.340 --> 00:45:58.615 So once diagnosed,
the treatment pattern of multiple myeloma consists of initial induction therapy with usual 3 drug,
or in 2021 you might choose a four drug regimen incorporating some of the monoclonal antibodies upfront,
This is then followed by high dose melphalan, an autologous stem cell rescue AKA auto transplant,
and then this is subsequently followed by maintenance therapy,
which is usually long term and mental progression of disease.
So the.
Treatment pattern resembles a marathon rather than the Sprint. As we continue maintenance for many years, every patient experiences their relapse, and this is due to branching pattern of clonal evolution or push persistence. Relapse patients have to sequence through the available therapies, each of which then leads to shorter and shorter overall duration of response, eventually leading to dismal prognosis for many of their refractory patients.
Fortunately, we've had number of drug approvals over the course of the past decade, which includes the novel proteasome inhibitors in the form of carfilzomib, novel immunomodulatory agents, pomalidomide, and others. The biggest breakthroughs came in 2015, when we had several approvals, namely IgG monoclonal antibody targeting CD38 receptor on the plasma cell daratumumab, as well as checkpoint inhibitor, targeting SLAM F7 or CS1 elotuzumab. Orally available Proteus inhibitor.
We also had approval for histone deacetylase inhibitor PANOBINOSTAT which is orally available and more recently a drugs of completely different mechanism affection such as selenexor. This is a selective nuclear export inhibitor promoting some of the tumor suppressor gene action and within the past year approval of another IgG monoclonal antibody. Against CD 38 is I took up with slightly different mechanism of action, but very similar to daratumumab and more recently just summer of last year. First antibody drug conjugate.
got FDA approval.

This is Bill on time Out method Odin targeting B cell maturation antigen which is currently approved for relapsed refractory multiple myeloma.

Beyond four prior lines of therapy next. So despite these therapeutic advances, number of clinical challenges remain primarily the concerns about what to do for refractory myeloma. Are we able to, in the novel immunotherapy era, replaced the high dose melphalan by some of the novel strategies such as car T cell or other therapeutics?
How best to continue the maintenance therapy for patients particularly high risk data genetic subsets? And we're far from understanding the disease biology and choosing this sequential therapy based on biology of disease and its selection of therapy. Any cases is random and this is a clinical challenge. Same is true for a llama. Lodosa is, as you know, about 10% of multiple myeloma patients will develop concurrent soft tissue deposition of the Lambda or Kappa light chains.
and this is an unmet need in all of gammopathy world as these patients, the care is not well defined, so it remains to be established what is the optimal first line, second line and beyond therapy for patients with this next please. So while facing these challenging issues in clinic we tried to. Integrate our clinical expertise and incorporate some of the Noble research therapeutic strategies and continue to educate our patients as well as our trainees next. In terms of individual focus of
clinical expertise, Terry Parker has
an outstanding clinical expertise in
a llama low doses having served as.
National Pi and number of a alloy doses,
trials as well As for relapsed refractory myeloma.
Nofar nofar sparkles has been mostly on high dose melphalan.
Autologous stem cell transplant for myeloma as well as cellular therapies in myeloma.
Sabrina also has an excellent expertise
in a llama low doses shoot the separate fellowship in amyloidosis at the Boston Medical Center.
She also focuses on understanding the biology of some of the
00:50:28.748 --> 00:50:30.658 precursor States and my own.

00:50:30.660 --> 00:50:33.205 Clinical research has focused On’s

00:50:33.205 --> 00:50:35.241 to understanding clonal heterogeneity

00:50:35.241 --> 00:50:38.164 and incorporating advanced imaging in

00:50:38.164 --> 00:50:40.508 this disease therapeutic assessment.

00:50:40.510 --> 00:50:43.107 So education is of course an integral

00:50:43.107 --> 00:50:45.049 component of our daily work.

00:50:45.050 --> 00:50:47.535 This is a group of current fellows

00:50:47.535 --> 00:50:49.079 rotating through myeloma clinics

00:50:49.079 --> 00:50:50.719 during this academic year,

00:50:50.720 --> 00:50:53.331 and many of them have been involved

00:50:53.331 --> 00:50:55.249 in research projects in myeloma.

00:50:55.250 --> 00:50:56.294 So, for instance,

00:50:56.294 --> 00:50:58.382 Weixin Lou has been working on

00:50:58.382 --> 00:51:00.168 myeloma Yale Database project.
On looking at outcomes in patients treated with monoclonal antibodies, Talib Dasani has the bone disease in myeloma cohort and looking to develop quality improvement project started. Targeting this Eric Chang will be involved in this promise study, which is the screening and community outreach project for patients with gammopathy’s and their families, and it has an important outreach project, and we’re partnering with Dana Farber Doctor Ghobrial on this project. And finally, Rose Mirkin will be involved in the collaborative effort where we
may study the antigenic targets of monoclonal antibodies of myeloma with the help of immunology. Lab of.

So our research consists of different missions. On one hand, we always try to enhance our clinical trial portfolio. We have ongoing trials, both investigator initiated as well as industry and collaborative group trials. In every space of this disease, including early stage, small during as well as newly diagnosed and late relapsed refractory patients.
And we always try to develop additional therapeutic concepts for refractory myeloma.

Andale Emily doses.

On the other hand, we do have certain outcomes, research initiatives and this include both myeloma as well as Mgas database as well as bone disease.

In myeloma collaboration with our copper team and finally on the research basic Science Research front, we’ve been trying to build teams to collaborate with researchers to study monoclonal antibodies.

They’re driving antigens and Additionally mean profiling to understand predictive
biomarkers of disease response.

And study extramedullary disease next please.

So on the basic science research front,

Stephanie Collins, Biobank project for hematology has been really instrumental in our program at North Haven Myeloma program.

We’ve essentially biobank the bone marrow and peripheral blood on every single patient that we diagnosed with multiple myeloma or gammopathy, and we hope to build collaboration with the help from David Shatz lab to develop ex vivo multiple myeloma.
mouse models where we can develop robust preclinical models using this. In addition, studying monoclonal antibodies and understanding the antigenic targets and what drives the disease in collaboration with Aaron Rings lab on clinical research front, we've had active clinical collaboration and couple of IIT’s. Closely working with musculoskeletal radiology and in this and haimson ending lishouk, both have been instrumental as well as our pathology colleagues. Mean issue and sang and pen know.
Farhan Sabrina have ongoing collaboration with an Habermann from lab laboratory medicine to understand immune profiling of patients to understand the biologic response in different subsets of myeloma patients treated with monoclonal antibodies in terms of outcomes research. We’ve had active collaboration with hematology copper team. With the leadership of Shammai, my and Scott Huntington, we have several ongoing projects, and one of them a recent one using flat iron database and one particular
myeloma project examining patterns of care, especially in the context of kovid period. And Lastly, we've been trying to build this disease team, which we refer to myeloma bone disease program collaboration, under which we try to bring together different departments, including Endocrinology, Carlin, Sonia, as well as orthopedic and spine center, Dieter, Lindskog, and Louise called. And as I said, Talib Dosani has an ongoing outcomes project looking into bone disease in myeloma. So with that I will stop. I’m exciting to have such great
collaborators and colleagues on the campus.

And I will leave a couple of minutes for questions.

Thank you so much.

Awesome, alright, so I’ll share my screen.

Just people know we have hematology joins 2021.

We have a Twitter handle so we’ll try and use it,

so I’m going to unshare so we can see everybody and let me pull up the

chat to see if there are questions.

Are there questions?

Would anybody like to raise their
00:55:34.240 --> 00:55:36.430 hand and ask the question?
NOTE Confidence: 0.8470994
00:55:36.430 --> 00:55:39.121 If not, I can start with one and then
NOTE Confidence: 0.8470994
00:55:39.121 --> 00:55:41.483 maybe I’ll ask Francine since I’m
NOTE Confidence: 0.8470994
00:55:41.483 --> 00:55:44.225 always a molecular person and I know
NOTE Confidence: 0.8470994
00:55:44.225 --> 00:55:46.271 that we have this phenomenal effort
NOTE Confidence: 0.8470994
00:55:46.271 --> 00:55:49.066 to get on this goal panel and whole
NOTE Confidence: 0.8470994
00:55:49.066 --> 00:55:50.470 exome sequencing tower patients.
NOTE Confidence: 0.8470994
00:55:50.470 --> 00:55:52.576 Do you think that that will
NOTE Confidence: 0.8470994
00:55:52.576 --> 00:55:53.629 fulfill your need?
NOTE Confidence: 0.8470994
00:55:53.630 --> 00:55:57.490 Or what do we need to do to improve on that?
NOTE Confidence: 0.886944
00:55:58.450 --> 00:56:00.118 I think that’s certainly a start.
NOTE Confidence: 0.886944
00:56:00.120 --> 00:56:01.788 You know, as you know Stephanie,
NOTE Confidence: 0.886944
00:56:01.790 --> 00:56:03.930 we see some of these.
NOTE Confidence: 0.886944
00:56:03.930 --> 00:56:06.352 We are rare lymphoma patients for whom
NOTE Confidence: 0.886944
00:56:06.352 --> 00:56:08.131 there’s really no treatment algorithm
NOTE Confidence: 0.886944
00:56:08.131 --> 00:56:10.843 and which is kind of picking out of
a hat to try to figure out what to do in the error of precision medicine. I really think that we should be profiling these patients and sequencing them and rationally developing strategies, so we would love to be able to start doing that. I think the critical component of that is, you know, is being able to get hold of that issue when it’s biopsied, and to get it to the right place as soon as possible. We’re also very interested in looking at circulating tumor cells and whether that’s a strategy that
we could use as well.

OK fantastic exciting.

Are there any other questions?

I don’t see any in the chat.

So one of the I’m sorry Charlie friends, you

I think you each allude to it,

I don’t see any in the chat.

So one of the I’m sorry Charlie friends, you

I think you each allude to it,

So where do you think cell therapy is going?

And obviously it’s relevant to each

And obviously it’s relevant to each

So where do you think five years from now

So where do you think five years from now

Well, we talk

a lot about that with respect

to transplant that we were just
NOTE Confidence: 0.80907613
00:57:19.299 --> 00:57:21.177 talking out at BMT rounds today.
NOTE Confidence: 0.80907613
00:57:21.180 --> 00:57:23.178 About whether we’ll be doing autologous
NOTE Confidence: 0.80907613
00:57:23.178 --> 00:57:24.930 transplants in the future or not.
NOTE Confidence: 0.80907613
00:57:24.930 --> 00:57:26.835 I think that that’s kind
NOTE Confidence: 0.80907613
00:57:26.835 --> 00:57:28.359 of where we’re heading.
NOTE Confidence: 0.80907613
00:57:28.360 --> 00:57:30.320 I just wanted to say that one of
NOTE Confidence: 0.80907613
00:57:30.320 --> 00:57:31.958 the things that yell is doing,
NOTE Confidence: 0.80907613
00:57:31.960 --> 00:57:33.738 by the way, is we are developing
NOTE Confidence: 0.80907613
00:57:33.738 --> 00:57:35.574 our own party and we didn’t get
NOTE Confidence: 0.80907613
00:57:35.574 --> 00:57:37.098 a chance to talk about that.
NOTE Confidence: 0.80907613
00:57:37.100 --> 00:57:38.355 But Doctor Katz and colleagues
NOTE Confidence: 0.80907613
00:57:38.355 --> 00:57:39.920 have developed the RNA car strategy
NOTE Confidence: 0.80907613
00:57:39.920 --> 00:57:41.205 and we’re working on that.
NOTE Confidence: 0.80907613
00:57:41.210 --> 00:57:42.978 I know he has a B cell construct
NOTE Confidence: 0.80907613
00:57:42.978 --> 00:57:44.809 as well as one vertice Oklahoma,
NOTE Confidence: 0.80907613
96
so I think the whole car world is undergoing evolution as well.

And hopefully we’re going to be at the forefront of some of that.

Don’t there’s a question from the audience will then from Doctor Chi with a new drug substitute of chemotherapy. In treatments.

Also, with the new drug substitute chemotherapy treatment. From Doctor Chi.

Yeah, I mean, I think that’s a question to everybody.

Can we get rid of chemo? This toxic stuff?

We are that we added Temple where we were able to do that,
like I acute firestick leukemias.

I mean, that’s that’s a poster child, I know.

But potentially yes, that can be.

That can be the one of the goal we have.

We should not basically forget that

chemotherapy is AB side effects,

but targeted therapies have side effects too,

Situation where we are really,

completely disparaging.

Chemo too much as it’s as some real

basically benefits for for some of

the patient and for chemotherapy

as well As for cell therapy.

I think we need to see the big picture
and how we can sequence and combine these different modalities of treatment. Rather than being one again the again the other. Awesome, so I think we have reached 1:00 o’clock or one minute past 1:00 o’clock tomorrow. Francine Natalia thank you for the fantastic presentation, Charlie. Thank you for hosting us and thank you for everything on behalf of everybody in immortality. And thank you to everybody who’s attending. Yeah, phenomenal work. I just think about in my tenure what’s been going on in hematology.
Yeah, it’s really exciting.

So congratulations to all the speakers and all the investigators and staff working on this. No. OK.