

Monthly Steering  
Committee Meetings

June 29  
2022 3-4PM ET

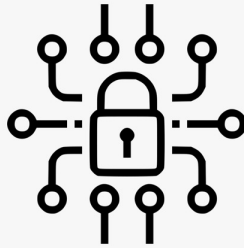
Pathology Innovation Collaborative Community



# FDA Guidance



Circulating tumor  
DNA (ctDNA)



Cybersecurity in  
medical devices



Quantitative imaging  
in radiological  
devices



# ctDNA – Out of the Dark Guidance Review Session

Plcc project aims to provide:

- A brief summary of the document
- relevant references
- And consider providing comment on the draft guidance

Join on Thursday, July 21 at 11AM ET

Comments due July 2

GUIDANCE DOCUMENT

## Use of Circulating Tumor Deoxyribonucleic Acid for Early-Stage Solid Tumor Drug Development; Draft Guidance for Industry; Availability

MAY 2022

Download the Draft Guidance Document

Draft

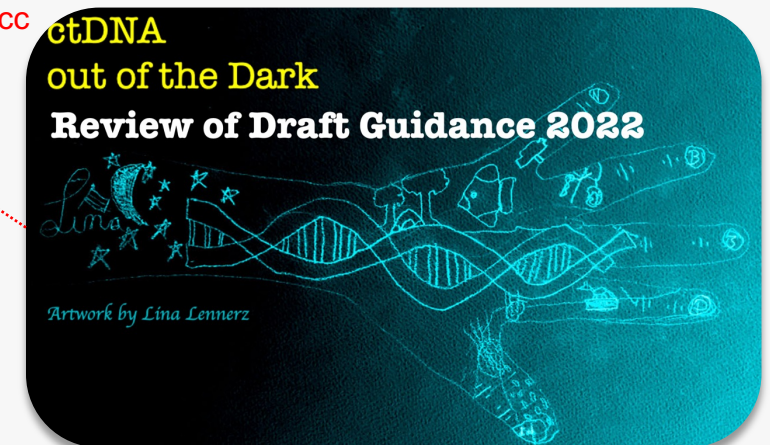
Not for implementation. Contains non-binding recommendations.

**FRIENDS**  
of CANCER  
RESEARCH

## ctMoniTR Project

- FOCR Virtual Meeting - Expediting Drug Development: Use of ctDNA as an Early Endpoint  
**Wednesday, July 20 at 11:30AM ET**

More info on Plcc project page



# Cybersecurity in medical devices

March 23, 2022

Bill: Healthcare  
Cybersecurity Act  
of 2022

April 7, 2022

FDA Document  
Details Cyber  
Expectations for  
Device Makers

May 26, 2022

Bill: Strengthening  
Cybersecurity for  
Medical Devices  
Act

GUIDANCE DOCUMENT

## Cybersecurity in Medical Devices: Quality System Considerations and Content of Premarket Submissions

*Draft Guidance for Industry and Food and Drug Administration Staff*

APRIL 2022

[Download the Draft Guidance Document](#)

[Read the Federal Register Notice](#)

Draft

Comment period open through July 7, 2022





# Cybersecurity Guidance Details

- The FDA's 49-page draft guidance covers a wide range of cybersecurity considerations and actions that medical device makers are recommended to address and document for premarket submissions, including:
- **Threat modeling**: This means identifying the security objectives, risks and vulnerabilities of a device system and then defining countermeasures to prevent, or mitigate the effects of, threats to the system throughout its life cycle.
- **Third-party software components**: This includes providing to the FDA - and to customers as part of product "labeling" - a software bill of materials, or **SBOMs**, containing information about device maker-developed components, as well as third-party purchased, licensed and open-source software.
- **Security risk management**: This includes providing a report summarizing the manufacturer's risk evaluation methods and processes and including details of the security risk assessment and risk mitigation activities undertaken, controls and the testing to ensure a device is reasonably secure.
- **Implementation of security controls**: This includes **authentication**; authorization; cryptography; code, data and execution integrity; confidentiality; event detection and logging; resiliency and recovery; updatability and patching.
- **Cybersecurity testing**: This includes security requirements, threat mitigation, vulnerability testing and penetration testing.
- **Cybersecurity transparency**: This includes transparency in labeling relevant security information for users, such as device instructions and product specifications related to cybersecurity controls, a list of network ports and other interfaces that are expected to receive and/or send data, and SBOMs that are available on a continuous basis and are machine-format readable.
- **Vulnerability management plans**: This means submitting plans for vulnerability communication so that the FDA can assess whether the manufacturer has sufficiently addressed how to maintain the safety and effectiveness of the device after marketing authorization is achieved.

# Quantitative imaging in radiological devices

- FDA Issues Final Guidance For Radiological Devices Using Quantitative Imaging
- This document is an update to FDA 2019 guidance

GUIDANCE DOCUMENT

## **Technical Performance Assessment of Quantitative Imaging in Radiological Device Premarket Submissions**

*Guidance for Industry and Food and Drug Administration Staff*

JUNE 2022

[Download the Final Guidance Document](#) [Read the Federal Register Notice](#)

Final

# European Artificial Intelligence Act (AIA)



EUROPEAN COMMISSION

Brussels, 21.4.2021

COM(2021) 206 final

2021/0106(COD)

Proposal for a

**REGULATION OF THE EUROPEAN PARLIAMENT AND OF THE COUNCIL**

**LAYING DOWN HARMONISED RULES ON ARTIFICIAL INTELLIGENCE (ARTIFICIAL INTELLIGENCE ACT) AND AMENDING CERTAIN UNION LEGISLATIVE ACTS**

{SEC(2021) 167 final} - {SWD(2021) 84 final} - {SWD(2021) 85 final}

- April 2021
- First **law on AI** by major regulator



# HHS withdrawing the **SUNSET** final rule

## Withdrawing Rule on Securing Updated and Necessary Statutory Evaluations Timely

**AGENCY:** Department of Health and Human Services.

**ACTION:** Final rule; withdrawal.

**SUMMARY:** The Department of Health and Human Services (HHS or Department) is issuing a final rule withdrawing a rule entitled “Securing Updated and Necessary Statutory Evaluations Timely” (SUNSET final rule), which published in the Federal Register of January 19, 2021. The SUNSET final rule was originally scheduled to take effect on March 22, 2021. However, after a lawsuit was filed on March 9, 2021, seeking to overturn the SUNSET final rule, HHS extended

“Had this withdrawal of the SUNSET rule not gone through, the FDA, according to HHS, would have faced the massive task of reviewing over 7,000 sections of the CFR that were promulgated by the agency and which are more than 10 years old or would become more than 10 years old during the first five years the rule would be in effect, representing over 95% of the FDA’s current regulations.” –*Endpoints News*

## Economic Impact Analyses of FDA Regulations

[f Share](#) [t Tweet](#) [in LinkedIn](#) [✉ Email](#) [🖨 Print](#)

The Food and Drug Administration conducts economic analyses of all important proposed and final regulations. Each economic analysis includes an assessment of the costs, benefits, and cost-effectiveness of the action, as well as assessments of the costs, benefits and cost-effectiveness of the most promising alternative actions. The full economic impact analyses of significant FDA regulations are no longer (as of April 2012) published in the Federal Register but are available on this site.



# AMA CPT Announces New Digital Pathology Codes



## CPT® Editorial Summary of Panel Actions May 2022

Tab #	Name	Codes	Request-Description	Effective Date
37	Post Operative Low Level Laser Therapy	●9X022	Accepted addition of code 9X022 code to report low-level laser therapy for post operative pain reduction	January 2024
38	Electroencephalogram Monitoring	-----	WITHDRAWN	
39	Human Milk Donation Services	-----	REJECTED	
40	Therapeutic Monitoring Services	-----	WITHDRAWN	
41	Vaccine Counseling	-----	WITHDRAWN	
42	Respiratory Syncytial Virus (RSV) Vaccine	●9X018	Accepted addition of code 9X018 to report RSV vaccine product and administration	January 2023
43	Cat II - Medication Adherence for Opioid Use Disorder	-----	WITHDRAWN	
44	Cat III - Digital Pathology	+●X018T +●X025T +●X019T +●X026T +●X020T +●X027T +●X021T +●X028T +●X022T +●X029T +●X023T +●X030T +●X024T	Accepted addition of add-on codes X018T-X030T to report additional clinical staff work and service requirements associated with digitizing glass microscope slides for primary diagnosis; and addition of a new heading in the Category III section and guidelines to define digital pathology digitization procedures	January 2023
45	Cat III - AI Analysis for Cardiac Function Services	+●X044T ●X045T	Accepted addition of add-on codes X044T and child code X045T to report assistive algorithmic electrocardiogram risk assessment for cardiac dysfunction	January 2023

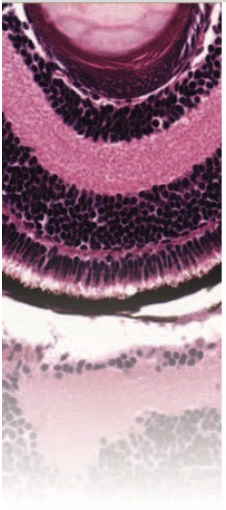






# CPT codes per January 2022

- Accepted addition of add-on codes X018T X030T to report additional **clinical staff** work and **service requirements** associated with **digitizing glass** microscope slides for primary diagnosis; and addition of a new heading in the **Category III** section and guidelines to define digital pathology digitization procedures



# What is DPA working on

- Allow labs to record activity and **begin tracking the value** of DP
- Categorization, cross walk, PC and/or TC
- Work with organizations like the **AMA and CAP** to determine the best approach to value DP solutions (including AI) and more
- DPA-F is seeking funding to focus on **outcome studies**

# CURE ID

- collaboration between FDA and the National Center for Advancing Translational Sciences (NCATS), part of the National Institutes of Health (NIH)



## What is the value of CURE ID?

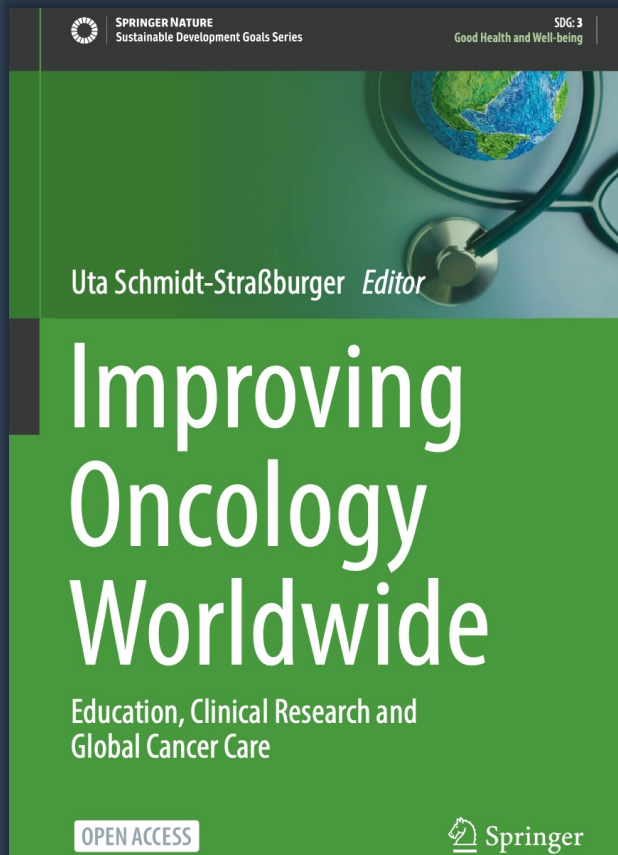
The CURE ID platform will enable collection of real-world treatment experience directly from healthcare providers in a systematic, accurate, and detailed manner. Information from CURE ID can be used to inform ongoing and future clinical trials and potentially drug labeling.

For example, it could:

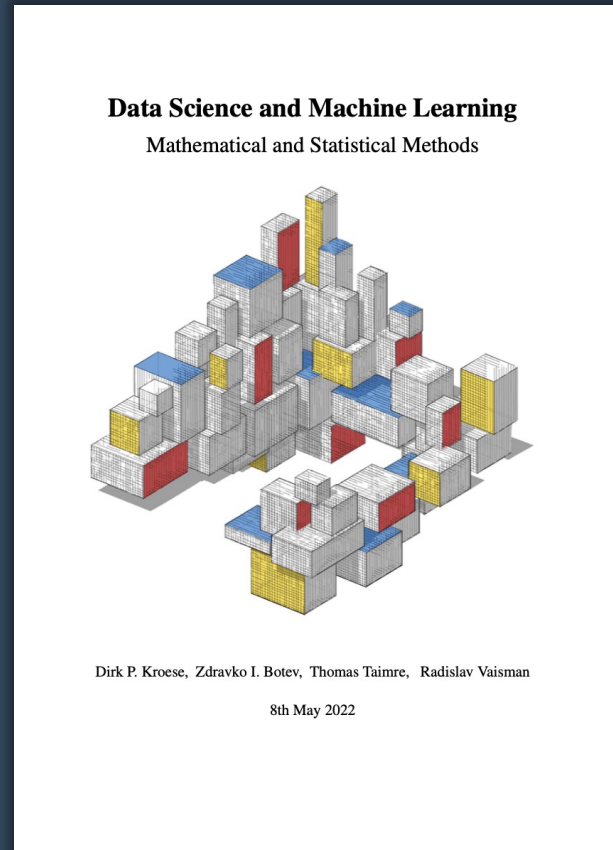
- Serve as a platform for healthcare providers to exchange their treatment experiences about challenging diseases.
- Spur clinical trial development by collecting better information regarding positive and negative health outcomes about existing products, which may help drug developers narrow potential drug candidates for study. Ultimately, this program could help enrich the armamentarium of drugs available for diseases that are difficult to treat.
- Serve as a resource for healthcare providers to share information where a disease lacks adequate approved treatment options.
- Allow for the rapid identification of promising treatment approaches in urgent situations, such as during outbreaks of emerging infectious threats and use this information to inform clinical trials.
- Enable collection of real-world treatment experience directly from healthcare providers in a systematic, accurate, and detailed manner. Information from CURE ID can be used to inform ongoing and future clinical trials and potentially drug labeling.



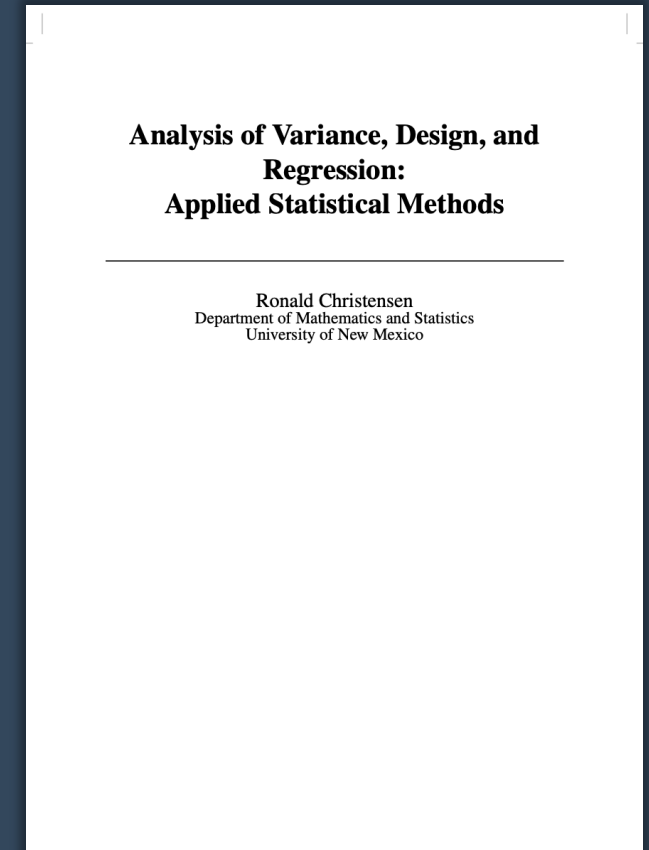
# Books of interest



**Uta Schmidt-Strasßburger**  
Improving Oncology Worldwide



**Kroese et al.**  
Data Science and Machine Learning  
Mathematical and Statistical Methods

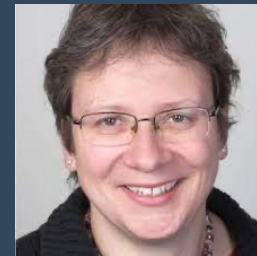


**Ronald Christensen**  
Analysis of Variance, Design, and  
Regression: Applied Statistical  
Methods





# Continuing Medical Education Program "Master Online Advanced Oncology" (part-time)



## Type of study

Continuing medical education programme/part-time (M.Sc.)

## Language

english

## Credit Points

363 CME points

60 ETCS points

## Number of students/year

20

## Start of the study programme

Winter semester (1 October)

application deadline 15 May

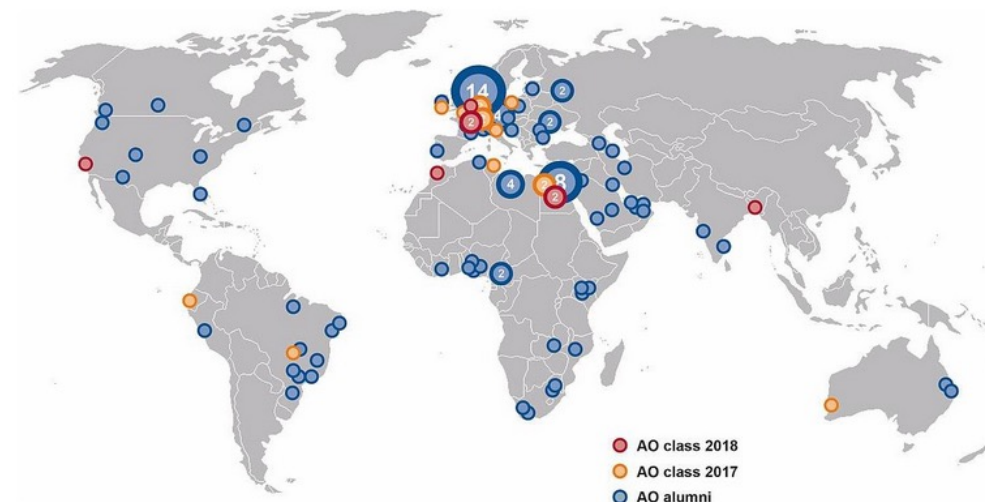
## Regular duration of the study

4 semesters



## Internationality

Our students and lecturers are from all over the world. Join the Advanced Oncology Network today and meet at high-ranking international meetings!





# History of AI



**CRITICAL FOCUS**

Brian J. Ford

featured work

## AI: Artificial, Yes. Intelligent, Not.

*Artificial intelligence and machine learning are wonderfully efficient, but they pale in comparison to the fine intelligence and complicated mechanisms of the living cell.*



Eric was one of the first autonomous robots and rose to give the opening speech at a major engineering exhibition in London in 1928. American newspapers described this robot as “an almost perfect man.” A working replica was reconstructed by the Science Museum in London in 2017.



Eric, the talking Robot, is shown above making an ora-

Press reports aroused international interest in Eric, who was designed to respond to verbal commands and could move his limbs and eyes. After being exhibited in London in 1928, the original Eric toured the world, but within a decade, he disappeared and was never seen again.



# QUAREP-LiMi

Become a member

Join the QUAREP community and become a member.

Click Here

## Quality Assessment and Reproducibility for Instruments & Images in Light Microscopy

The QUAREP-LiMi is a group of enthusiastic light microscopists from Academia and Industry all interested in

improving quality  
on-line Web  
has grown to  
communities  
standardizat  
agencies (se

### Please select the Working Group(s) you would like to join \*

- |   |   |
|---|---|
| <input type="checkbox"/> WG 1 - Illumination power                | <input type="checkbox"/> WG 2 - Detection system performance                    |
| <input type="checkbox"/> WG 3 - Uniformity of field - flatness    | <input type="checkbox"/> WG 4 - System chromatic aberration and co-registration |
| <input type="checkbox"/> WG 5 - Lateral and axial resolution      | <input type="checkbox"/> WG 6 - Stage and focus                                 |
| <input type="checkbox"/> WG 7 - Metadata                          | <input type="checkbox"/> WG 8 - White paper                                     |
| <input type="checkbox"/> WG 9 - Over all planning + funding       | <input type="checkbox"/> WG 10 - Image Quality                                  |
| <input type="checkbox"/> WG 11 - Microscopy publication standards | <input type="checkbox"/> WG 12 - Image Visualization and Analysis               |
| <input type="checkbox"/> WG 13 - Phototoxicity                    | <input type="checkbox"/> Currently not actively want to join a WG               |

featured work

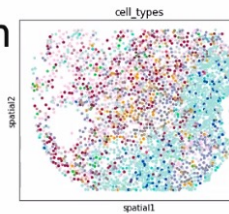
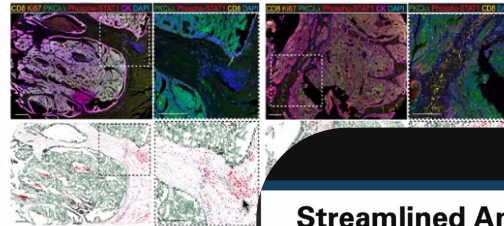
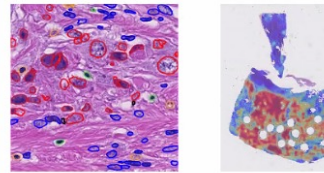




# PathML

## How PathML is Being Used Today

- Prostate cancer research, H&E images
  - Multiple instance learning
  - Cell segmentation and classification
- Colorectal cancer research, 7-channel IF images
  - Cell segmentation and rules-based phenotyping
  - Spatial biology
- Image quantification pipelines used in production at DFCI core facility, CODEX spatial proteomics
  - Improving operational efficiency of key institutional resource



## PathML: An open-source software toolkit for computational pathology research



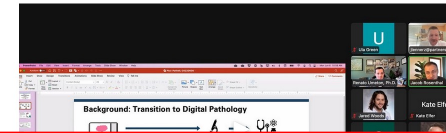
### Summary:

Imaging datasets in cancer research have grown exponentially in size and information density in recent years, driven chiefly by two trends:

- Increasing adoption of digital pathology workflows at departmental and institutional scale (large  $n$  datasets)
- Emerging technologies in highly multiplexed imaging and spatial omics (high dimensional datasets)

The unprecedented scale of today's datasets may enable derivation of insights for cancer research and clinical care, but only if researchers are equipped with the tools to leverage advanced computational approaches from machine learning and computer vision. PathML is a software toolkit designed to lower the barrier to entry for computational pathology, enabling researchers to develop streamlined, scalable, fully customized end-to-end image analysis pipelines, with a unified framework for brightfield, multiplexed immunofluorescence, and spatial omics images and support for 160+ file formats. Developed at Dana-Farber Cancer Institute and Weill Cornell Medicine, PathML is currently being used by 7+ research groups and 2 imaging core facilities across the two institutions. PathML is an open-source project freely available on GitHub, with complete documentation, tutorials, and example vignettes and more than 15,000 downloads worldwide. We welcome anyone interested in collaborating or learning more to contact us at [PathML@dfci.harvard.edu](mailto:PathML@dfci.harvard.edu) or visit [www.pathml.org](http://www.pathml.org) for more information.

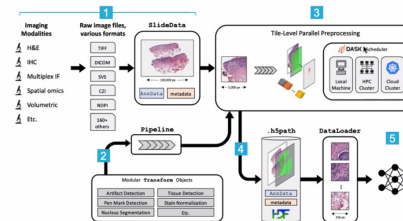
### Presentation recording:



June 6<sup>th</sup> presentation  
Available on website

## Streamlined Analysis Workflows

- Complete end-to-end pipelines in ~10 lines of code
- Lower barrier to entry for image analysis research
- Enables rapid prototyping/development
- Built-in support for HPC and cloud clusters



```

1 # Load the Image
slide = CODEXSLIDE("path/to/image_001.tif")

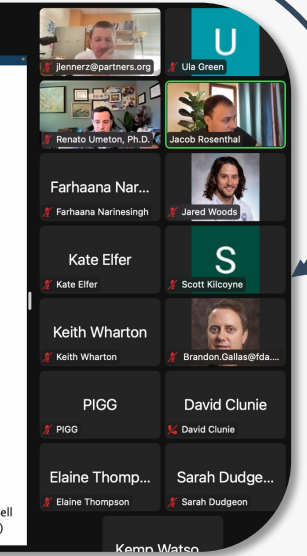
# Define a pipeline
pipe = Pipeline[
  ColLapsRuns(CODEXz = 0),
  SegmentMF(model = "mesher",
             nuclear_channel = 0,
             cytoplasm_channel = 11,
             image_resolution = 0.5),
  QuantifyMF("cell_segmentation")
]

# Run pipeline with distributed computing
slide.run(pipe, title_size = 1024)

# Save output
slide.write("path/to/image_h5path")

# PyTorch DataLoader
dataset = TILoader("path/to/image_h5path")
data_loader = DataLoader(dataset, batch_size = 16)
    
```

Full code for analysis of mIF images (including ML-powered cell segmentation, marker quantification, PyTorch DataLoader)



featured work



# MedTech Color Collaborative Community Annual Report

featured work

1



MedTech Color Collaborative Community Annual Report

## MedTech Color Collaborative Community Annual Report

### Collaborative Community Overview

MedTech Color has convened this [Collaborative Community](#) to create a forum in which a diverse cross-section of MedTech industry stakeholders can work together to identify issues and opportunities for improvement of current practices, share and curate existing knowledge, and develop evidence-based solutions to address racial and ethnic minority health issues in medical device product development and clinical research. The Collaborative Community will work to develop targeted strategies, including education and awareness activities, guidelines, best practices, and thought leadership to increase the awareness, understanding, access, engagement, and participation of racial and ethnic minorities.

Our community is comprised of volunteers united by our common drive, passion, and dedication for creating more diverse and representative populations in clinical research. The potential to support better patient outcomes by promoting greater diversity in medical device development fuels our mission.

Thank you to the MedTech Color  
Collaborative Community Sponsors





# American Board of Pathology Annual Report

featured work

2021

## Annual Report

**American  
Board of  
Pathology**

One Urban Centre, Suite 690  
4830 West Kennedy Boulevard  
Tampa, Florida 33609  
<https://www.abpath.org>



## 2021 Subspecialty Examinations

	Total Candidates		First-Time Takers			Repeaters		
	#	% Pass	#	# Pass	% Pass	#	# Pass	% Pass
<b>BB/TM</b>	48	81	45	38	84	3	1	33
<b>CH</b>	-	-	-	-	-	-	-	-
<b>CI</b>	35	77	30	25	83	5	2	40
<b>CYP</b>	124	86	117	103	88	7	4	57
<b>DP</b>	45	87	43	38	88	2	1	50
<b>FP</b>	38	95	33	31	94	5	5	100
<b>HEM</b>	112	94	108	103	95	4	2	50
<b>MMB</b>	18	100	17	17	100	1	1	100
<b>MGP</b>	52	94	52	49	94	-	-	-
<b>NP</b>	31	90	31	28	90	-	-	-
<b>PP</b>	22	91	21	20	95	1	0	0
<b>FPclinMB*</b>	-	-	-	-	-	-	-	-

\*FPclinMB = Focus Practice Designation-Clinical Microbiology



# The Commonwealth Fund Meeting America's Public Health Challenge

featured work



## MEETING AMERICA'S PUBLIC HEALTH CHALLENGE

Recommendations for Building a National Public Health System That Addresses Ongoing and Future Health Crises, Advances Equity, and Earns Trust

The Commonwealth Fund Commission on a National Public Health System

JUNE 2022



### Congress should:

- Establish a position, such as an undersecretary for public health at the U.S. Department of Health and Human Services (HHS), to oversee and coordinate the development of the national public health system.
- Provide adequate and reliable public health infrastructure funding, paired with expectations that states, localities, tribes, and territories meet standards for protecting their communities.
- Fund a modern public health information technology system and provide HHS with the authority to make it work.

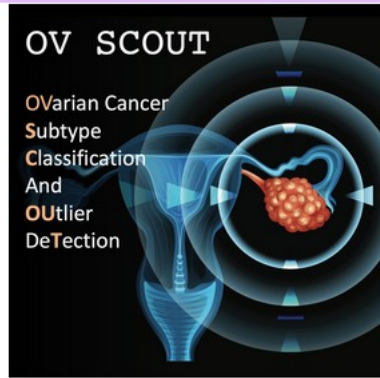
### The Administration should:

- Set parameters for use of available funds to systematically build public health infrastructure, with an initial focus on workforce and data systems.
- Support revision of accreditation standards for state, local, tribal, and territorial health departments to focus on basic capabilities for public health protection.
- Establish a council to coordinate federal public health action with states, localities, tribes, and territories.
- Reconvene the National Prevention and Public Health Council to guide an all-of-government approach to the drivers of health.
- Embrace ethics, integrity, and transparency in decision-making in public health.

### States, localities, tribes, and territories should:



# Other project updates pushed to July



5/25/22

## OV SCOUT

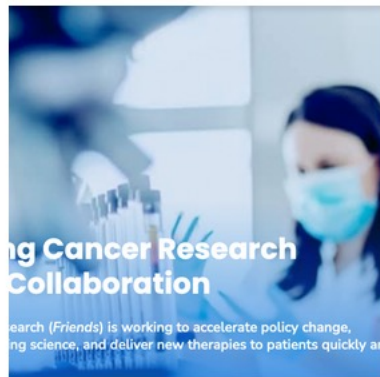
[Read More](#)



5/24/22

## CPIM: WSI for nonclinical development

[Read More](#)



5/24/22

## Advancing Cancer Research Through Collaboration

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5/19/22

## SEER 2.0

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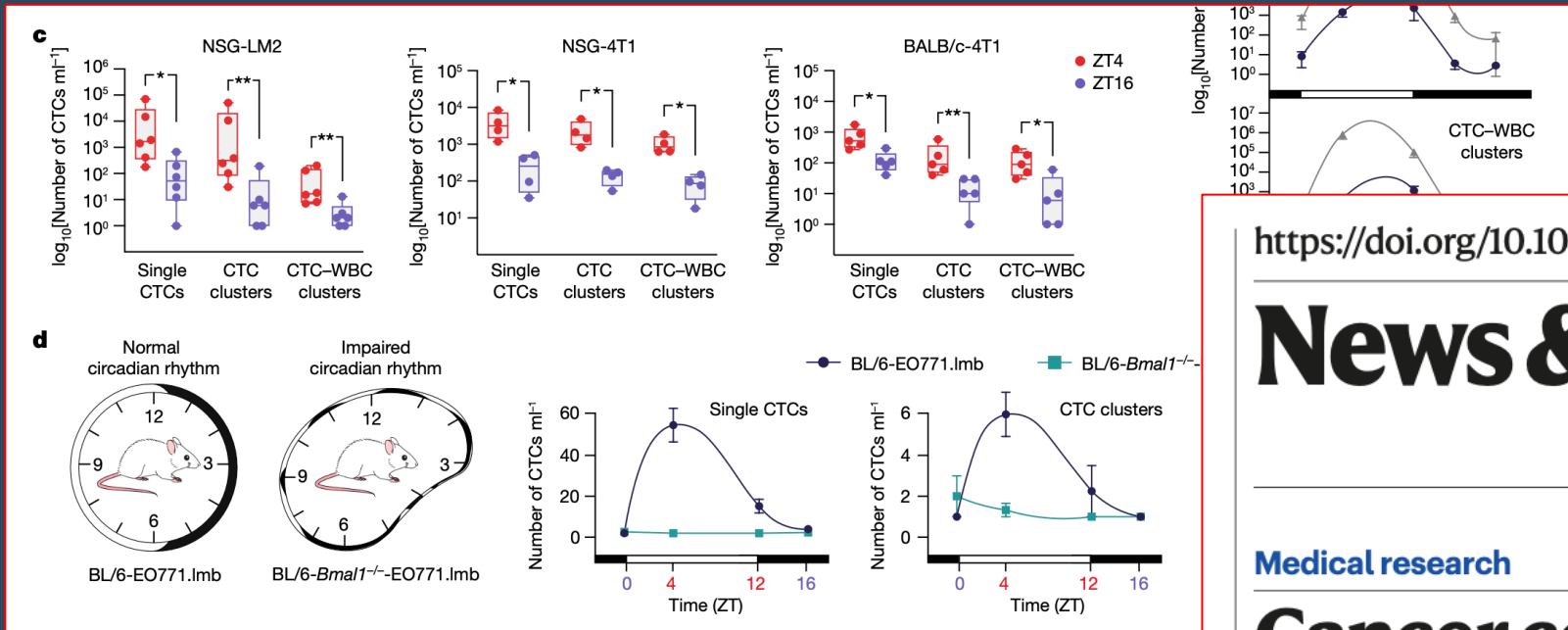


Pre-Analytics • 3/9/22

## Yin and Yang Study

# Diamantopoulou et al. ETH Zurich

## The metastatic spread of breast cancer accelerates during sleep



<https://doi.org/10.1038/d41586-022-01639-6>

## News & views

Medical research

## Cancer cells spread aggressively during sleep

Harrison Ball & Sunitha Nagrath

The deadly spread of cancer occurs predominantly during

featured papers







# Jan Trøst Jørgenson (Denmark)

## Missing companion diagnostic for US FDA Approved

abstract

Within hematology and oncology, companion diagnostics (CDxs) play an increasing role in securing an optimal therapy for individual patients, and the US Food and Drug Administration (FDA) consider this type of assay essential for the safe and effective use of a corresponding therapeutic product. Most CDxs are developed prospectively using the drug-diagnostic codevelopment model, which normally secures the simultaneous approval of both drugs and diagnostics. A CDx assay is an important treatment decision tool that needs to be available simultaneously with the drug. However, within the past few years, several targeted drugs and new indications have been approved by the FDA without a CDx, despite the use of a predictive biomarker assay for patient selection during clinical development. A missing analytical and clinically validated CDx assay could affect the correct use of these drugs and ultimately patient safety. An alternative to FDA-approved or FDA-cleared CDxs could be to use a laboratory-developed test, which will normally miss documentation on the clinical validity. On the basis of the information available from different publicly available FDA databases, this article briefly discusses the issue of missing CDx assays in relation to the approval of hematological and oncological drugs and indications.

J Clin Oncol 6:e2200100. © 2022 by American Society of Clinical Oncology

featured papers

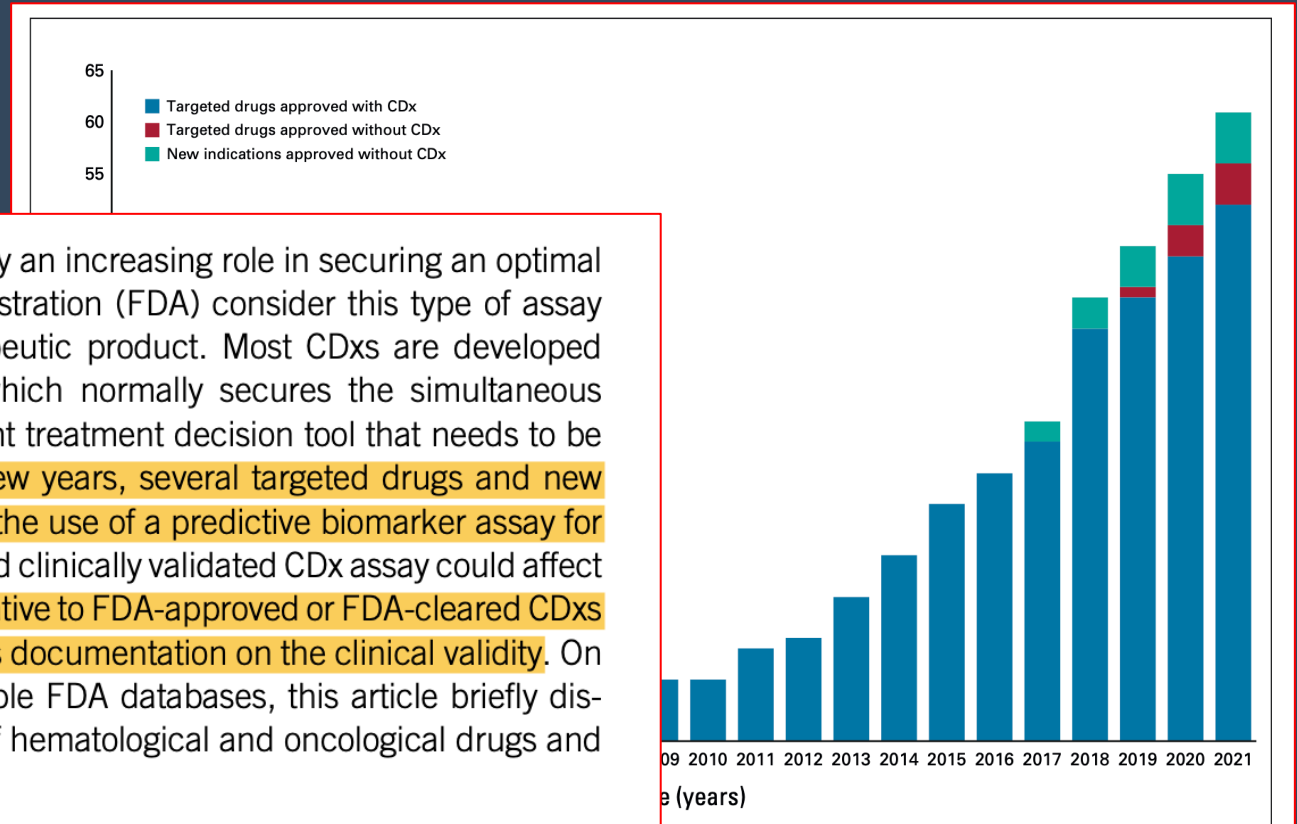


Figure 1. Number of FDA-approved CDx-drug combinations within hematology and oncology from 1998 to present. The figure also includes new drugs and indications that have been approved without a CDx assay. CDx, companion diagnostic.



# Lips et al. \* Amsterdam UK etc...

## Genomic analysis defines clonal relationships of ductal carcinoma in situ and recurrent invasive breast cancer

nature  
genetics

ARTICLES

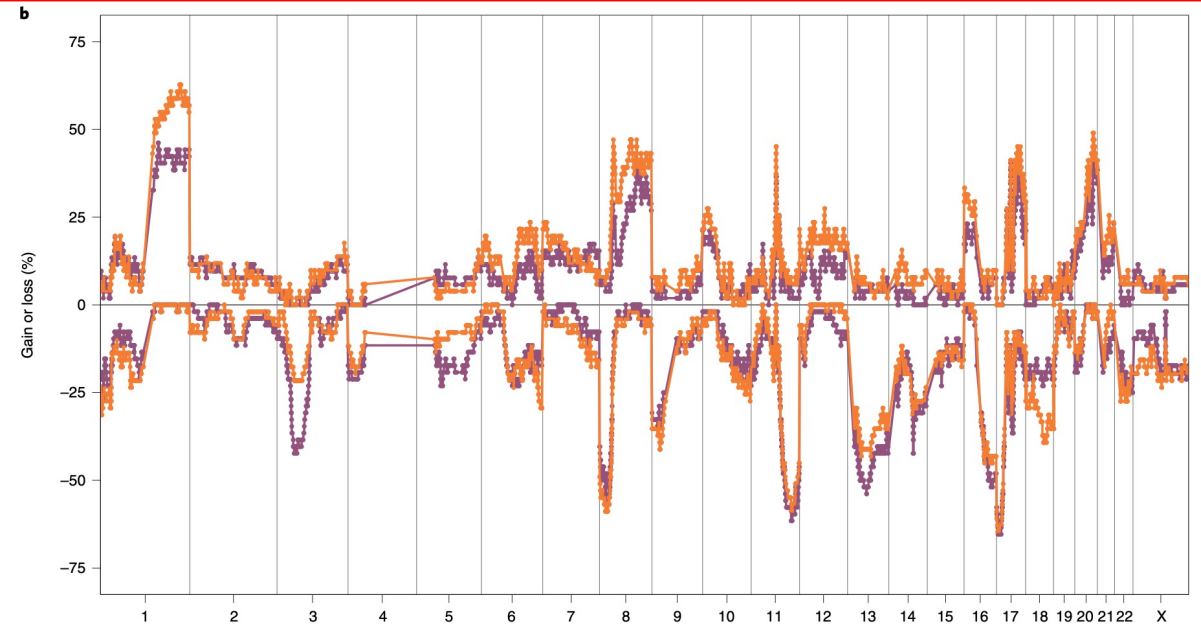
<https://doi.org/10.1038/s41588-022-01111-1>

Check for updates

OPEN

### Genomic analysis defines clonal relationships of ductal carcinoma in situ and recurrent invasive breast cancer

Esther H. Lips <sup>1,2,4</sup>, Tapsi Kumar <sup>2,3,4,24</sup>, Anargyros Megalios <sup>5,24</sup>, Lindy L. Visser <sup>1</sup>, Michael Sheinman<sup>6</sup>, Angelo Fortunato<sup>7,8</sup>, Vandna Shah<sup>5</sup>, Marlous Hoogstraat <sup>6</sup>, Emi Sei<sup>3</sup>, Diego Mallo <sup>7,8</sup>, Maria Roman-Escorza <sup>5</sup>, Ahmed A. Ahmed <sup>5</sup>, Mingchu Xu<sup>2</sup>



**Fig. 6 | Mutations and copy number alterations in primary DCIS and subsequent clonally related invasive recurrences.** **a**, Oncoplots for primary DCIS samples (left) and invasive recurrences (right) based on WES and targeted sequencing. Of the 45 genes covered by all sequencing platforms, only genes mutated in more than 3% of the primary DCIS or invasive recurrence samples are shown. We removed C>T mutations with allele frequency < 0.1 and fewer than three entries in the COSMIC database. **b**, Frequency plot of genome-wide copy number alterations in clonally related DCIS and invasive recurrences ( $n=55$ ) showing primary DCIS (purple) and its paired ipsilateral invasive recurrence (orange). The y axis shows the percentage of samples with gains (above zero line) and losses (below zero line). The genomic position is indicated by chromosome 1 on the left and up to chromosome X on the right with chromosome boundaries indicated by vertical lines.

featured papers



# Maier-Hein et al. (German/Swiss/Czech)

## BIAS: Transparent reporting of biomedical image analysis challenges



Contents lists available at [ScienceDirect](#)

### Medical Image Analysis

journal homepage: [www.elsevier.com/locate/media](http://www.elsevier.com/locate/media)



## BIAS: Transparent reporting of biomedical image analysis challenges



Lena Maier-Hein<sup>a,\*</sup>, Annika Reinke<sup>a</sup>, Michal Kozubek<sup>b</sup>, Anne L. Martel<sup>c,d</sup>, Tal Arbel<sup>e</sup>,  
Matthias Eisenmann<sup>a</sup>, Allan Hanbury<sup>f,g</sup>, Pierre Jannin<sup>h</sup>, Henning Müller<sup>i,j</sup>, Sinan Onogur<sup>a</sup>,  
Julio Saez-Rodriguez<sup>k,l,m</sup>, Bram van Ginneken<sup>n</sup>, Annette Kopp-Schneider<sup>o</sup>,  
Bennett A. Landman<sup>p</sup>

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# Mund et al.

## Unbiased spatial proteomics with single-cell resolution in tissues

Molecular Cell

CellPress

Technology review

### Unbiased spatial proteomics with single-cell resolution in tissues

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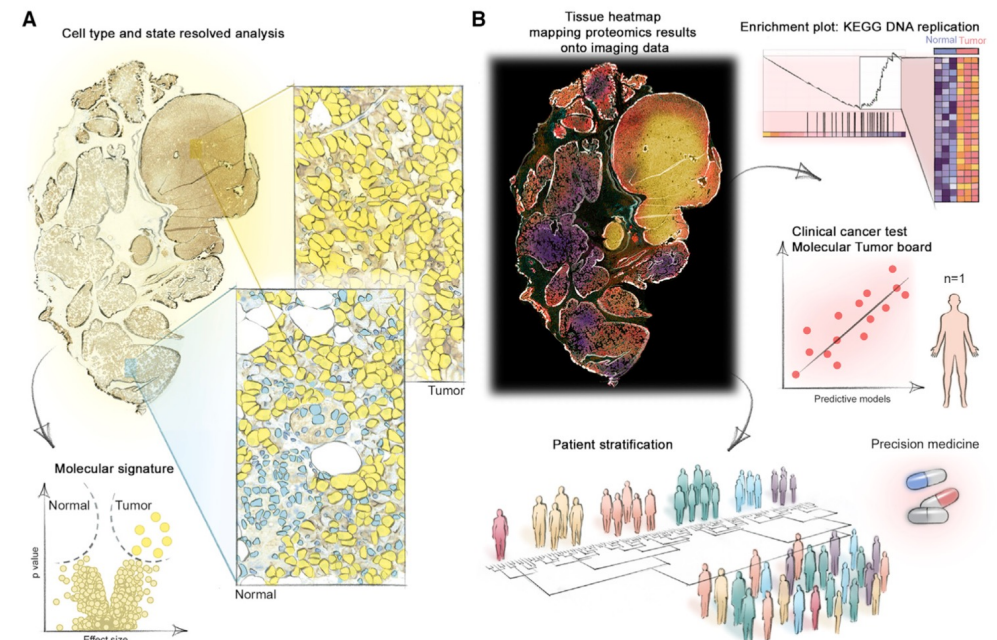
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<https://doi.org/10.1016/j.molcel.2022.05.022>

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**Figure 5. Clinical applications of spatial proteomics for patient phenotyping**

(A) High-resolution tissue maps allow machine-learning-based accurate cell segmentation and classification. Spatial proteomics analysis reveals disease-specific molecular signatures in their native tissue context, directly from normal or tumor FFPE tissue slices.

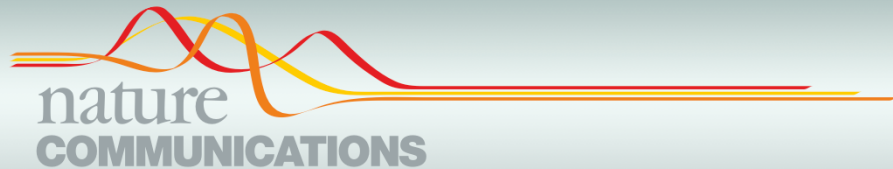
(B) Combining unbiased proteomics with high-content imaging generates a phenotype map including the tissue microenvironment. Out of the detailed and quantitative proteomic map of the tissue, matrices, profiles, enrichment plots, and neighborhood analysis can be generated to define phenotypic relationships and mine the spatial correlations in the data to provide diagnostic decision support.





# Nacev et al. (MSKCC)

## Clinical sequencing of soft tissue and bone sarcomas delineates diverse genomic landscapes and potential therapeutic targets



ARTICLE

<https://doi.org/10.1038/s41467-022-30453-x>

OPEN

Clinical sequencing of soft tissue and bone sarcomas delineates diverse genomic landscapes and potential therapeutic targets

Benjamin A. Nacev<sup>1,2,3,15</sup>, Francisco Sanchez-Vega<sup>4,14,15</sup>, Shaleigh A. Smith<sup>4,5</sup>,  
Ana R. Antonescu<sup>6</sup>, Evan Rosenbaum<sup>1,2</sup>, Hongyu Shi<sup>7</sup>, Cerise Tang<sup>6,8</sup>, Nicholas D. Socci<sup>5,9</sup>, Satshil Rana<sup>6</sup>,  
Gularte-Mérida<sup>4</sup>, Ahmet Zehir<sup>6</sup>, Mrinal M. Gounder<sup>1,2</sup>, Timothy G. Bowler<sup>1</sup>, Anisha Luthra<sup>5,10</sup>,  
1,2,3,4,5,6,7,8,9,10,11,12,13,14,15

Because a pan-cancer MSK-IMPACT analysis identified *TERT* promoter mutations in a subset of sarcomas<sup>22</sup>, we investigated *TERT* alteration frequency as a function of sarcoma subtype (Fig. 4C). We identified oncogenic *TERT* amplifications in 44% (8/18) of intimal sarcoma (INTS) and *TERT* promoter mutations in 79% (38/48) of MRLS, 46% (24/52) of SFT, and 35% (5/14) of dedifferentiated chondrosarcoma (DDCHS). In DDLS, oncogenic *TERT* promoter alterations were present in 16% of samples (27/167) and were almost entirely amplifications ( $n = 24$ ). *TERT* copy

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# Park et al. (Seoul, Korea; Lunit)

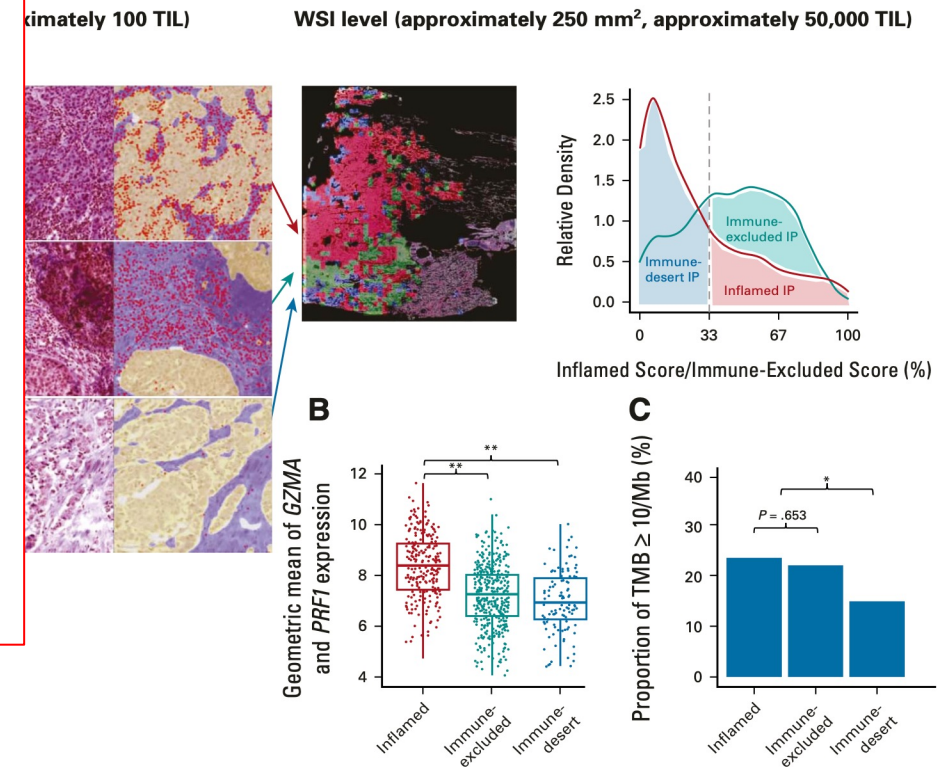
## Artificial Intelligence-Powered spatial analysis of TILs as complementary biomarker for immune checkpoint inhibition in NSCL

original reports

### Artificial Intelligence–Powered Spatial Analysis of Tumor-Infiltrating Lymphocytes as Complementary Biomarker for Immune Checkpoint Inhibition in Non–Small-Cell Lung Cancer

Sehhoon Park, MD, PhD<sup>1</sup>; Chan-Young Ock, MD, PhD<sup>2</sup>; Hyojin Kim, MD, PhD<sup>3</sup>; Sergio Pereira, PhD<sup>2</sup>; Seonwook Park, PhD<sup>2</sup>; Minuk Ma, MS<sup>2</sup>; Sangjoon Choi, MD<sup>4</sup>; Seokhwi Kim, MD, PhD<sup>5</sup>; Seunghwan Shin, MD<sup>2</sup>; Brian Jaehong Aum, PhD<sup>2</sup>; Kyunghyun Paeng, MS<sup>2</sup>; Donggeun Yoo, PhD<sup>2</sup>; Hongui Cha, PhD<sup>1</sup>; Sunyoung Park, PhD<sup>1</sup>; Koung Jin Suh, MD<sup>6</sup>; Hyun Ae Jung, MD, PhD<sup>1</sup>; Se Hyun Kim, MD, PhD<sup>6</sup>; Yu Jung Kim, MD, PhD<sup>6</sup>; Jong-Mu Sun, MD, PhD<sup>1</sup>; Jin-Haeng Chung, MD, PhD<sup>3</sup>; Jin Seok Ahn, MD, PhD<sup>1</sup>; Myung-Ju Ahn, MD, PhD<sup>1</sup>; Jong Seok Lee, MD, PhD<sup>6</sup>; Keunchil Park, MD, PhD<sup>1</sup>; Sang Yong Song, MD, PhD<sup>4</sup>; Yung-Jue Bang, MD, PhD<sup>7</sup>; Eun-La Choi, MD, PhD<sup>4</sup>; Tony S. Mok, MD<sup>8</sup>; and Se-Hoon Lee, MD, PhD<sup>1,9</sup>

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# Reinke et al. (DKFZ, and many others)

## Common limitations of image processing metrics: a picture story

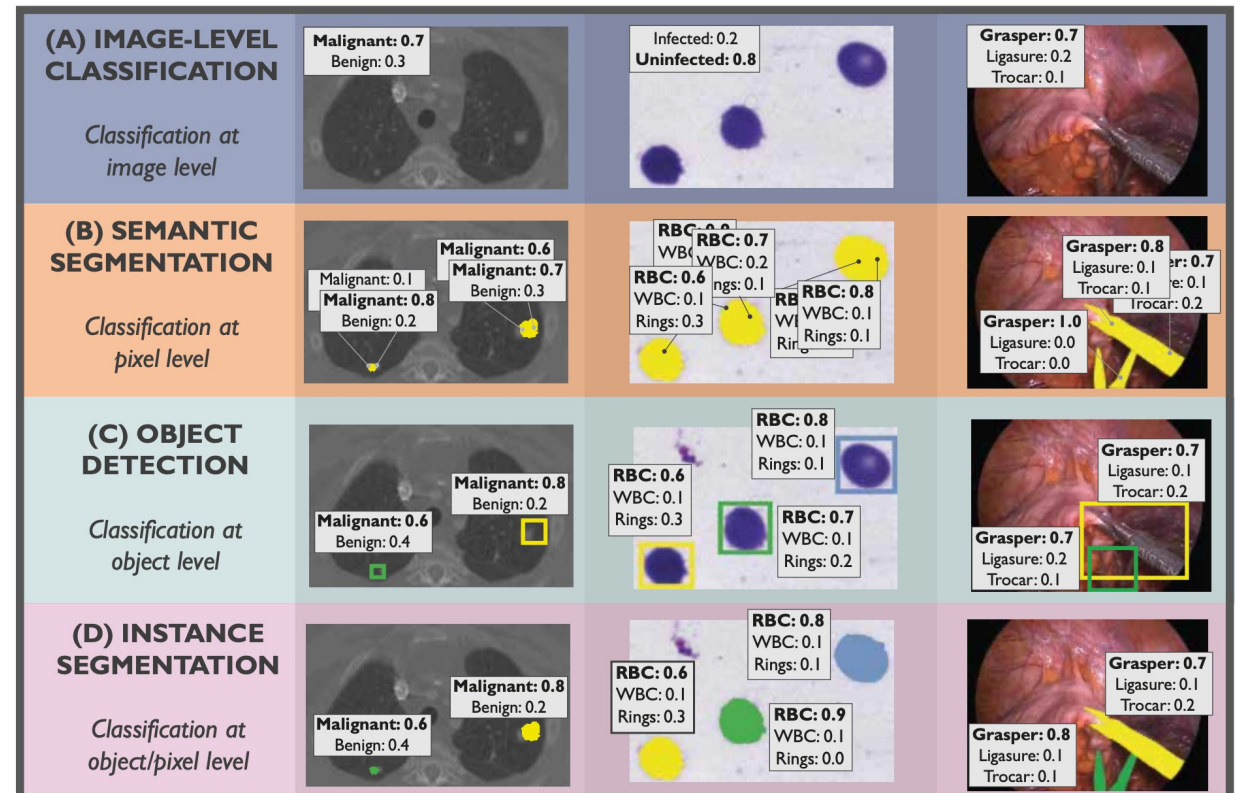
### Common Limitations of Image Processing Metrics: A Picture Story

ANNIKA REINKE\*, German Cancer Research Center (DKFZ), Germany and Heidelberg University, Germany  
 MINU D. TIZABI, German Cancer Research Center (DKFZ), Germany  
 CAROLE H. SUDRE, University College London, UK and King's College London, UK  
 MATTHIAS EISENMANN, German Cancer Research Center (DKFZ), Germany  
 TIM RÄDSCH, German Cancer Research Center (DKFZ), Germany and University of Tübingen, Germany

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Problem categories addressed by this paper



# Sherman et al.

## Genome-wide mapping of somatic mutation rates uncovers drivers of cancer

nature  
biotechnology

ARTICLES

<https://doi.org/10.1038/s41587-022-01353-8>

 Check for updates

OPEN

## Genome-wide mapping of somatic mutation rates uncovers drivers of cancer

Maxwell A. Sherman <sup>1,2,3,4,10</sup>, Adam U. Yaari<sup>1,4,5,10</sup>, Oliver Priebe<sup>1,4,6,10</sup>, Felix Dietlein <sup>4,7,9</sup>, Po-Ru Loh <sup>3,4</sup>   
and Bonnie Berger <sup>1,2,4,8</sup> 

Identification of cancer driver mutations that confer a proliferative advantage is central to understanding cancer; however, searches have often been limited to protein-coding sequences and specific non-coding elements (for example, promoters) because of the challenge of modeling the highly variable somatic mutation rates observed across tumor genomes. Here we present Dig, a method to search for driver elements and mutations anywhere in the genome. We use deep neural networks to map cancer-specific mutation rates genome-wide at kilobase-scale resolution. These estimates are then refined to search for evidence of driver mutations under positive selection throughout the genome by comparing observed to expected mutation counts. We mapped mutation rates for 37 cancer types and applied these maps to identify putative drivers within intronic cryptic splice regions, 5' untranslated regions and infrequently mutated genes. Our high-resolution mutation rate maps, available for web-based exploration, are a resource to enable driver discovery genome-wide.

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# Thank you

Next month's steering committee on  
July 27, 2022 at 3:00-4:00 PM Eastern Time

Things to expect by then

- 1) Update MDUFA/VALID act
- 2) Updates from projects and workgroups
- 3) Please send links/papers/other

