

Poll Everywhere for Q&A pollev.com/unclcn

Upcoming Live Webinars learn.unclcn.org/live

For technical difficulties (919) 445-1000 unclen@unc.edu **Sound Check**



Start Time



Contact UNCLCN

Phone:

(919) 445-1000

Email:

unclcn@unc.edu Website:

unclen.org

This program co-provided with UNC Digital and Lifelong Learning

1

POLL EVERYWHERE

Join by Web



- 1 Go to PollEv.com
- 2 Enter UNCLCN
- Respond to activity

TO CLAIM CE CREDIT

Participants must attend using one of the following:

- · Zoom with the slides and video components
- · At a designated site with a site coordinator

The following do NOT qualify for CE credit:

- Joining Zoom using Phone audio only (using the Zoom Android or iPhone app is fine)
- · Watching with MediaSite

To claim CE credit

- · View 50 minutes or more
- · Fill out an evaluation and select a certificate
- · Claim credit within seven days

3

AFTER THE WEBINAL

To collect CE Credit, please wait for UNCLCN to end the Zoom video.

The Learning Portal only receives Zoom attendance data after the Zoom video has ended.



CONTINUING EDUCATION CREDITS

FREE CE Credits with Live Webinars

Only Available at the Day and Time Indicated



FREE CE Credits with Self-Paced, Online Courses

Available any Day and Time

learn.unclcn.org





Yuri Fedoriw, MD

Yuri Fedoriw, MD is a Distinguished Professor and Vice Chair for Clinical Research and Academic Affairs Co-Director, UNC Project-Malawi Cancer Program.

Since 2011, he has been involved in the UNC Project-Malawi Cancer Program, developing and supporting diagnostic services in support of the Kamuzu Central Hospital Lymphoma Study, AIDS Malignancy Consortium and NIH-funded clinical trials. He now serves as the co-Director of the UNC Project-Malawi Cancer Program, the co-Pl of the UNC-Malawi-South Africa Cancer Consortium (U54CA254564) and direct the translational lymphoma studies of samples submitted to UNC through this ongoing work. We have established weekly diagnostic telepathology conferences between Malawian pathologists and clinical teams to improve diagnostic accuracy and build regional capacity for cancer care. We support training in global cancer research through the recently awarded Malawi Cancer Outcomes Research Program (D43CA260641 Fedoriw, MPI) that aims to develop global cancer research leaders in Malawi and UNC. in Malawi and UNC.

Dr. Fedoriw's research interests focus on the immunologic and genetic mechanisms of lymphomagenesis, particularly in the setting of HIV infection. While hematologic malignancies and lymphoproliferative disorders (including Multicentric Casteman Disease) in sub-Saharan Africa arise under intrinsic and extrinsic pressures very different from those in the United States, comprehensive analyses of these diseases have not been performed. Our laboratory group uses advanced sequencing, immunophenotypic and cellular analyses to address gaps in our understanding of lymphomagenesis and tumor microenvironment in the context of HIV-associated immune dysregulation. Deciphering these tumor-host interactions is critical to better tailor treatment and improve outcomes, particularly in the era of cancer immunotherapy.

Dr. Fedoriw's clinical efforts revolve around the diagnosis and classification of benign and Dr. Heorin's Clinical efforts revoive around the diagnosis and classification of benigh and malignant hematolymphoid disorders. He served as the Director of Hematopathology at UNC from 2012-2019, and previously as the Director of the Hematopathology Fellowship Program. He has served on the NCI Lymphoma Clinical Trials Planning Group and the Hematology and Clinical Microscopy Committee for the College of American Pathologists. He is a member of the United States and Canadian Academy of Pathology Education Committee.

7

UR PRESENTER

He has been a proud member of the UNC Cancer Community and LCCC since 2008

OUR PRESENTER

9

He has been a proud member of the UNC Cancer Community and LCCC since 2008

OUR PRESENT

He is passionate about providing optimal diagnosis for cancer care

R PRESENTER

- He has been a proud member of the UNC Cancer Community and LCCC since 2008
- He is passionate about providing optimal diagnosis for cancer care
- He directed the division of Hematopathology from 2012-2019 before starting a research lab and taking over as the co-Director of the Malawi Cancer Program

11

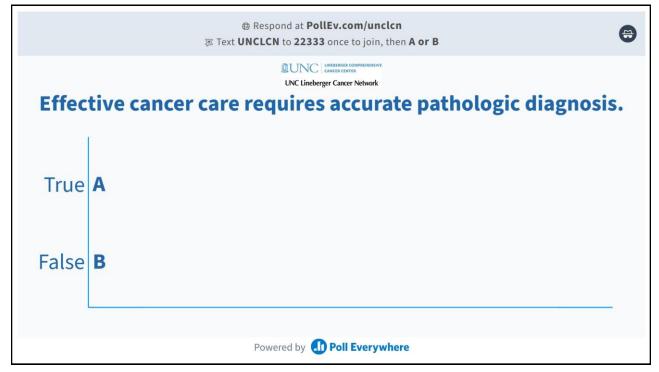
PRESENTER

- He has been a proud member of the UNC Cancer Community and LCCC since 2008
- He is passionate about providing optimal diagnosis for cancer care
- He directed the division of Hematopathology from 2012-2019 before starting a research lab and taking over as the co-Director of the Malawi Cancer Program
- He enjoys interaccting with clinical teams, trainees, staff and medical students

PRESENTER

He has been a proud member of the UNC Cancer Community and LCCC since 2008

- He is passionate about providing optimal diagnosis for cancer care
- He directed the division of Hematopathology from 2012-2019 before starting a research lab and taking over as the co-Director of the Malawi Cancer Program
- He enjoys interaccting with clinical teams, trainees, staff and medical students
- 1 Married to a UNC Family Medicine physician which adds an important primary care perspective to cancer diagnosis

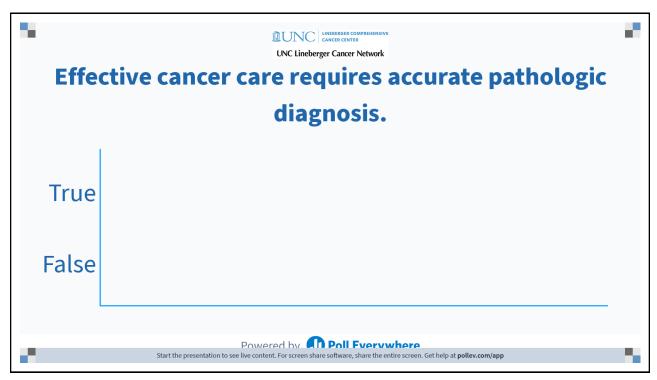


SCLOSURE

This activity has been planned and implemented under the sole supervision of the Course Director, William A. Wood, MD, MPH, in association with the UNC Office of Continuing Professional Development (CPD). The course director and CPD staff have no relevant financial relationships with ineligible companies as defined by the ACCME.

The University of North Carolina at Chapel Hill is accredited with distinction as a provider of nursing continuing professional development by the American Nurses Credentialing Center's Commission on Accreditation.

A potential conflict of interest occurs when an individual has an opportunity to affect educational content about health-care products or services of a commercial interest with which they have a financial relationship. The speakers and planners of this learning activity have not disclosed any relevant financial relationships with any commercial interests pertaining to this activity.



Cancer Pathology

Yuri Fedoriw, MD
LabCorp Distinguished Professor of Pathology & Laboratory Medicine
Lineberger Comprehensive Cancer Center

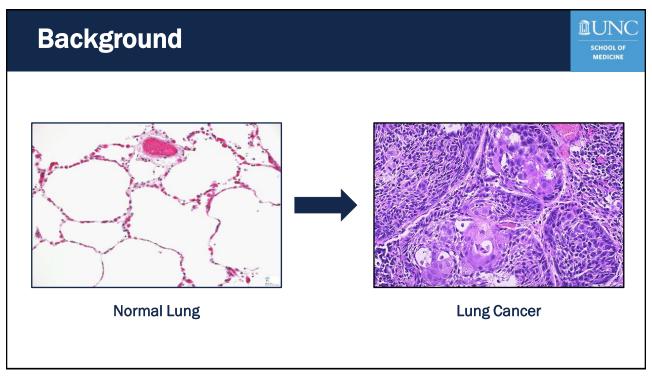


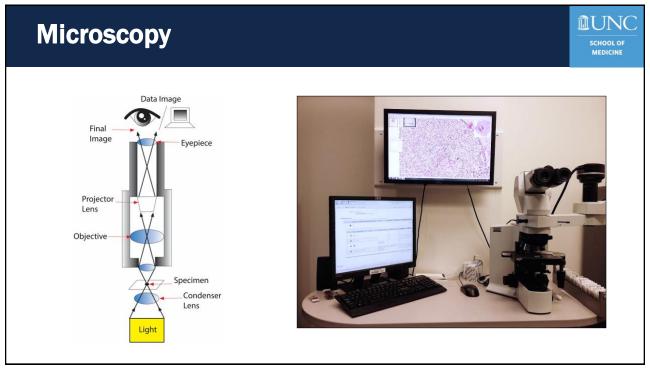
17

Objectives



- 1. Describe the evolution of testing methods for cancer diagnosis.
- 2. Recognize how classification systems are developed to uniformly diagnose tumors.
- Select appropriate tissue sampling method for cancer diagnosis.
- 4. Describe the development and application of CAP synoptic reporting for cancer diagnoses



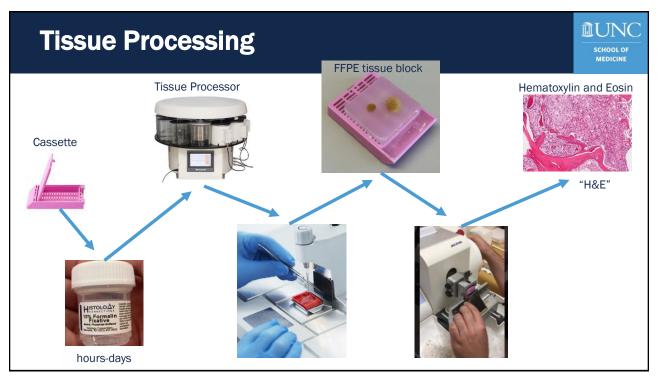


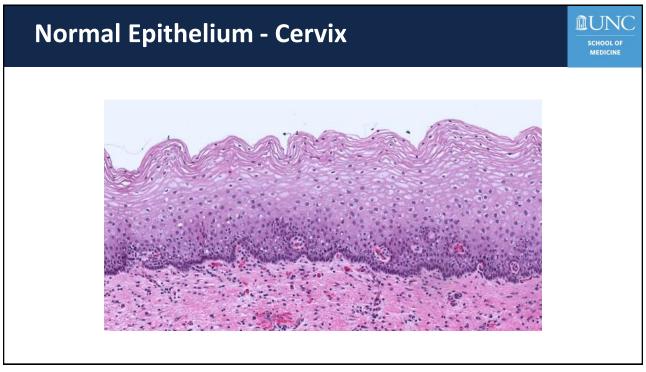
Question 1

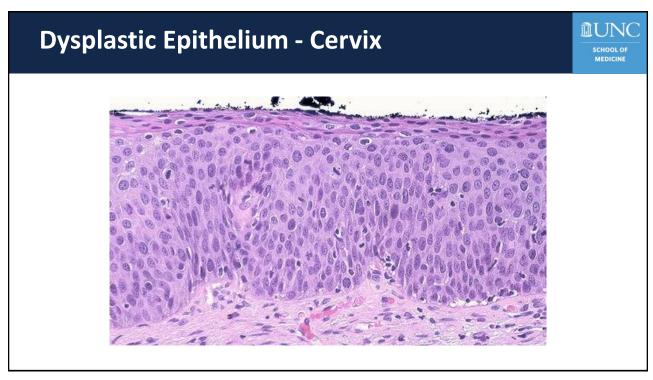


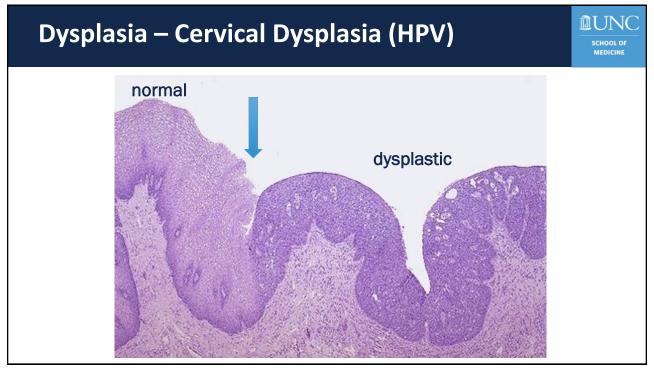
When a patient has a breast lumpectomy, when are the first slides ready to review by a pathologist?

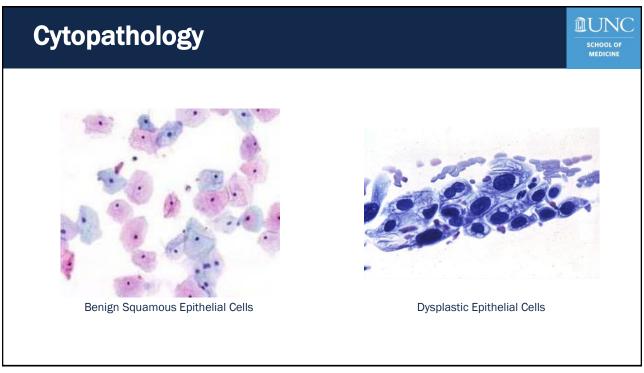




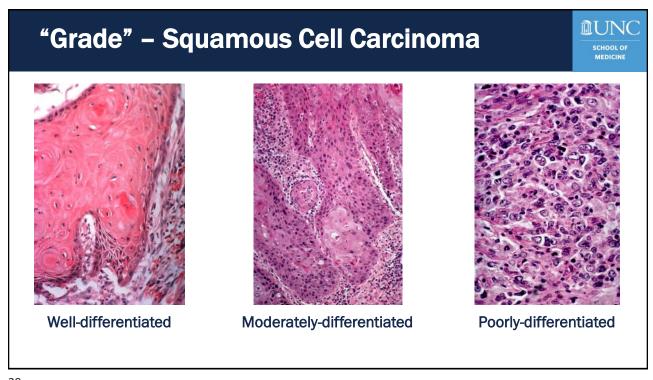


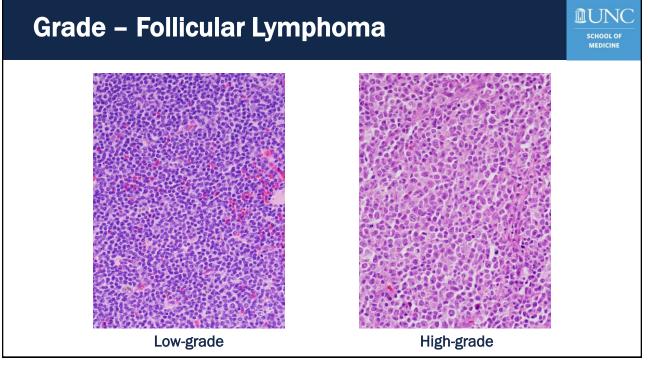








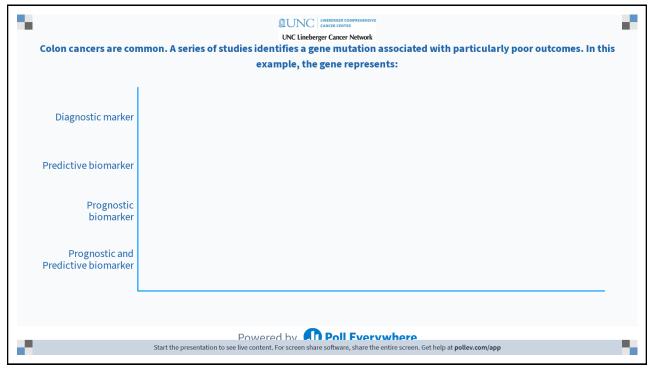




Question 2



Colon cancers are common. A series of studies identifies a gene mutation associated with particularly poor outcomes. In this example, the gene represents:



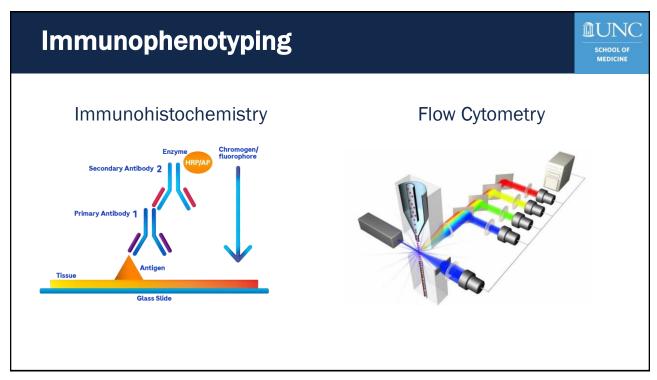
Evolution of Diagnostic Testing

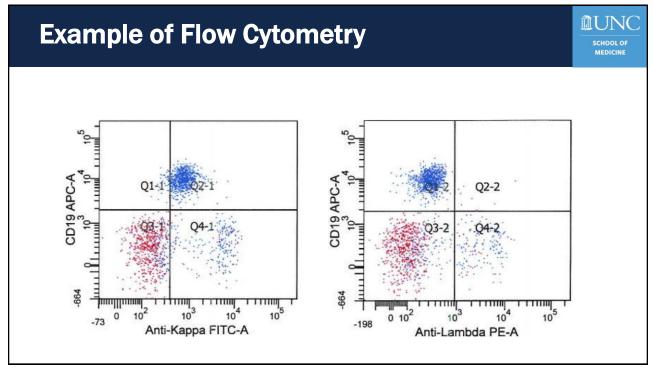


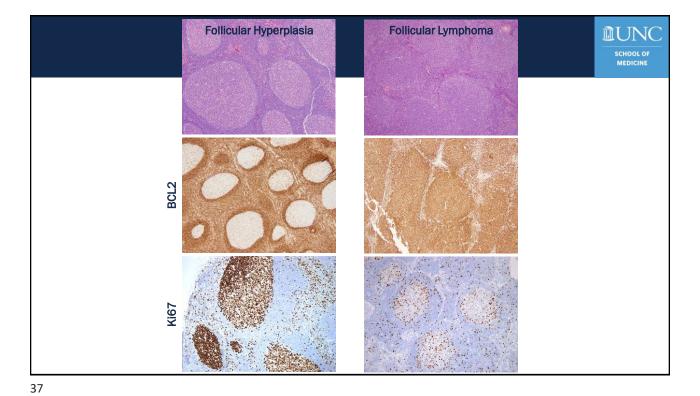
- Not all cancers that looked the same acted similarly
- Cellular and molecular biology research → mechanisms of disease and features that are:
 - Diagnostic
 - Prognostic
 - Predictive
- Technology and understanding of biology → clinical care

33

B-cell Lymphomas -MUNC from maturation and classification TdT HLA-DR CD10 CD19 CD20 CD22 CD23 CD24 CD37 CD38 CD72 CD138 * () * () * () Shaffer AL 3rd, Young RM, Staudt LM. Pathogenesis of human B cell lymphomas. Annu Rev Immunol. 2012;30:565-610.



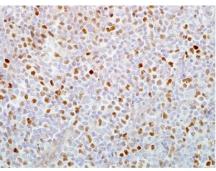




... the staining is rarely "binary"

SCHOOL OF MEDICINE

- Some stains work better than others
 - Biologic
 - Technical
- Reporting for many clinically actionable/important stains is subjective



cMYC expression:

- >40% is considered +
- What would YOU call this?

Other testing methods



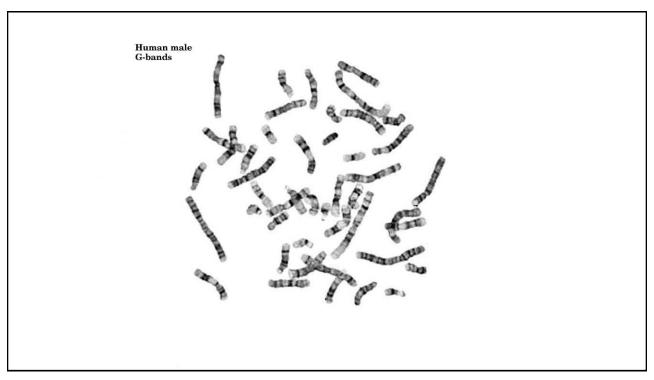
- Karyotype ("routine cytogenetics")
- Fluorescence in situ hybridization (FISH)
- Sequencing

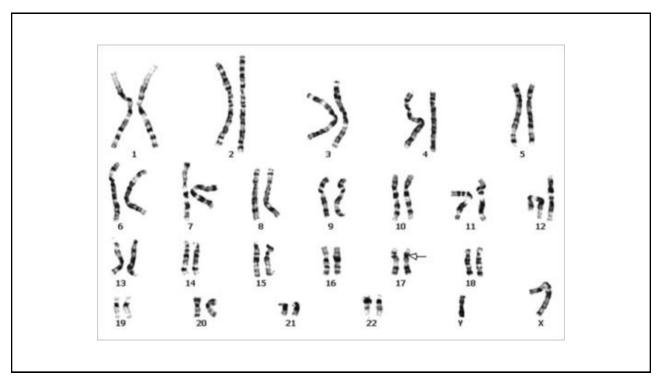
39

Karyotype



- Culture cells → arrest in metaphase → stain
- Advantages:
 - Unbiased look at the entire genome
 - Identify large aberrations
- Disadvantages:
 - Poor resolution





• Fluorescent DNA probes are hybridized to specific sequences of interest 5'— AGGATCCGTAG —3' 3'— TCCTAGGCATC —5'

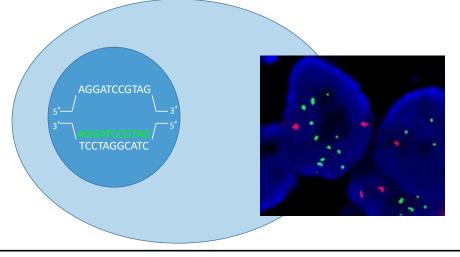
43

• Fluorescent DNA probes are hybridized to specific sequences of interest AGGATCCGTAG TCCTAGGCATC

Fluorescence in situ hybridization

SCHOOL OF MEDICINE

Fluorescent DNA probes are hybridized to specific sequences of interest

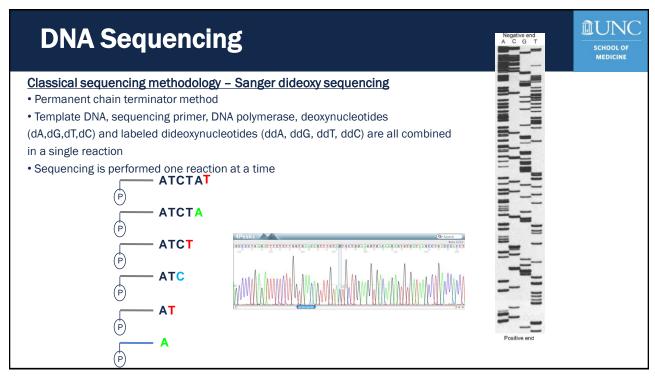


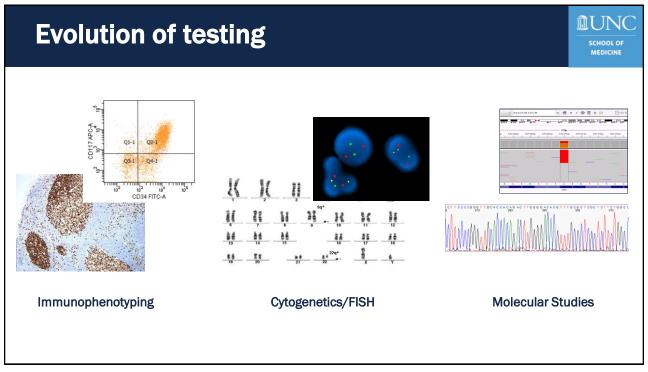
45

Fluorescence in situ hybridization



- Advantages:
 - Increased resolution compared to karyotype
- Disadvantages:
 - User defines the target

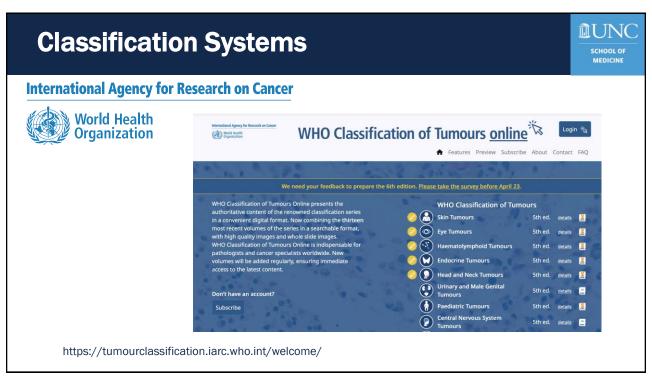


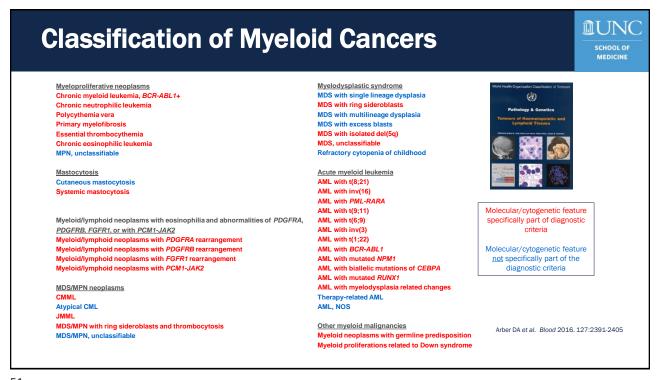


Why is Uniform Classification Important?



- Patient Care
 - Diagnosis
 - Prognostic outcome
 - Predictive predict response to therapies
- Clinical Trials
- Public Health and Policy (accurate cancer registries)





Review of Referral Cases



- Inconsistencies in diagnosis
 - Some could be true "errors"
 - Many reflect access to improved technologies, updated clinical history and imaging, new laboratory findings from presentation to referral, etc.
- Requirement for review of diagnostic material before treatment

Discordance with Non-Hodgkin Lymphoma Diagnoses



- National Comprehensive Cancer Network NHL database
- 731 patients with the 5 most common lymphoma types
- 43 (6%) were discordant from primary to NCCN center review
- 35 of 43 may have had a change in treatment!
- Depending on tumor type, discordance can be >25%

Comparison of Referring and Final Pathology for Patients With Non-Hodgkin's Lymphoma in the National Comprehensive Cancer Network

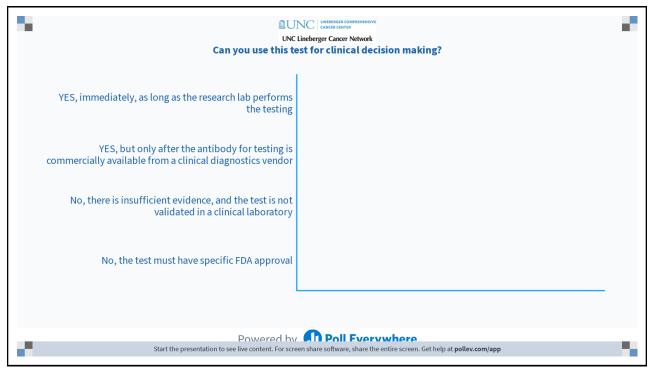
Ann S. LaCasce, Michelle E. Kho, Jonathan W. Friedberg, Joyce C. Niland, Gregory A. Abel, Maria Alma Rodriguez, Myron S. Czuczman, Michael M. Millenson, Andrew D. Zelenetz, and Jane C. Weeks Journal of Clinical Oncology 2008 26:31, 5107-5112

53

Question 3



A study of human pancreatic tumors implanted into mice identifies a new prognostic biomarker by immunohistochemistry. The findings are reported in a high-impact journal and antibody for testing is available from the research lab. Can you use this test for clinical decision making?



College of American Pathologists

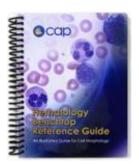


- Council on Accreditation
- Council on Education
- Council on Government and Professional Affairs
- Council on Membership and Professional Development
- Council on Scientific Affairs
 - Anatomic Pathology, Chemistry, Hematology, Informatics, Laboratory General, Molecular Pathology
- Committees of the Board of Governors

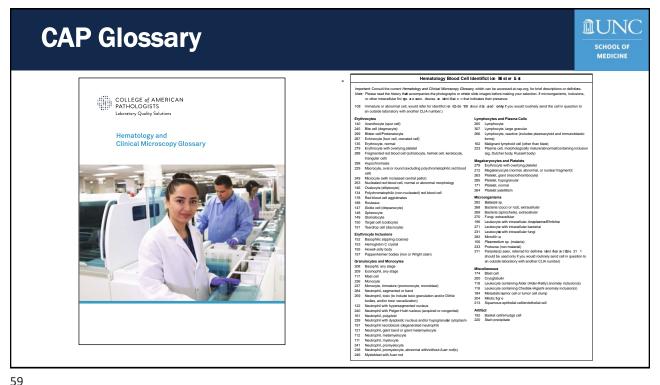
College of American Pathologists



- Primary responsibilities:
 - Survey development and review (100's/year)
 - Revisions of inspection checklist
 - Glossary upkeep



				2014				2015				2016				2017
				year				year			2016	year				year
Survey Description	2014 A	2014 B	2014 C	total	2015 A	2015 B	2015 C	total	2016 A	2016 B	С	total	2017 A	2017 B	2017 C	total
ABF1 - Automated Body Fluid, Bayer	32	32		64	36	35		71	36	35		71	41	40		81
ABF2 - Automated Body Fluid,																
Sysmex/Beckman	1337	1337		2674	1484	1483		2967	1484	1483		2967	1717	1717		3434
ABF3- Automated Body Fluid, Iris	317	317		634	339	330		669	339	330		669	348	343		691
AFL - Amniotic Fluid Leakage	662	662		1324	663	661		1324	663	661		1324	777	777		1554
APT - Fetal Hemoglobin	404	403		807	394	389		783	394	389		783	362	356		718
BCP - Blood Cell ID Photopages	192	190	190	572	188	184	183	555	188	184	183	555	238	196	193	627
BCP2 - Blood Cell ID, Limited													116	114	114	344
BCR - Bile Crystal	91	91		182	91	91		182	91	91		182	90	90		180
BFC- Body Fluid Crystals	1737	1737		3474	1820	1821		3641	1820	1821		3641	1869	1869		3738
BMD - Bone Marrow Differential	312	312		624	318	317		635	318	317		635	331	324		655
CMP- Clinical Microscopy																
Photopages CMP1- Clinical Microscopy iCHEM	5789	5840		11629	5805	5827		11632	5805	5827		11632	4538	4538		9076
								١								
Photopages CMP2- Clinical Microscopy	575	578		1153	903	997		1900	903	997		1900	1364	1393		2757
													1500	1 457 :		1.00-
Urinalysis Basic													1508	14574		16082
CMP3- Urinalysis with Clinical																
Microscopy Photopages CMMP- Clinical Microscopy Misc.													13335	1313		14648
Photopages and CD	3987	3988		7975	4129	4147		8276	4129	4147		8276	4590	4563		9153
DSC - Dipstick Confirmatory Testing EHE1 - Extended Hematology	1497	1497		2994	1374	1373		2747	1374	1373		2747	1152	1150		2302
Exercise ESR - Erythrocyte Sedimentation	236	236		472	217	215		432	217	215		432	245	244		489
Rate (ESR)	2010	2000			20.40	2020		7000	20.40	2020		7000	4010	2007		
ESR1 - ESR Sedimat 15	3810	3809		7619	3942	3938		7880	3942	3938		7880	4010	3996		8006 947
ESR2-Alifax	581 68	581 68		1162	543	541		1084 232	543	541 116		1084 232	475	472 193		386
ESR3-ALCOR	33	33	-	136 66	116 95	116 95		190	116 95	95		190	193 328	338		666
FH1 - Hematology and Differential	267	266	266	799	317	333	331	981	317	333	331	981	528 529	527	524	1580
FH1P - Hematology and Differential	20/	200	200	/77	31/	ుుు	331	701	31/	ుు	33 I	701	327	32/	324	1360
Photopages	72	72	72	216	81	86	86	253	81	86	86	253	168	163	161	492
FH2 - Hematology and Differential	185	185	185	555	178	196	195	569	178	196	195	569	450	447	443	1340
FH2P - Hematology and Differential	100	103	103	999	1/0	170	173	307	1/0	170	173	307	430	44/	440	1340
Photopages	66	66	66	198	58	56	55	169	58	56	55	169	161	163	163	487
FH3 - Hematology and Differential	97	97	97	291	108	109	109	326	108	109	109	326	107	109	109	325
FH3P - Hematology and Differential		- //	- //		.00	.57	.57	- 520	.00	.57	.57	520	.57	.07	.57	020
Photopages	315	318	318	951	303	306	302	911	303	306	302	911	228	230	229	687
FH4 - Hematology and Differential	72	68	67	207	70	81	81	232	70	81	81	232	61	59	59	179
FH4P - Hematology and Differential		50	3,							- 51	- 51		<u> </u>		/	
Photopages	318	295	299	912	262	274	274	810	262	274	274	810	200	197	197	594
FH6 - Hematology and Differential	261	261	259	781	309	311	308	928	309	311	308	928	70	60	59	189
FH6P - Hematology and Differential																
Photopages	2082	2089	2089	6260	2015	2053	2038	6106	2015	2053	2038	6106	392	325	322	1039
FH9 - Hematology and Differential	568	517	517	1602	665	676	667	2008	665	676	667	2008	761	750	742	2253



Synoptic Reporting for Cancer Diagnosis



College of American Pathologists (CAP)

- One of the leading pathology organizations
- Oversee laboratory accreditation (... more on this soon!)
- Many studies in the 90's revealed significant variation in cancer reporting → CAP cancer committee reporting checklists

https://www.cap.org/protocols-and-guidelines/cancer-reporting-tools/cancer-protocol-templates

COLLEGE of AMERICAN PATHOLOGISTS	SCHOOL OF MEDICINE
Protocol for the Examination of Specimens fro Cutaneous Squamous Cell Carcinoma of the H	
Procedure (select all that apply) Excision, ellipse Excision, wide Excision, other (specify): Re-excision, ellipse Re-excision, wide Re-excision, other (specify): Lymphadenectomy, sentinel node(s) Lymphadenectomy, regional nodes (specify): Other (specify): Not specified TUMOR Tumor Focality Unifocal Multiflocal: Cannot be determined: Multiple Primary Sites Not applicable (no additional primary site(s) present)	Histologic Type (Note B) Squamous cell carcinoma, not otherwise specified Keratoacanthoma Acantholytic squamous cell carcinoma Spindle cell squamous cell carcinoma Verrucous squamous cell carcinoma Verrucous squamous cell carcinoma Adenosquamous carcinoma Clear cell squamous cell carcinoma Squamous cell carcinoma with sarcomatoid differentiation Squamous cell carcinoma with osteoclast-like giant cells Pseudovascular squamous cell carcinoma Lymphoepithelioma-like carcinoma Other (specify): Histologic Grade (Note C) GX: Cannot be assessed G1: Well differentiated G2: Moderately differentiated G2: Moderately differentiated
Not applicable (no additional primary site(s) present) Present: Please complete a separate checklist for each primary site if required as above. Tumor Site Specify site:	G3: Poorly differentiated G4: Undifferentiated Other (specify): Not applicable
Not specified Tumor Laterality (select all that apply) Right Left Midline Not specified	Tumor Depth of Invasion (DOI) (Note D) Not applicableSpecify depth in Millimeters (mm): mmAt least (mm): mmCannot be determined (explain):

Synoptic Reporting for Cancer Diagnosis



- Checklists improve completeness
- Improved accuracy (to a point)
- For those who read them:
 - Consistent formatting
 - Columned vs. justified
 - Single-line vs. multiple lined

Synoptic Reporting: Evidence-Based Review and Future Directions

Andrew A. Renshaw, Mercy Mena-Allauca, Edwin W. Gould, and S. Joseph Sirintrapun JCO Clinical Cancer Informatics 2018:2, 1-9

Tissue Sampling for Diagnosis



- Specimen adequacy is a major challenge in clinical practice
- Limited sampling can lead to missed or delayed diagnosis
- Insufficient tissue for study enrollment or correlative science
- In response to the trend, other diagnostic methods (liquid biopsies, etc) are being developed

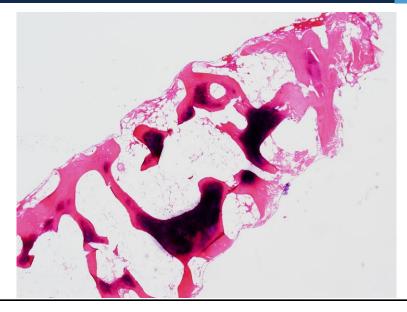
63

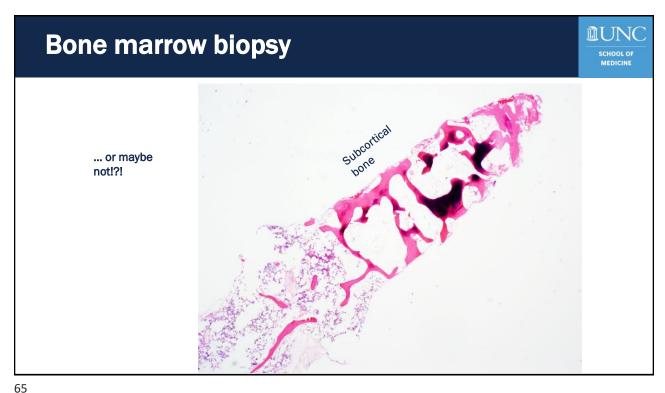
Bone marrow biopsy



Hypocellular bone marrow?

- Aplastic anemia
- Drug/toxin effect
- Hypocellular MDS
- GVHD



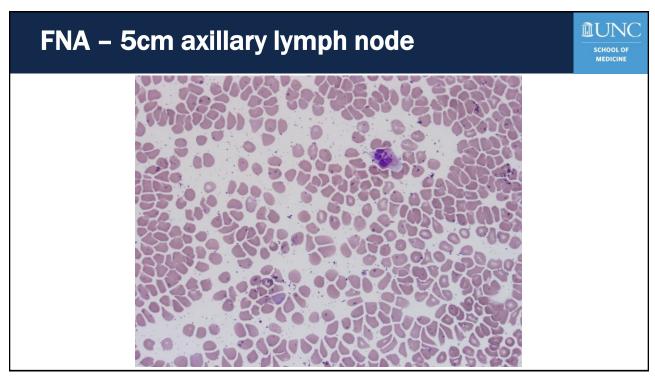


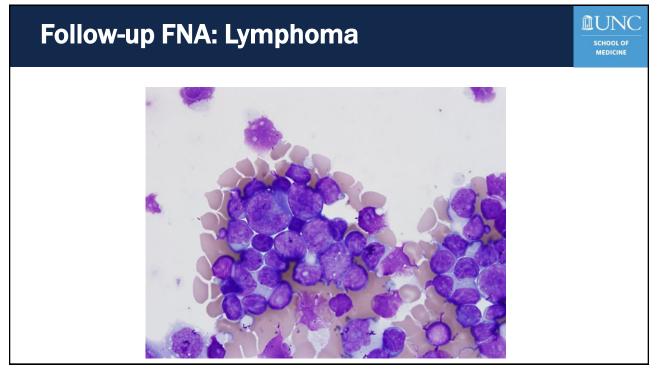
-

Fine Needle Aspirate vs Tissue Biopsy



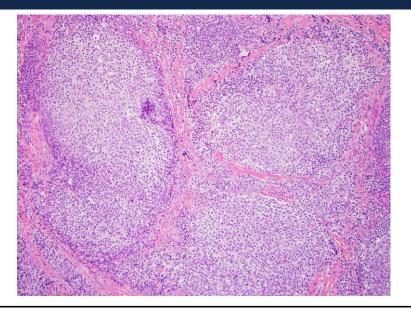
- Advantages of FNA
 - Simple procedure
 - Low morbidity
 - · Efficiently guides patient triage
- Disadvantages of FNA
 - Sampling
 - No histologic architecture
 - Frequently needs follow-up excisional biopsy





Excisional biopsy - Follicular Lymphoma





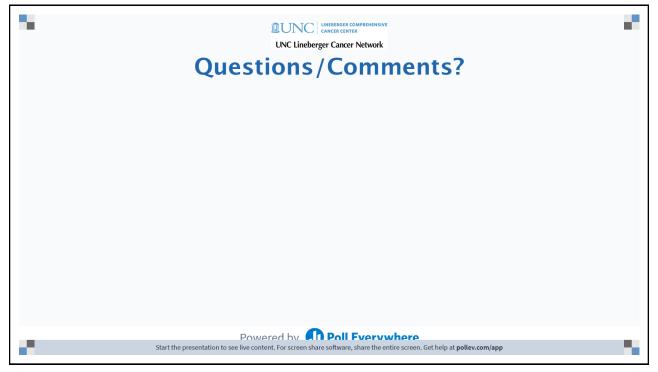
69

Conclusions



- Effective cancer care requires accurate pathologic diagnosis
- Testing methods have evolved that reflect our understanding of cancer biology and allow for improved reporting of prognostic and predictive biomarkers
- While new technologies are being developed to do 'more with less' adequate tissue biopsies are necessary (perhaps more than ever before)
- Synoptic reporting allows for discrete data elements to be provided for consistency and data collection.





University Cancer Research Fund



LINEBERGER COMPREHENSIVE CANCER CENTER

UNC Lineberger Cancer Network

The Telehealth Team

Tim Poe, Director

Veneranda Obure, Technology Support Specialist
Jon Powell, PhD, Continuing Education Specialist
Oliver Marth, Technology Support Technician

Andrew Dodgson, DPT, Continuing Education Specialist
Nadja Brown, Interim Administrative Support Specialist
Patrick Muscarella, Technology Support Technician

73

OMING LIVE WEBINARS



ADVANCED PRACTICE PROVIDER

Oncologic Emergencies: Neutropenic Fever, Cord Compression, Tumor Lysis Syndrome

Laura Blanchard, MPAP, PA-C



RESEARCH TO PRACTICE

Radiation Oncology Management of Lung Cancer in NC: Update on Small-Cell Lung Cancer Ashley Weiner, MD, PhD



PATIENT CENTERED CARE June 14 12:00 PM

May 17

4:00 PM

May 24 12:00 PM

Psychotherapy for Cancer-Related Distress Melissa Holt, DNP, PMHNP-BC, MSW Lisa Stewart, Psy.D.

Complete details on upcoming Live Webinars: learn.unclcn.org/live-webinars

SELF-PACED, ONLINE COURSES





What Is Cancer Rehabilitation and How Can it Help My Patients Sasha Knowlton, MD





Clinical Updates in Breast Oncology Emily Ray, MD, MPH





Integrating the Caregiver as a Member of the Multidisciplinary Care Team

Erin E. Kent, PhD, MSc Loretta Muss, RN, BA

Complete details on upcoming Live Webinars: learn.unclcn.org/live-webinars

75

HANK YOU FOR PARTICIPATING!

UNC Lineberger Cancer Network

Email: unclcn@unc.edu Call: (919) 445-1000

Send us an email to sign up for our monthly e-newsletter.

Check us out at unclcn.org



