

0:00:00 -> 0:00:02.54 Funding for Yale Cancer Answers
0:00:02.54 -> 0:00:05.08 is provided by Smilow Cancer
0:00:05.167 -> 0:00:07.367 Hospital and AstraZeneca.
0:00:07.37 -> 0:00:09.404 Welcome to Yale Cancer Answers with
0:00:09.404 -> 0:00:11.818 your host doctor Anees Chagpar.
0:00:11.82 -> 0:00:13.69 Yale Cancer Answers features the
0:00:13.69 -> 0:00:15.985 latest information on cancer care by
0:00:15.985 -> 0:00:17.457 welcoming oncologists and specialists
0:00:17.457 -> 0:00:19.903 who are on the forefront of the
0:00:19.903 -> 0:00:21.682 battle to fight cancer. This week,
0:00:21.682 -> 0:00:23.387 it's a conversation about the
0:00:23.387 -> 0:00:25.217 role of pathology and cancer
0:00:25.217 -> 0:00:26.705 with Doctor Angelique Levi.
0:00:26.71 -> 0:00:28.873 Dr. Levi is an associate professor of
0:00:28.873 -> 0:00:31.26 pathology at the Yale School of Medicine,
0:00:31.26 -> 0:00:33.27 where Doctor Chagpar is a
0:00:33.27 -> 0:00:34.61 professor of surgical oncology.
0:00:36 -> 0:00:37.614 Angelique, maybe we can start off
0:00:37.614 -> 0:00:39.72 by you telling us a little bit about
0:00:39.72 -> 0:00:41.39 yourself and what it is you do.
0:00:41.82 -> 0:00:44.865 So as a pathologist I am anatomic
0:00:44.865 -> 0:00:46.72 and clinical pathology trained
0:00:46.72 -> 0:00:50.196 so a PCP for short and
0:00:50.196 -> 0:00:52.048 have received fellowship training
0:00:52.048 -> 0:00:54.74 and in Cytopathology,
0:00:54.74 -> 0:00:57.9 a subspecialty of the discipline.
0:00:57.9 -> 0:01:00.81 I have also received some extra
0:01:00.81 -> 0:01:04.03 training and expertise in GU pathology,
0:01:04.03 -> 0:01:06.02 all of which I did
0:01:06.02 -> 0:01:09.54 way back at Hopkins and
0:01:09.54 -> 0:01:12.62 it's a combined program.

0:01:12.62 -> 0:01:14.5 If you do both anatomic
0:01:14.5 -> 0:01:15.628 and clinical pathology
0:01:15.63 -> 0:01:17.75 it used to be five years and now is
0:01:17.75 -> 0:01:20.394 4 but you can do one of the two
0:01:20.394 -> 0:01:22.77 disciplines for a little less time now.
0:01:22.77 -> 0:01:26.508 And the anatomic focuses
0:01:26.508 -> 0:01:29.544 mostly on the study of tissue
0:01:29.544 -> 0:01:31.484 and working with a microscope,
0:01:31.49 -> 0:01:33.041 fluids and cells,
0:01:33.041 -> 0:01:35.626 whereas the clinical pathology focuses
0:01:35.626 -> 0:01:38.748 a bit more on laboratory testing,
0:01:38.75 -> 0:01:41 blood tests for example.
0:01:41.27 -> 0:01:43.682 So let's dive a little bit more into that.
0:01:43.69 -> 0:01:46.712 I mean, when we think about the role
0:01:46.712 -> 0:01:49.067 of pathology and cancer automatically,
0:01:49.07 -> 0:01:51.778 our brain kind of goes to, Oh yeah,
0:01:51.778 -> 0:01:53.198 it's the pathologists who kind
0:01:53.198 -> 0:01:55.112 of look at the biopsy and tell
0:01:55.112 -> 0:01:57.07 me whether or not I have cancer.
0:01:57.07 -> 0:01:59.016 Can you flesh out a little bit
0:01:59.016 -> 0:02:01.039 more about what it is you do and
0:02:01.039 -> 0:02:02.949 how you come up with that answer?
0:02:02.95 -> 0:02:06.846 I mean everything hinges on what you say,
0:02:06.85 -> 0:02:08.51 how much pressure is that,
0:02:08.51 -> 0:02:09.842 and how do you actually come
0:02:09.842 -> 0:02:11.24 up with the correct diagnosis?
0:02:12.51 -> 0:02:14.74 It's certainly a team
0:02:14.74 -> 0:02:16.524 from the very beginning,
0:02:16.53 -> 0:02:19.49 patients will go to either a hospital
0:02:19.49 -> 0:02:22.587 or a physician office and will have
0:02:22.587 -> 0:02:25.428 a procedure done so the procedure

0:02:25.428 -> 0:02:28.572 could be either a Pap test,
0:02:28.572 -> 0:02:30.687 screening test for cervical cancer,
0:02:30.69 -> 0:02:32.574 it could be a fine needle
0:02:32.574 -> 0:02:33.83 aspiration of a breast mass,
0:02:33.83 -> 0:02:36.742 or it could be a surgical procedure
0:02:36.742 -> 0:02:39.414 in the operating room where
0:02:39.414 -> 0:02:42.597 a tumor or an organ is removed, so
0:02:42.597 -> 0:02:46.674 all of those tissues come to the lab from
0:02:46.674 -> 0:02:50.54 those scenarios and in the lab, the
0:02:50.54 -> 0:02:54.7 histology component is where that
0:02:54.7 -> 0:02:58.06 tissue is transformed into a medium
0:02:58.06 -> 0:03:03.024 where it is put onto a glass slide and
0:03:03.024 -> 0:03:06.032 that process itself is quite intense.
0:03:06.032 -> 0:03:08.708 We have pathology assistants who help
0:03:08.708 -> 0:03:11.374 in the gross examination of these
0:03:11.374 -> 0:03:14.3 tissues when they come to the lab,
0:03:14.3 -> 0:03:17.264 especially the larger ones where they
0:03:17.264 -> 0:03:21.489 may note sizes of lesions they may sample.
0:03:21.49 -> 0:03:23.058 Areas that are critical,
0:03:23.058 -> 0:03:24.744 close to margins, etc.
0:03:24.744 -> 0:03:27.264 And those sections are then
0:03:27.264 -> 0:03:30.23 submitted in cassettes and processed,
0:03:30.23 -> 0:03:33.206 in an automated lab in a way that
0:03:33.206 -> 0:03:36.66 they are sliced and stained and put
0:03:36.66 -> 0:03:39.275 on glass slides for pathologists
0:03:39.368 -> 0:03:42.98 to then review at the time of a
0:03:42.98 -> 0:03:45.35 case review and in community practice,
0:03:45.35 -> 0:03:47.387 often it is just a pathologist,
0:03:47.39 -> 0:03:50.738 but here at academic centers we
0:03:50.738 -> 0:03:52.412 have trainees, residents,
0:03:52.42 -> 0:03:54.238 who are involved in that process.

0:03:54.24 -> 0:03:57.607 We have many sets of eyes that
0:03:57.61 -> 0:03:59.782 we call preview slides and then
0:03:59.782 -> 0:04:01.874 the pathologist sits down at a
0:04:01.874 -> 0:04:03.432 microscope to sign out.
0:04:03.432 -> 0:04:05.08 And that's actually transforming as well.
0:04:05.08 -> 0:04:06.837 Soon we might say we don't sit
0:04:06.837 -> 0:04:09.099 down at a microscope to sign out,
0:04:09.1 -> 0:04:12.636 but we may sit at a computer screen
0:04:12.636 -> 0:04:16.032 if we transform into the digital era,
0:04:16.032 -> 0:04:18.18 but we're not quite there yet.
0:04:22.46 -> 0:04:26.16 Then with a microscope is where we
0:04:26.16 -> 0:04:28.22 really do what we were trained to do,
0:04:28.22 -> 0:04:31.972 and you use your trained eye to look
0:04:31.972 -> 0:04:34.856 at the morphology of the tissue and
0:04:34.856 -> 0:04:37.784 see where it differs from what you
0:04:37.784 -> 0:04:40.99 have trained yourself to know what's normal.
0:04:40.99 -> 0:04:43.186 So identifying what's abnormal
0:04:43.186 -> 0:04:46.48 disease and in that then deciding
0:04:46.572 -> 0:04:48.887 whether it's cancer or not.
0:04:48.89 -> 0:04:50.816 So not every disease is cancer,
0:04:50.82 -> 0:04:52.758 and it's important in some cases
0:04:52.76 -> 0:04:54.044 where the presumption clinically
0:04:54.044 -> 0:04:56.41 might be a mass because of cancer,
0:04:56.41 -> 0:04:58.702 it's a really important piece to
0:04:58.702 -> 0:05:01.337 be able to say this isn't cancer,
0:05:01.337 -> 0:05:03.92 and so therefore no treatment is necessary.
0:05:03.92 -> 0:05:05.68 But at a Cancer Center,
0:05:05.68 -> 0:05:10.16 many of the referrals that come here often
0:05:10.16 -> 0:05:12.415 perhaps already with a preliminary
0:05:12.415 -> 0:05:15.13 diagnosis on a small biopsy of
0:05:15.13 -> 0:05:17.434 cancer and then our job sometimes,

0:05:17.44 -> 0:05:19.948 as pathologists, in a larger procedure, or a
0:05:19.95 -> 0:05:24.36 resection is to then go ahead and stage that,
0:05:24.36 -> 0:05:26.765 which means assign some more
0:05:26.765 -> 0:05:28.689 parameters around that diagnosis.
0:05:28.69 -> 0:05:31.122 So not only is it cancer,
0:05:31.122 -> 0:05:33.866 but it's a type of cancer that
0:05:33.866 -> 0:05:36.868 you want to kind of classify.
0:05:36.87 -> 0:05:40.206 It's given a grade as we call it,
0:05:40.21 -> 0:05:42.694 well differentiated, poorly differentiated.
0:05:42.694 -> 0:05:45.799 It might be given certain
0:05:45.8 -> 0:05:49.37 other parameters regarding size or margin.
0:05:49.37 -> 0:05:50.778 Different cancers have different
0:05:50.778 -> 0:05:52.186 parameters that are important,
0:05:52.19 -> 0:05:55.928 and all of those details are important
0:05:55.928 -> 0:05:58.4 in prognosis prediction and then
0:05:58.4 -> 0:06:00.065 treatment and usually
0:06:00.065 -> 0:06:02.285 associated then with outcome.
0:06:03.43 -> 0:06:05.149 So I want to pick up on a few
0:06:05.149 -> 0:06:06.457 things that you said there.
0:06:06.46 -> 0:06:09.412 So one was this whole process
0:06:09.412 -> 0:06:11.99 that really goes on that
0:06:11.99 -> 0:06:14.654 many people who have never stepped into a
0:06:14.654 -> 0:06:17.207 pathology lab might not know about which is
0:06:17.21 -> 0:06:20.034 when you have a biopsy done
0:06:20.04 -> 0:06:23.36 and your surgeon, your radiologist,
0:06:23.36 -> 0:06:25.33 whoever has done the biopsy,
0:06:25.33 -> 0:06:27.706 sends that specimen away.
0:06:27.706 -> 0:06:29.476 Oftentimes, it's the greatest
0:06:29.476 -> 0:06:31.436 amount of patient anxiety waiting
0:06:31.436 -> 0:06:33.599 for that result to come back.
0:06:33.6 -> 0:06:36.28 And sometimes it can take a few days,

0:06:36.28 -> 0:06:38.926 but there is all of this
0:06:38.926 -> 0:06:41.289 preprocessing that needs to go on.
0:06:41.29 -> 0:06:44.878 Can you give us a sense of how long
0:06:44.878 -> 0:06:47.854 these biopsy results can sometimes take,
0:06:47.86 -> 0:06:50.537 and why it's important to really be
0:06:50.537 -> 0:06:52.499 patient and wait for your pathologist
0:06:52.499 -> 0:06:54.857 to give you the right answer because
0:06:54.857 -> 0:06:57.553 as you say so much of treatment really
0:06:57.553 -> 0:06:59.77 rests on what the pathologist says.
0:07:00.52 -> 0:07:02.232 Absolutely, that pre-analytical phase
0:07:02.232 -> 0:07:05.483 that you're talking about is a big part
0:07:05.483 -> 0:07:07.923 of our processing in the lab and
0:07:07.992 -> 0:07:10.404 that's kind of a traditional laboratory
0:07:10.404 -> 0:07:12.568 setting where you know pathologists when
0:07:12.568 -> 0:07:14.76 we talk about where do you work,
0:07:14.76 -> 0:07:16.084 you work in the lab, well no,
0:07:16.084 -> 0:07:17.708 we actually work mostly in our offices,
0:07:17.71 -> 0:07:19.09 but much of what's happening
0:07:19.09 -> 0:07:20.85 before we even see that glass slide.
0:07:29.425 -> 0:07:32.05 An Accessioner is the first person in the
0:07:32.05 -> 0:07:34.358 laboratory that basically does the
0:07:34.358 -> 0:07:38.17 patient registration that assigns that
0:07:38.17 -> 0:07:39.658 specimen a unique number.
0:07:39.658 -> 0:07:42.246 Every specimen in pathology is assigned a
0:07:42.246 -> 0:07:44.59 unique number and that's how we identify it.
0:07:45.046 -> 0:07:46.414 The patient information,
0:07:46.414 -> 0:07:48.694 clinical identifiers are then entered,
0:07:48.7 -> 0:07:51.535 and that's a really important step in
0:07:51.535 -> 0:07:53.771 terms of specimens being identified
0:07:53.771 -> 0:07:57.089 properly and assigned to the right person.
0:07:57.09 -> 0:08:00.219 That is the first thing that happens and

0:08:00.22 -> 0:08:04.532 the next step is it goes to the gross
0:08:04.532 -> 0:08:08.517 Histology bench and so for small biopsies
0:08:08.52 -> 0:08:11.256 that are cores or maybe liquid,
0:08:11.26 -> 0:08:12.756 or a pap smear,
0:08:12.756 -> 0:08:13.878 just single cells,
0:08:13.88 -> 0:08:15.76 fixation is something that
0:08:15.76 -> 0:08:17.64 doesn't take as long,
0:08:17.64 -> 0:08:20.065 so fixation is something that
0:08:20.065 -> 0:08:23.058 happens in different chemicals,
0:08:23.058 -> 0:08:26.193 alcohol and or formalin.
0:08:26.2 -> 0:08:28.606 Now when these tissues are larger,
0:08:28.61 -> 0:08:31.306 as in the case of a large tumor
0:08:31.306 -> 0:08:33.369 or resection or a large organ,
0:08:33.37 -> 0:08:36.618 that fixation process can
0:08:36.618 -> 0:08:39.01 happen over a 12 hour period.
0:08:39.01 -> 0:08:39.754 Sometimes overnight,
0:08:39.754 -> 0:08:40.87 so for example,
0:08:40.87 -> 0:08:44.206 a prostate that is removed whole
0:08:44.206 -> 0:08:47.55 or a large breast excision,
0:08:47.55 -> 0:08:49.488 those are examples of tissues that
0:08:49.488 -> 0:08:52.05 take a long time to fix in formalin.
0:08:52.05 -> 0:08:54.458 So before those sections can even be
0:08:54.458 -> 0:08:57.25 taken to embed in those paraffin blocks,
0:08:57.25 -> 0:08:59 that process has to happen,
0:08:59 -> 0:09:00.755 and it's critically important for
0:09:00.755 -> 0:09:03.099 that process to happen
0:09:03.1 -> 0:09:05.656 because these tissues need to be
0:09:05.66 -> 0:09:07.676 able to be examined in sections
0:09:07.676 -> 0:09:09.96 in a way where the margins
0:09:09.96 -> 0:09:12.005 and all of those distinctions
0:09:12.005 -> 0:09:14.05 between things that are critically

0:09:14.114 -> 0:09:15.838 important for patient care,
0:09:15.84 -> 0:09:17.855 whether the person gets radiation
0:09:17.855 -> 0:09:20.62 or not is the margin positive.
0:09:20.62 -> 0:09:22.76 Those delineations are critically
0:09:22.76 -> 0:09:25.435 dependent on that fixation step,
0:09:25.44 -> 0:09:28.072 and that step is where we really
0:09:28.072 -> 0:09:30.32 need to wait, and we can't rush it.
0:09:30.32 -> 0:09:33.239 So we have some technologies,
0:09:33.24 -> 0:09:35.18 microwave assistance and other things,
0:09:35.18 -> 0:09:38.558 but in that process there
0:09:38.558 -> 0:09:40.81 are still very manual
0:09:40.81 -> 0:09:43.492 pieces that take time and then
0:09:43.492 -> 0:09:46.898 by the time that slide comes out,
0:09:46.9 -> 0:09:48.664 if the surgery was on a Monday,
0:09:48.67 -> 0:09:51.256 that glass slide may not even
0:09:51.256 -> 0:09:54.253 come to a pathologist's desk until
0:09:54.253 -> 0:09:56.029 the following afternoon.
0:09:56.03 -> 0:09:57.735 And if that following afternoon
0:09:57.735 -> 0:09:59.899 is the first time a pathologist
0:09:59.899 -> 0:10:01.849 is looking at a cancer,
0:10:01.85 -> 0:10:04.906 whether it's a complicated case or even
0:10:06.04 -> 0:10:08.865 a standard morphologic diagnosis of,
0:10:08.87 -> 0:10:11.18 let's say, breast cancer, there are still
0:10:11.18 -> 0:10:13.668 additional tests that will have to get done,
0:10:13.67 -> 0:10:16.328 and so those tests will include
0:10:16.328 -> 0:10:18.1 immunostains and other markers
0:10:18.175 -> 0:10:20.401 that are all very important that
0:10:20.401 -> 0:10:23.067 need to be included in the report.
0:10:24.54 -> 0:10:27.699 So a lot of those those markers are
0:10:27.699 -> 0:10:30.38 things that we have to then order,
0:10:30.38 -> 0:10:32.54 and again it's another day

0:10:32.54 -> 0:10:33.836 or overnight processing,
0:10:33.84 -> 0:10:36.612 and so each of these steps
0:10:36.612 -> 0:10:39.12 requires kind of another decision
0:10:39.12 -> 0:10:41.596 and potentially another test
0:10:41.6 -> 0:10:44.51 or stain or molecular marker, for example.
0:10:45.12 -> 0:10:47.633 So important for people not
0:10:47.633 -> 0:10:49.955 to rush the pathologist because as
0:10:49.955 -> 0:10:53.08 I tell my patients,
0:10:53.08 -> 0:10:55.81 everything rests on what they say.
0:10:55.81 -> 0:10:57.81 But having said that,
0:10:57.81 -> 0:11:00.722 many people nowadays are
0:11:00.722 -> 0:11:02.634 talking about second opinions,
0:11:02.64 -> 0:11:04.232 either a second opinion
0:11:04.232 -> 0:11:05.426 from their clinician,
0:11:05.43 -> 0:11:07.548 but also getting their pathology that
0:11:07.548 -> 0:11:10.22 may have been reviewed at one institution
0:11:10.22 -> 0:11:12.21 re-reviewed at another institution.
0:11:12.21 -> 0:11:13.122 So for example,
0:11:13.122 -> 0:11:14.946 if they get a second opinion,
0:11:14.95 -> 0:11:18.968 their outside pathology is often re reviewed.
0:11:18.97 -> 0:11:21.426 So can you talk about the importance of
0:11:21.426 -> 0:11:24.348 that and how often do pathologists disagree?
0:11:24.35 -> 0:11:25.955 I mean, are these diagnoses
0:11:25.955 -> 0:11:27.99 things that are black and white?
0:11:27.99 -> 0:11:29.58 That is pretty crystal clear when
0:11:29.58 -> 0:11:31.708 you see a cancer that it's a cancer.
0:11:31.71 -> 0:11:34.188 Or are there some nuances that
0:11:34.188 -> 0:11:36.37 allow for some variability in
0:11:36.37 -> 0:11:38.17 terms of pathologic opinion?
0:11:38.41 -> 0:11:42.323 I'll start by saying second opinions within
0:11:42.323 -> 0:11:45.987 any scenario are always a good thing.

0:11:45.99 -> 0:11:48.118 I think for another set of eyes to
0:11:48.118 -> 0:11:53.82 take a look at a cancer case is
0:11:53.82 -> 0:11:57.096 always good and in the vast majority of
0:11:57.096 -> 0:12:00.136 cases a confirmation is what you'll find.
0:12:00.14 -> 0:12:03.698 The confirmation of the original diagnosis.
0:12:03.7 -> 0:12:06.526 It becomes more important in certain
0:12:06.526 -> 0:12:10.078 scenarios, so certain cancers
0:12:10.08 -> 0:12:12.672 have required subspecialty training
0:12:12.672 -> 0:12:15.912 that not all pathologists have,
0:12:15.92 -> 0:12:18.356 where you practice,
0:12:18.36 -> 0:12:19.503 where you've trained,
0:12:19.503 -> 0:12:21.789 and what you've become an expert
0:12:21.789 -> 0:12:24.483 in really does matter and standards
0:12:24.483 -> 0:12:26.723 are different for different places.
0:12:26.796 -> 0:12:28.468 In the community setting,
0:12:28.47 -> 0:12:30.205 while there's very high
0:12:30.205 -> 0:12:31.593 standards of care there,
0:12:31.6 -> 0:12:35.704 they may not always see all of the unique
0:12:35.704 -> 0:12:40.492 rare tumors that we might have in a tertiary
0:12:40.492 -> 0:12:41.66 academic center.
0:12:41.66 -> 0:12:44.6 Whereas in an academic center like Yale
0:12:44.6 -> 0:12:48.42 we would be able to kind of explain a bit
0:12:48.42 -> 0:12:51.456 more if there are nuances to a tumor.
0:12:51.46 -> 0:12:52.724 So black and white,
0:12:52.724 -> 0:12:54.066 yes cancer or not,
0:12:54.066 -> 0:12:56.118 in the vast majority of cases.
0:12:56.12 -> 0:12:58.26 But for challenging cases,
0:12:58.26 -> 0:13:01.47 I think second opinions are definitely
0:13:01.554 -> 0:13:04.926 helpful with expert review and consensus.
0:13:04.93 -> 0:13:06.826 Daily Conference is something
0:13:06.826 -> 0:13:09.67 that is part of our routine,

0:13:09.67 -> 0:13:11.33 not always in all practices.
0:13:11.64 -> 0:13:14.496 So it's important to kind of
0:13:14.496 -> 0:13:16.759 understand the nuances of pathology.
0:13:16.76 -> 0:13:18.506 We're going to pick up this
0:13:18.506 -> 0:13:20.251 conversation right after we take a
0:13:20.251 -> 0:13:21.835 short break for a medical minute.
0:13:21.84 -> 0:13:23.856 Please stay tuned to learn more about
0:13:23.856 -> 0:13:25.806 the role of pathology in cancer
0:13:25.806 -> 0:13:27.846 with my guest Dr. Angelique Levi.
0:13:28.3 -> 0:13:30.29 Support for Yale Cancer Answers
0:13:30.29 -> 0:13:32.28 comes from Smilow Cancer Hospital,
0:13:32.28 -> 0:13:33.995 where a dedicated team approach
0:13:33.995 -> 0:13:36.144 is used to diagnose liver cancer
0:13:36.144 -> 0:13:38.346 early when treatment is optimal and
0:13:38.346 -> 0:13:40.283 new, more effective treatments are
0:13:40.283 -> 0:13:42.587 being developed. To learn more visit
0:13:44.69 -> 0:13:47.934 Yalecancercenter.org/liver
0:13:47.934 -> 0:13:50.526 Over 230,000 Americans will be diagnosed with lung
0:13:50.526 -> 0:13:52.957 cancer this year and in Connecticut
0:13:52.957 -> 0:13:56.392 alone there will be over 2700 new cases.
0:13:56.392 -> 0:13:58.912 More than 85% of lung cancer
0:13:58.912 -> 0:14:01.12 diagnosis are related to smoking and
0:14:01.193 -> 0:14:03.761 quitting even after decades of use
0:14:03.761 -> 0:14:05.951 can significantly reduce your risk
0:14:05.951 -> 0:14:08.25 of developing lung cancer. Each day,
0:14:08.25 -> 0:14:10.32 patients with lung cancer are surviving
0:14:10.32 -> 0:14:12.822 thanks to increased access to advanced
0:14:12.822 -> 0:14:14.574 therapies and specialized care,
0:14:14.58 -> 0:14:16.265 new treatment options and surgical
0:14:16.265 -> 0:14:17.95 techniques are giving lung cancer
0:14:17.95 -> 0:14:19.63 survivors more hope than they

0:14:19.63 -> 0:14:20.974 have ever had before.
0:14:20.98 -> 0:14:22.896 Clinical trials are currently
0:14:22.896 -> 0:14:24.812 underway at federally designated
0:14:24.812 -> 0:14:26.29 Comprehensive cancer centers,
0:14:26.29 -> 0:14:28.502 such as the battle two trial at
0:14:28.502 -> 0:14:30.129 Yale Cancer Center and Smilow
0:14:30.129 -> 0:14:32.257 Cancer Hospital to learn if a drug
0:14:32.257 -> 0:14:34.555 or combination of drugs based on
0:14:34.555 -> 0:14:36.485 personal biomarkers can help to
0:14:36.49 -> 0:14:39.22 control non small cell lung cancer.
0:14:39.22 -> 0:14:42.21 More information is available at
0:14:42.21 -> 0:14:43.98 yalecancercenter.org. You're listening
0:14:43.98 -> 0:14:46.34 to Connecticut Public Radio.
0:14:46.84 -> 0:14:48.56 Welcome back to Yale Cancer Answers.
0:14:48.56 -> 0:14:50.387 This is doctor Anees Chagpar
0:14:50.387 -> 0:14:52.255 and I'm joined tonight by my
0:14:52.255 -> 0:14:53.619 guest doctor Angelique Levi.
0:14:53.62 -> 0:14:56.05 We're talking about the role of
0:14:56.05 -> 0:14:58.48 pathology in cancer and Angelique,
0:14:58.48 -> 0:15:00.44 you know one of the things that you
0:15:00.44 -> 0:15:01.843 mentioned before the break that I
0:15:01.843 -> 0:15:03.43 was interested in is that you said
0:15:03.43 -> 0:15:05.886 we are getting close to a
0:15:05.886 -> 0:15:08.5 time when pathologists might not be
0:15:08.5 -> 0:15:10.905 looking down the microscope anymore.
0:15:10.91 -> 0:15:13.297 They might be looking at a computer
0:15:13.297 -> 0:15:15.585 screen and that brought to mind
0:15:15.585 -> 0:15:17.53 this whole concept of digital
0:15:17.606 -> 0:15:20.201 pathology and potentially the role
0:15:20.201 -> 0:15:22.796 of artificial intelligence in helping
0:15:22.87 -> 0:15:25.25 pathologists make that diagnosis.

0:15:25.25 -> 0:15:27.35 You talked a little bit before the
0:15:27.35 -> 0:15:29.198 break about some of the nuances.
0:15:29.2 -> 0:15:31.664 Can you talk a little bit about
0:15:31.67 -> 0:15:34.856 where you see digital pathology and
0:15:34.86 -> 0:15:37.079 the role of AI kind of playing
0:15:37.079 -> 0:15:39.089 into pathology as we move forward?
0:15:39.44 -> 0:15:44.389 Absolutely, the landscape is already
0:15:44.389 -> 0:15:50.476 changing and the field is rapidly evolving.
0:15:50.48 -> 0:15:53.78 And pathologists, I think are
0:15:53.78 -> 0:15:57.868 definitely stepping up and wanting to
0:15:57.868 -> 0:16:03.378 not just join this era of digital and
0:16:03.378 -> 0:16:05.318 artificial intelligence as you say,
0:16:05.32 -> 0:16:07.685 machine learning, but hopefully
0:16:07.685 -> 0:16:11.2 also take a role in leading that
0:16:11.2 -> 0:16:13.84 charge and for pathology there are
0:16:13.84 -> 0:16:16.246 so many potential applications as
0:16:16.246 -> 0:16:19.258 with everything AI is everywhere.
0:16:19.26 -> 0:16:20.692 We don't necessarily appreciate
0:16:20.692 -> 0:16:23.22 it from our phones and our apps,
0:16:23.22 -> 0:16:26.64 or for many the interfaces we do each day,
0:16:26.64 -> 0:16:29.706 but it's a tool no different
0:16:29.706 -> 0:16:31.948 than for pathology,
0:16:31.948 -> 0:16:33.676 maybe an immunostain and
0:16:33.676 -> 0:16:35.339 molecular marker or genetic
0:16:35.34 -> 0:16:38.706 profile and how we use that tool
0:16:38.706 -> 0:16:42.135 is largely dependent on
0:16:42.14 -> 0:16:44.78 what help or guidance a particular
0:16:44.78 -> 0:16:47.654 practice might be looking for.
0:16:47.654 -> 0:16:50.51 One example of AI and pathology as
0:16:50.51 -> 0:16:53.079 you mentioned or alluded to would be,
0:16:53.08 -> 0:16:55.838 helping to make a diagnosis

0:16:55.838 -> 0:16:59.06 or grading a tumor.
0:16:59.06 -> 0:17:03.391 So an area of study that I
0:17:03.391 -> 0:17:06.076 have pursued in GU pathology
0:17:06.08 -> 0:17:07.481 and in prostate cancer
0:17:07.481 -> 0:17:10.992 this is a common application now
0:17:10.992 -> 0:17:15.097 and there are already software
0:17:15.097 -> 0:17:18.96 companies that are promoting
0:17:18.96 -> 0:17:22.596 AI programs and software that can
0:17:22.596 -> 0:17:25.742 reliably help predict grades or
0:17:25.742 -> 0:17:28.837 Gleason scoring of prostate cancer.
0:17:28.84 -> 0:17:33.57 But it's not that simple.
0:17:33.57 -> 0:17:38.085 Depending on the cancers that might be
0:17:38.09 -> 0:17:40.43 seen in a given institution,
0:17:40.43 -> 0:17:42.728 whether it's more common,
0:17:42.73 -> 0:17:46.178 lower grade, or in a tertiary care center,
0:17:46.18 -> 0:17:47.506 much more complicated,
0:17:47.506 -> 0:17:49.304 higher grade,
0:17:49.304 -> 0:17:52.47 algorithms are kind of taught to
0:17:52.47 -> 0:17:55.11 answer a specific question or grade.
0:17:55.11 -> 0:17:58.27 So if you're looking for well differentiated
0:17:58.27 -> 0:18:01.385 prostate cancer 3 + 3 Gleason score,
0:18:01.39 -> 0:18:03.688 that might be one training set,
0:18:03.69 -> 0:18:06.725 whereas if you're looking for
0:18:06.725 -> 0:18:09.153 high grade prostate cancer,
0:18:09.16 -> 0:18:11.164 that is
0:18:11.164 -> 0:18:13.168 amenable not to resection,
0:18:13.17 -> 0:18:14.433 but further treatment,
0:18:14.433 -> 0:18:16.538 that might be another training
0:18:16.538 -> 0:18:18.269 software kind of algorithm,
0:18:18.27 -> 0:18:20.502 so much depends on
0:18:20.502 -> 0:18:21.99 the question being asked,

0:18:21.99 -> 0:18:24.419 and it's not just help in grading,
0:18:24.42 -> 0:18:27.768 but it could also just be help in detection,
0:18:27.77 -> 0:18:30.63 so different programs can be
0:18:30.63 -> 0:18:34.22 taught how to do different tasks,
0:18:34.22 -> 0:18:36.518 and another program might be in
0:18:36.518 -> 0:18:38.87 a better setting for community
0:18:38.87 -> 0:18:41.696 practice not to miss cancer
0:18:41.696 -> 0:18:43.896 as much as
0:18:43.896 -> 0:18:46.647 focusing on the grade because
0:18:46.65 -> 0:18:48.898 detection and preventing false
0:18:48.898 -> 0:18:52.27 negatives would really be the key
0:18:52.878 -> 0:18:54.398 perhaps in a Community setting
0:18:54.398 -> 0:18:56.07 with a lower cancer rate,
0:18:56.07 -> 0:18:58.92 whereas at the tertiary setting something
0:18:58.92 -> 0:19:03.05 that would be more helpful is perhaps
0:19:03.05 -> 0:19:05.918 an AI software algorithm that not
0:19:05.918 -> 0:19:08.639 just helps with detection or grade,
0:19:08.64 -> 0:19:10.216 but maybe with prognosis.
0:19:10.216 -> 0:19:12.991 And that's really the key,
0:19:12.991 -> 0:19:15.639 trying to discern what this AI can help
0:19:15.639 -> 0:19:18.158 with and how we'd like to apply it,
0:19:21.01 -> 0:19:22.459 tailoring the solution to the problem.
0:19:22.46 -> 0:19:25.886 But one of the questions is this.
0:19:25.886 -> 0:19:28.184 Are these technologies in use now?
0:19:28.19 -> 0:19:30.71 And is there a way for patients to
0:19:30.71 -> 0:19:32.742 know whether a particular pathology
0:19:32.742 -> 0:19:35.262 department is using that or not?
0:19:35.27 -> 0:19:36.155 For example,
0:19:36.155 -> 0:19:39.242 if I just had a biopsy at my
0:19:39.242 -> 0:19:41.454 Community Hospital and I want to make
0:19:41.528 -> 0:19:44.055 sure that they didn't miss a cancer,

0:19:44.06 -> 0:19:47.75 should I expect that they have that kind of
0:19:47.75 -> 0:19:50.598 technology that can help the pathologists?
0:19:50.6 -> 0:19:52.35 And if I'm not sure,
0:19:52.35 -> 0:19:53.519 is there a way to find out?
0:19:53.75 -> 0:19:55.864 There's always a way to find
0:19:55.864 -> 0:19:57.886 out and certainly just calling that
0:19:57.886 -> 0:19:59.992 pathologist on the bottom of the
0:19:59.992 -> 0:20:01.89 report would be the first step.
0:20:04.64 -> 0:20:07.028 Or wherever those procedures are done would
0:20:07.028 -> 0:20:09.35 certainly know within the department,
0:20:09.35 -> 0:20:12.519 I would say we're still on the cusp.
0:20:18.41 -> 0:20:21.834 I think right now in tertiary care centers,
0:20:21.84 -> 0:20:25.256 there are many kinds of testing and
0:20:25.256 -> 0:20:27.496 research scenarios and these
0:20:27.496 -> 0:20:30.392 are all kind of sprouting
0:20:30.392 -> 0:20:33.46 up now and it's not to
0:20:33.46 -> 0:20:35.8 be expected I would say because
0:20:35.8 -> 0:20:39.226 it requires so much investment and
0:20:39.226 -> 0:20:43.038 infrastructure.
0:20:46.523 -> 0:20:50.814 Whether it's cloud based
0:20:50.814 -> 0:20:54.52 memory or machine or human time,
0:20:54.52 -> 0:20:57.861 we can't expect that to all be there.
0:20:57.861 -> 0:21:01.298 Now I would say you know,
0:21:01.298 -> 0:21:03.496 in the future 5 to 10 years,
0:21:03.5 -> 0:21:05.198 10 to 15 years, I think
0:21:05.2 -> 0:21:07.208 then we can start
0:21:07.208 -> 0:21:09.89 to see where these
0:21:09.89 -> 0:21:12.23 applications are best suited,
0:21:12.23 -> 0:21:16.579 and imagine with all of this investment
0:21:16.579 -> 0:21:21.34 it would probably be helpful as a QC measure.
0:21:21.34 -> 0:21:24.322 You know there are always reimbursement

0:21:24.322 -> 0:21:26.848 codes for things that are additive,
0:21:26.848 -> 0:21:28.758 whether it's a stain or
0:21:28.758 -> 0:21:30.279 whether it's AI assisted.
0:21:30.28 -> 0:21:33.4 So I imagine in the future it would also be
0:21:33.481 -> 0:21:36.601 part of a report and so you know we're not
0:21:36.61 -> 0:21:40.01 there yet, but it does take a lot of time,
0:21:40.01 -> 0:21:40.462 infrastructure,
0:21:40.462 -> 0:21:42.27 and money frankly,
0:21:42.27 -> 0:21:44.79 and so until those costs come
0:21:44.79 -> 0:21:46.85 down or those partnerships are
0:21:46.85 -> 0:21:48.682 established, things
0:21:48.682 -> 0:21:50.514 may be commercially available
0:21:53.495 -> 0:21:56.588 at a price that is affordable for you.
0:22:03.976 -> 0:22:07.649 The other thing that is here now
0:22:07.65 -> 0:22:10.226 more and more in the cancer world is
0:22:10.226 -> 0:22:12.306 this whole concept of personalized
0:22:12.306 -> 0:22:15.024 medicine and so many clinicians are
0:22:15.024 -> 0:22:17.609 really now trying to unlock and
0:22:17.609 -> 0:22:19.669 understand the genomics of cancers.
0:22:19.67 -> 0:22:22.316 And we've certainly had guests on this
0:22:22.316 -> 0:22:24.924 show who talk about doing stains that
0:22:24.924 -> 0:22:27.75 look at a number of different
0:22:27.75 -> 0:22:31.155 genetic and genomic mutations that
0:22:31.155 -> 0:22:34.195 actually help in figuring out how
0:22:34.195 -> 0:22:37.4 a particular tumor may be treated.
0:22:37.4 -> 0:22:40.48 Is that done at the local pathology lab?
0:22:40.48 -> 0:22:43.128 What's the role of the pathologist in that?
0:22:43.13 -> 0:22:45.986 How do you decide which of these
0:22:45.986 -> 0:22:48.138 markers really needs to be done?
0:22:48.14 -> 0:22:50.772 What's the cost and is that standard
0:22:50.772 -> 0:22:53.691 of care or is that something that

0:22:53.691 -> 0:22:55.881 patients need to really individualize?
0:22:56.54 -> 0:22:59.852 So at the local level I don't
0:22:59.852 -> 0:23:03.218 think it's necessarily standard of care.
0:23:03.22 -> 0:23:05.611 Certainly immunostains,
0:23:08.8 -> 0:23:11.824 certain markers that are common
0:23:11.824 -> 0:23:14.832 to lay folks would be,
0:23:14.832 -> 0:23:16.668 we talk about estrogen and progesterone
0:23:16.668 -> 0:23:18.02 receptors for breast cancer.
0:23:18.02 -> 0:23:19.571 For example, ER,
0:23:19.571 -> 0:23:22.156 PR and certain molecular markers.
0:23:22.16 -> 0:23:24.9 I think in Community practice
0:23:24.9 -> 0:23:27.092 the idea is
0:23:27.1 -> 0:23:29.112 to partner often with
0:23:29.112 -> 0:23:30.976 another lab.
0:23:30.976 -> 0:23:33.116 Whether it's a tertiary center,
0:23:33.12 -> 0:23:36.054 a commercial lab that offers those
0:23:36.054 -> 0:23:39.177 tests because they are not able to
0:23:39.177 -> 0:23:42.3 have access to all of that in house,
0:23:42.3 -> 0:23:45.324 and so a lab like
0:23:45.324 -> 0:23:47.24 ours comes into play,
0:23:47.24 -> 0:23:50.194 where if we have something to offer,
0:23:50.2 -> 0:23:53.595 we can partner with other network hospitals,
0:23:53.6 -> 0:23:54.52 community hospitals,
0:23:54.52 -> 0:23:57.74 even other labs that might not have
0:23:57.74 -> 0:24:00.132 the same volume we do in a Cancer
0:24:00.132 -> 0:24:02.884 Center to provide all of these highly
0:24:02.884 -> 0:24:05.457 specialized tests that without a certain
0:24:05.457 -> 0:24:07.959 volume it's not affordable to run.
0:24:07.96 -> 0:24:12.064 So I think the same thing holds for
0:24:12.07 -> 0:24:14.566 additional molecular assays.
0:24:14.566 -> 0:24:19.558 Panel genetic profiling those are highly

0:24:19.558 -> 0:24:23.154 specialized areas and fields that
0:24:23.154 -> 0:24:25.739 without partnering with another kind
0:24:25.739 -> 0:24:29.01 of tertiary care center or larger
0:24:29.01 -> 0:24:31.735 lab specifically geared towards that,
0:24:31.74 -> 0:24:33.792 I think it's not
0:24:33.792 -> 0:24:35.71 expected at the local level.
0:24:36.06 -> 0:24:38.562 So are the decisions about what
0:24:38.562 -> 0:24:40.9 additional tests need to be done,
0:24:40.9 -> 0:24:42.112 so additional molecular
0:24:42.112 -> 0:24:44.77 tests and so on, EGFR VEGF,
0:24:44.77 -> 0:24:46.33 various mutational panels
0:24:46.33 -> 0:24:49.6 and so on are those decisions
0:24:49.6 -> 0:24:53.05 made by the pathologist, by the
0:24:53.05 -> 0:24:55.619 treating clinician, by a group?
0:24:55.62 -> 0:24:57.268 How are those decided?
0:24:57.93 -> 0:25:00.762 I think in the Community level
0:25:00.762 -> 0:25:03.517 the oncologist drives a lot of that because
0:25:03.517 -> 0:25:06.06 the oncologist sees on that leading edge
0:25:06.06 -> 0:25:09.756 what the potential drugs that are
0:25:09.756 -> 0:25:13.631 available that are targeted to a
0:25:13.631 -> 0:25:16.746 particular molecular change, and so in
0:25:16.746 -> 0:25:19.42 the Community setting,
0:25:19.42 -> 0:25:21.85 I think the oncologist takes that
0:25:21.85 -> 0:25:25.288 role more so in asking a pathologist,
0:25:25.29 -> 0:25:28.755 hey, there's a new drug and it targets
0:25:28.76 -> 0:25:29.657 this molecular marker.
0:25:29.657 -> 0:25:32.03 Is that something you do in your lab?
0:25:33.6 -> 0:25:35.376 Or is it something we can send out for?
0:25:35.38 -> 0:25:36.8 And then you know,
0:25:36.8 -> 0:25:38.22 the pathologist facilitates that.
0:25:38.22 -> 0:25:40.033 And so that I think happens

0:25:40.033 -> 0:25:41.759 more on the Community side,
0:25:41.76 -> 0:25:44.736 whereas I think in the tertiary care setting,
0:25:44.74 -> 0:25:45.412 like here,
0:25:45.412 -> 0:25:48.28 I think it is a bit more of a
0:25:48.28 -> 0:25:50.42 collaborative effort because there
0:25:50.42 -> 0:25:53.63 are there are the pathologists
0:25:53.712 -> 0:25:56.246 here who are doing those genetic tests
0:25:56.246 -> 0:25:59.399 and so we also have our tumor boards
0:25:59.399 -> 0:26:01.376 that while they have been outside
0:26:01.376 -> 0:26:03.14 at the Community level as well,
0:26:03.14 -> 0:26:04.976 I think in a Cancer Center,
0:26:04.98 -> 0:26:06.82 the tumor boards really are
0:26:06.82 -> 0:26:08.66 putting everyone at the table.
0:26:08.66 -> 0:26:11.54 Who has that subspecialty expertise?
0:26:11.54 -> 0:26:13.38 And so I think it's a bit more
0:26:13.38 -> 0:26:15.299 of a collaborative effort.
0:26:15.3 -> 0:26:18.485 And if there's something that is
0:26:18.485 -> 0:26:20.855 clinically warranted or a new drug,
0:26:20.86 -> 0:26:24.885 I think the pathologists here in a
0:26:24.885 -> 0:26:28.32 tertiary center are able to create
0:26:28.32 -> 0:26:30.396 these answers to some of
0:26:30.396 -> 0:26:32.14 those questions or research them,
0:26:32.14 -> 0:26:35.076 or they're already a line of research here
0:26:35.08 -> 0:26:39.385 in the department or collectively.
0:26:40.73 -> 0:26:43.51 Which segues nicely into you,
0:26:43.51 -> 0:26:44.908 know, one of my last questions,
0:26:44.91 -> 0:26:47.806 which is what are the exciting areas of
0:26:47.806 -> 0:26:50.309 research in pathology and cancer?
0:26:50.31 -> 0:26:52.89 I mean, it seems like pathology
0:26:52.89 -> 0:26:55.688 is so central to what we do.
0:26:55.69 -> 0:26:57.735 Are there some exciting developments

0:26:57.735 -> 0:27:00.389 that you see coming down the Pike
0:27:00.39 -> 0:27:02.616 in terms of pathology and cancer?
0:27:03.29 -> 0:27:06.398 Well, I definitely think the
0:27:06.398 -> 0:27:08.47 digital pathology component and
0:27:08.558 -> 0:27:10.898 the artificial intelligence piece
0:27:10.9 -> 0:27:12.42 is very exciting.
0:27:13.94 -> 0:27:15.94 It's entirely a new platform
0:27:15.94 -> 0:27:18.035 and revolution, so to speak.
0:27:18.035 -> 0:27:20.225 It's something that can be applied
0:27:20.225 -> 0:27:22.866 to all of the tools that we have
0:27:22.866 -> 0:27:25.199 and then it's a tool on its own.
0:27:25.2 -> 0:27:29.394 So what I mean by that is the
0:27:29.394 -> 0:27:32.786 ability to work with digital images,
0:27:32.786 -> 0:27:35.678 whether it's radiology or
0:27:35.678 -> 0:27:38.57 scanned pathology slide and
0:27:38.57 -> 0:27:44.12 with that scan slide use metrics or
0:27:44.12 -> 0:27:47.284 segmentation to detect changes
0:27:47.284 -> 0:27:50.39 that maybe even the human eye can't.
0:27:50.39 -> 0:27:52.686 And maybe it's not just about morphology,
0:27:52.69 -> 0:27:55.636 it's just a whole other level
0:27:55.636 -> 0:27:58.24 of detection
0:27:58.24 -> 0:28:00.905 in addition to our molecular
0:28:00.905 -> 0:28:03.037 assays and genetic profiles,
0:28:03.04 -> 0:28:05.912 is something that can on its own be
0:28:05.912 -> 0:28:08.8 additive and the exciting pieces when it
0:28:08.8 -> 0:28:12.316 is also its own prognostic indicator,
0:28:12.32 -> 0:28:14.852 and so we're always interested in
0:28:14.852 -> 0:28:17.14 knowing more about the meaning of
0:28:17.14 -> 0:28:19.18 the cancer and what effect
0:28:19.18 -> 0:28:21.558 that has on outcome and prognosis,
0:28:21.56 -> 0:28:23.96 and AI really has the potential

0:28:23.96 -> 0:28:28.39 to help each of these special
0:28:28.39 -> 0:28:30.124 techniques that we use and the
0:28:30.124 -> 0:28:31.81 ability to stand on its own.
0:28:32.49 -> 0:28:34.405 Doctor Angelique Levi is an
0:28:34.405 -> 0:28:35.937 associate professor of pathology
0:28:35.937 -> 0:28:38.086 at the Yale School of Medicine.
0:28:38.09 -> 0:28:39.254 If you have questions,
0:28:39.254 -> 0:28:40.418 the address is cancer
0:28:40.42 -> 0:28:43.465 Answers at yale.edu and past editions of
0:28:43.465 -> 0:28:46.532 the program are available in audio and
0:28:46.532 -> 0:28:48.112 written form at yalecancercenter.org.
0:28:48.112 -> 0:28:50.608 We hope you'll join us next week to
0:28:50.608 -> 0:28:52.511 learn more about the fight against
0:28:52.511 -> 0:28:54.046 cancer here on Connecticut Public Radio.
0:28:54.102 -> 0:28:55.562 Funding for Yale Cancer
0:28:55.562 -> 0:28:57.022 Answers is provided by Smilow
0:28:57.03 -> 0:29:00 Cancer Hospital and AstraZeneca.