

Development of Al-based diagnostic tools based on massive digital pathology datasets

LUMI AI Factory Launch @ Pikku-Finlandia 2.4.2025 Pekka Ruusuvuori, Associate Professor Institute of Biomedicine, University of Turku <u>pekka.ruusuvuori@utu.fi</u> LinkedIN: @PekkaRuusuvuori



Mira Valkonen – Sartaj Salman - Masi Valkonen – Pekka Ruusuvuori (PI) Hesam Hakimnejad – Muhammad Adnan – Hyder Abbas - Rimsha Kaokab – Henrique Hiram Libutti – Niloufar Rahimizadeh

Bioimage informatics research group

We develop computational methods and tools using modern machine learning to quantify and extract information from biomedical images for diagnostics, decision making and for data-driven cancer research and medicine.





#ERAPerMed







ComPatAl consortium

• ComPatAI:

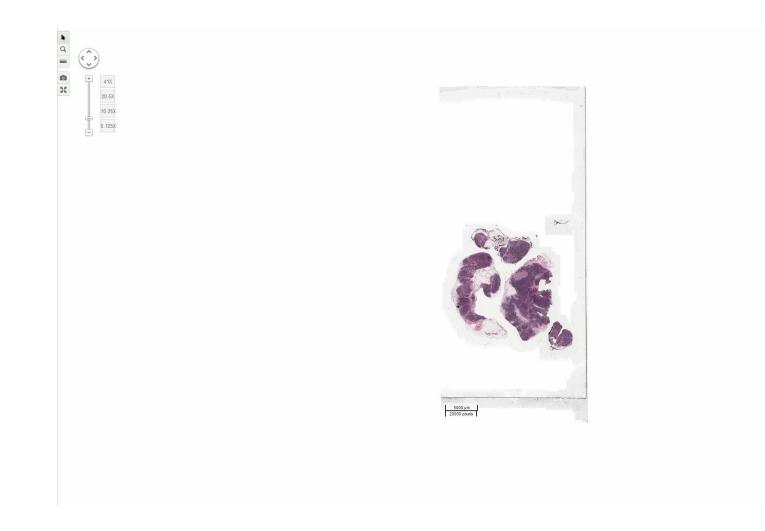
- Large scale AI-development for histopathology
- Extreme scale access projects for LUMI-G (2023, 2024, 2025)
- Research Council of Finland funded project "Computationally intensive modeling of histopathology using generative and predictive AI", 2024-2026 (Ruusuvuori & Latonen)
- People:
 - Assoc.Prof. Pekka Ruusuvuori, University of Turku, director
 - Al-development, generative Al, high-performance computing, computational pathology
 - Adj.Prof. Leena Latonen, University of Eastern Finland:
 - Virtual staining in histopathology, histopathology expertise, AI development
 - Adj. Prof. Teemu Tolonen, FIMLAB Laboratories & Tampere University Hospital
 - Pathology expertise, FIMLAB data access











One whole slide image: 97 000 x 220 000 pixels = **21.3 Gigapixels**

Al-based diagnostic systems with human expert level accuracy



Deep learning based detection and grading of cancer



Can AI detect and grade prostate cancer?

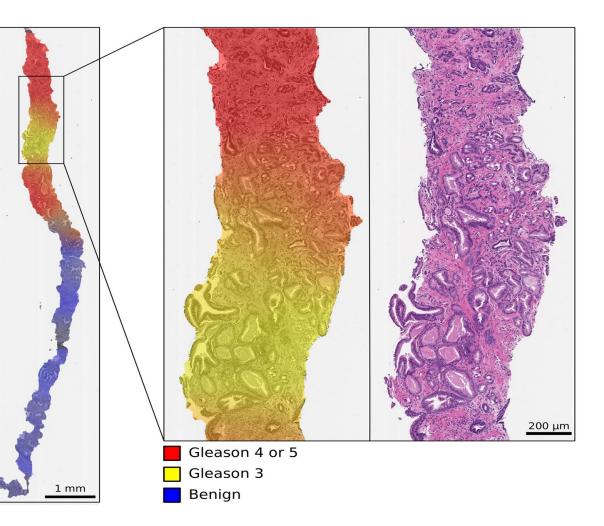
THE LANCET Oncology



Artificial intelligence for diagnosis and grading of prostate cancer in biopsies: a population-based, diagnostic study

Peter Ström, MSc[↑] • Kimmo Kartasalo, MSc[↑] • Henrik Olsson, MSc • Leslie Solorzano, MSc • Prof Brett Delahunt, MD • Prof Daniel M Berney, MD • Prof David G Bostwick, MD • Andrew J Evans, MD • Prof David J Grignon, MD • Prof Peter A Humphrey, MD • Prof Kenneth A Iczkowski, MD • Prof James G Kench, MD • Prof Glen Kristiansen, MD • Prof Theodorus H van der Kwast, MD • Prof Katia R M Leite, MD • Jesse K McKenney, MD • Jon Oxley, MD • Chin-Chen Pan, MD • Prof Hemamali Samaratunga, MD • Prof John R Srigley, MD • Hiroyuki Takahashi, MD • Prof Toyonori Tsuzuki, MD • Murali Varma, MD • Prof Ming Zhou, MD • Johan Lindberg, PhD • Cecilia Lindskog, PhD • Pekka Ruusuvuori, PhD • Prof Carolina Wählby, PhD • Prof Henrik Grönberg, MD • Mattias Rantalainen, PhD • Prof Lars Egevad, MD • Martin Eklund, PhD $\stackrel{\circ}{\sim}$ Show less • Show footnotes

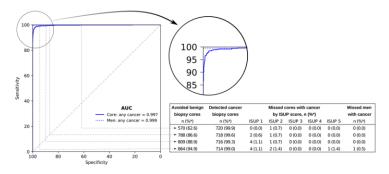
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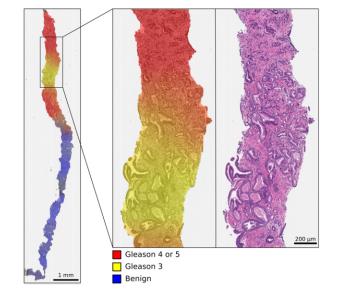


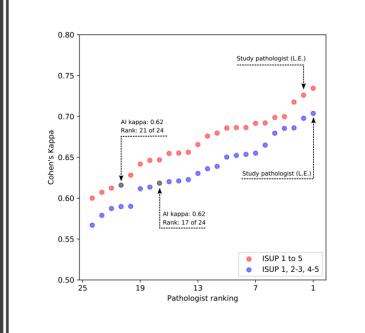


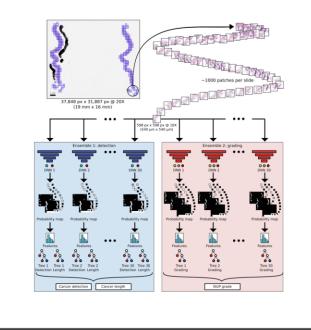
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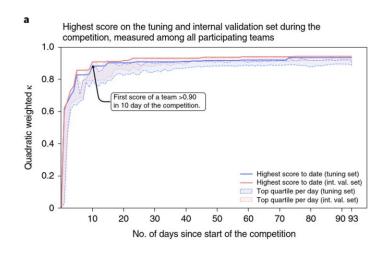




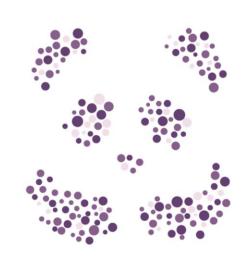




Al-system diagnoses prostate cancer from whole slide images on human expert level

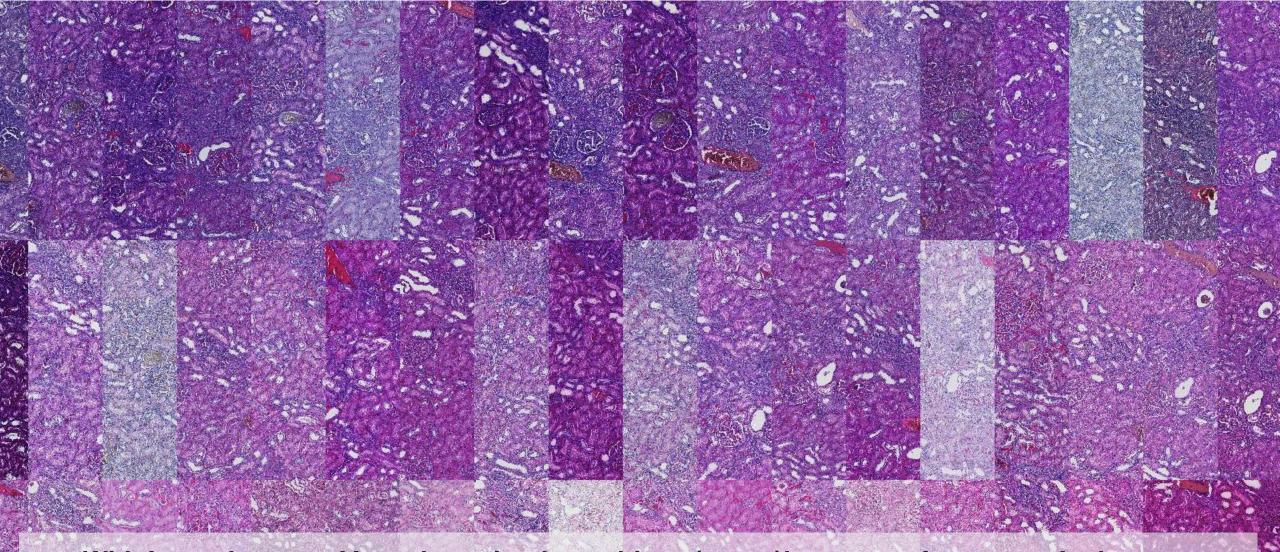






PANDA prostate cancer grade assessment

Crowdsourced AI development outperforms human experts in prostate cancer grading



With large datasets AI can be trained to achieve (super)human performance. At the same time, AI may fail to generalize in situations where humans can easily adjust. Figure from Khan, Härkönen, Friman, Latonen, Kuopio, Ruusuvuori. Staining normalization in histopathology: Multi-center dataset and method benchmarking (Submitted).

Towards foundational AI models using population level digital pathology cohort

- Major Finnish hospitals have adopted fully digital workflow in pathology diagnostics
 - Turku & Tampere university hospitals use digital pathology in routine diagnostics:
 - ~600 000 whole slide images annually
- Routine digitization of all biopsies leads to petabyte level data collections
- We currently have access to roughly ~1M WSI collection
- Can digital pathology data from routine diagnostics collected at hospitals be harnessed to build better diagnostic tools?

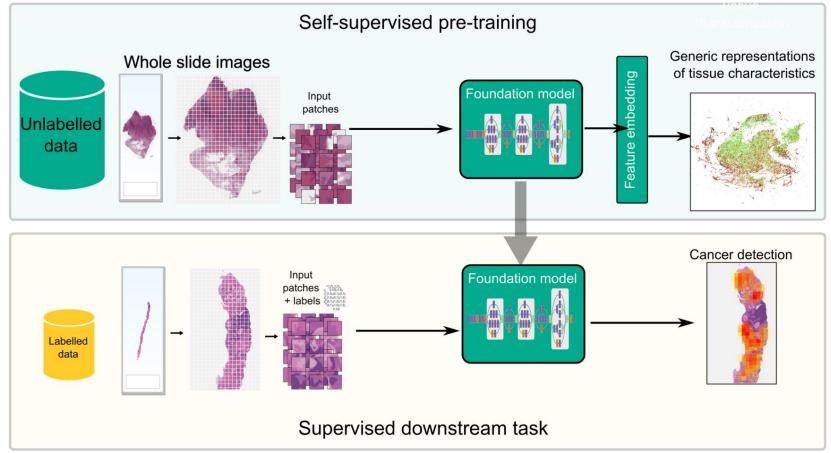


Massive digital pathology data + high performance computing + AI

- · Access to massive, high-quality digital pathology data
- State-of-the-art high-performance computing resources: LUMI supercomputer & CSC as a national service provider
- High level of computational data science & AI expertise
- Research Council of Finland and Cancer Foundation Finland fund the study



Towards generalizable AI for histopathology: foundation models using self-supervised learning (SSL)



Valkonen, M., Aho, A., Tolonen, T., Latonen, L., Kather, J., Ruusuvuori, P. Self-supervised learning architectures as the basis for foundation models in histopathology: A comparative study (Manuscript)

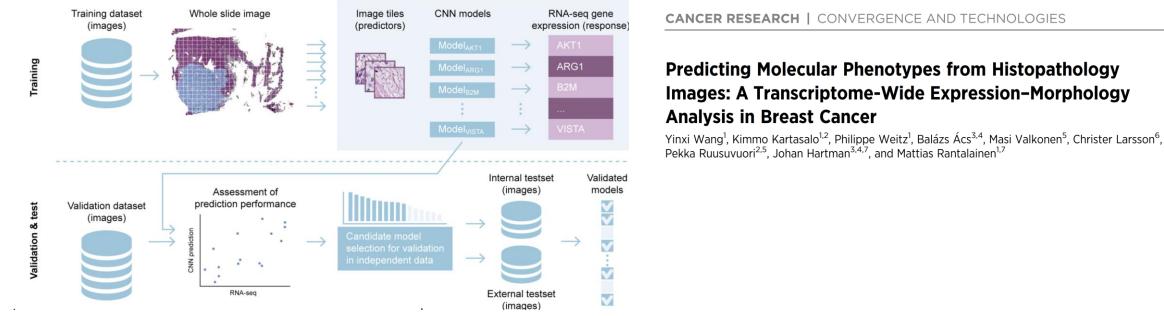
Combining pathology and genetics using AI

Prediction of gene expression from histopathology images



Computational pathology beyond human vision: Gene expression prediction directly from histology

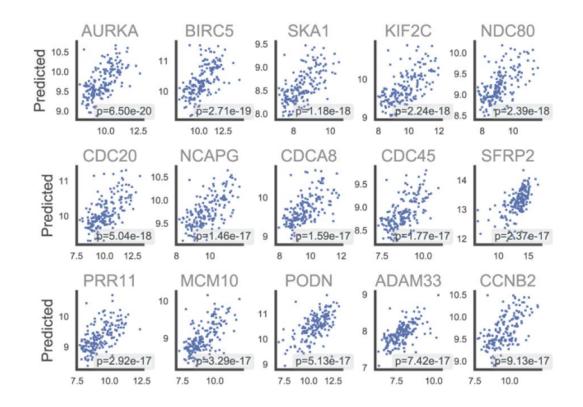
- CSC Grand Challenge Pilot Project
- Transcriptome-wide Expression-MOrphology (EMO) analysis in breast cancer
 - Individual deep convolutional neural networks (CNNs) were optimised to predict the expression of 17,695 genes from hematoxylin and eosin (HE) stained whole slide images (WSIs).
 - Ensembles of 30 CNN x 18 000 genes = huge computational effort •

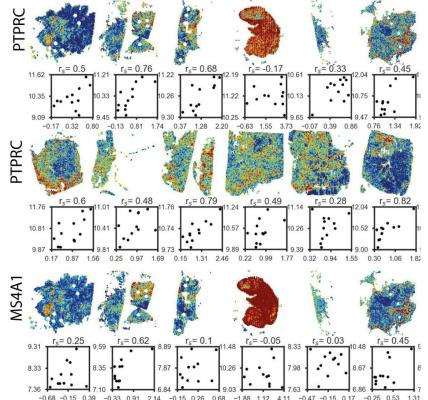


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Computational pathology beyond human vision: Gene expression prediction directly from histology

- Prediction of both tumour average gene expression and intra-tumour spatial expression is possible directly from tissue morphology
- Predicted expressions in 9,334 (52.75%) genes were significantly associated with RNA sequencing estimates internal test set

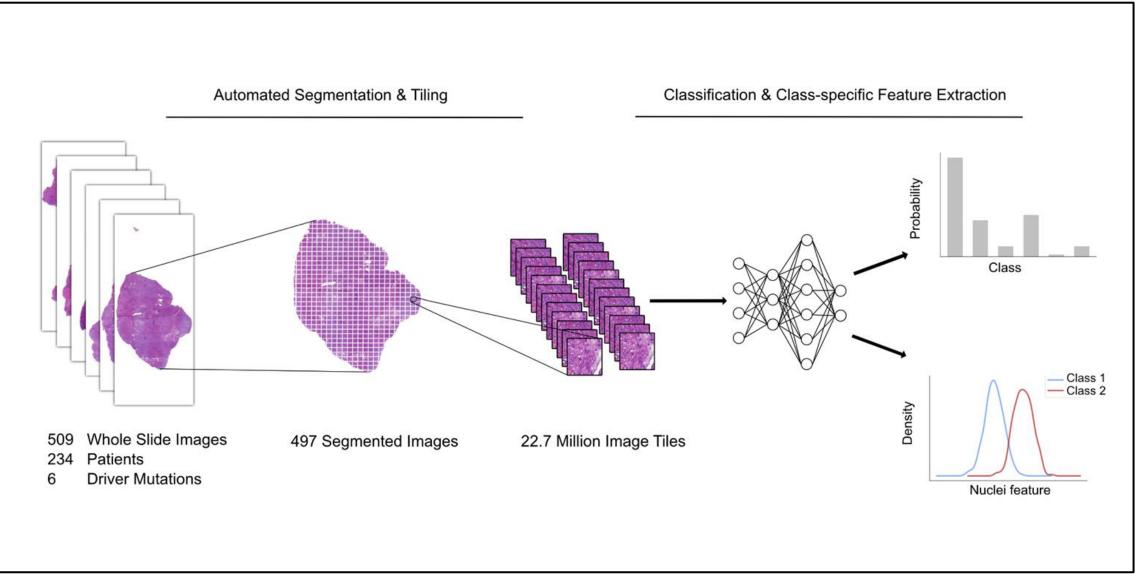




Computational pathology beyond human vision

Prediction of mutations from histopathology images

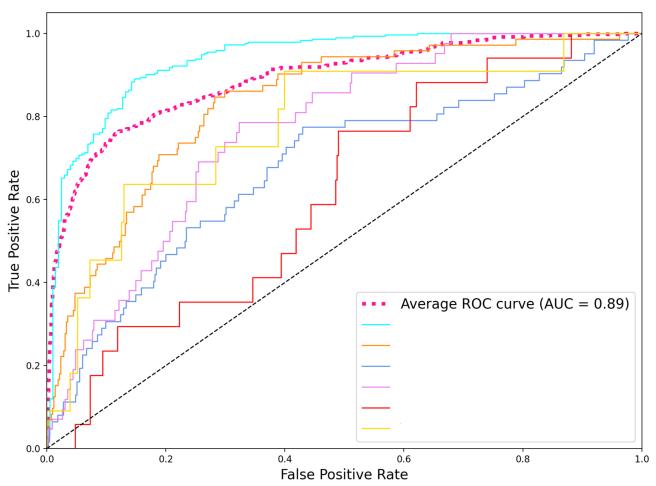




Kiviaho, Nykter, Aaltonen, Ruusuvuori et al. Manuscript.

Predicting mutational status in Uterine Leiomyoma (Fibroids)

- Benign tumors of the uterine wall
- Affect one in four women
- Characterized by few driver mutations
- Good performance on mutations with reasonable sample size
- Less accurate for mutations with less samples
- Overall AUC 0.89



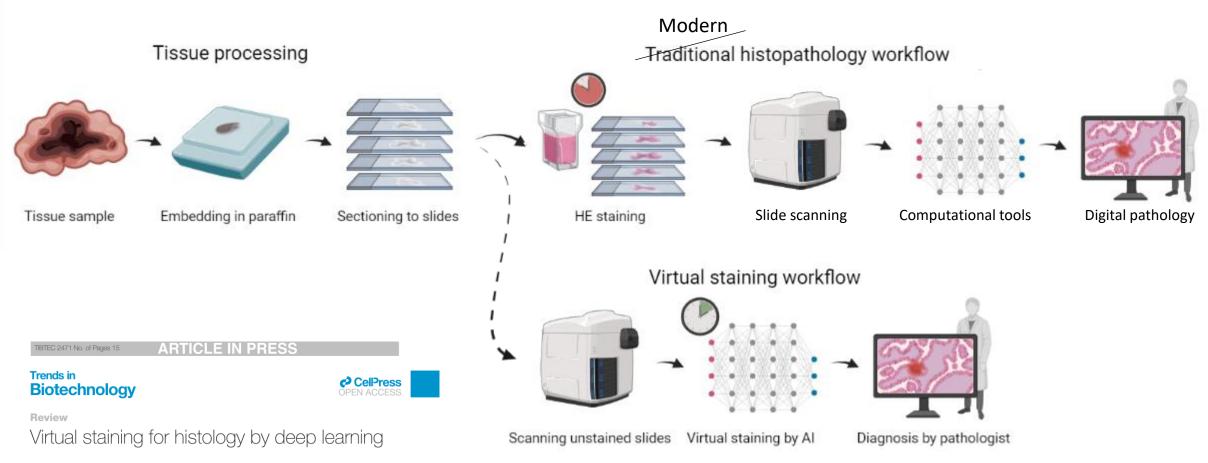
Kiviaho, Nykter (TUNI), Aaltonen (UH), Ruusuvuori et al. Manuscript.

Computational pathology beyond human vision

Virtual staining of unstained tissue using generative AI

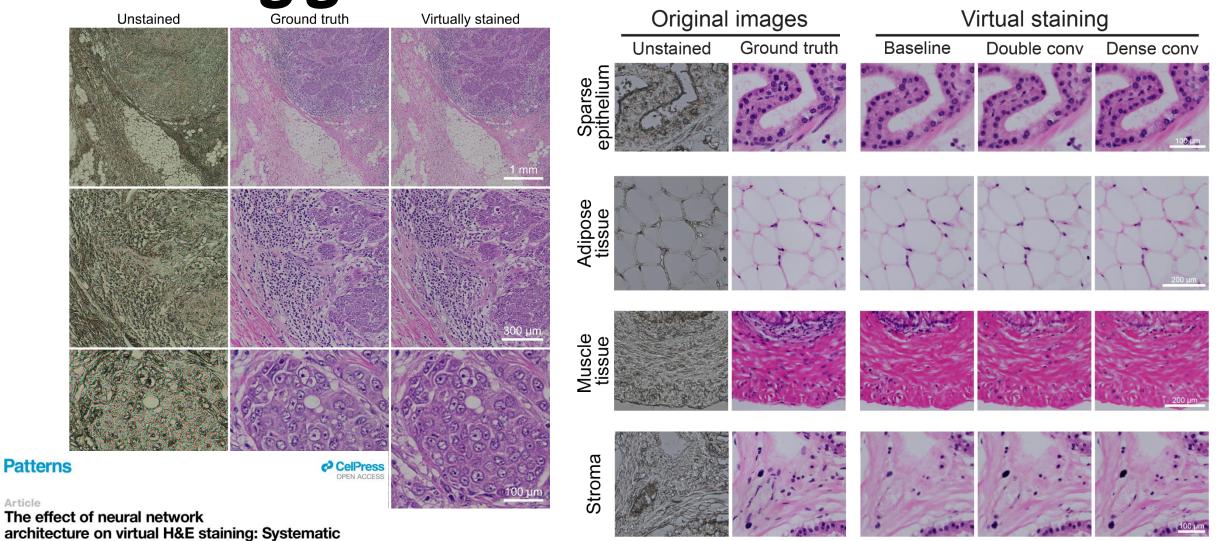


Histopathological workflow - how much can be replaced virtually?



Leena Latonen ¹,*,[@] Sonja Koivukoski,¹ Umair Khan,² and Pekka Ruusuvuori ²,[@]

Virtual staining of unstained tissue using generative Al

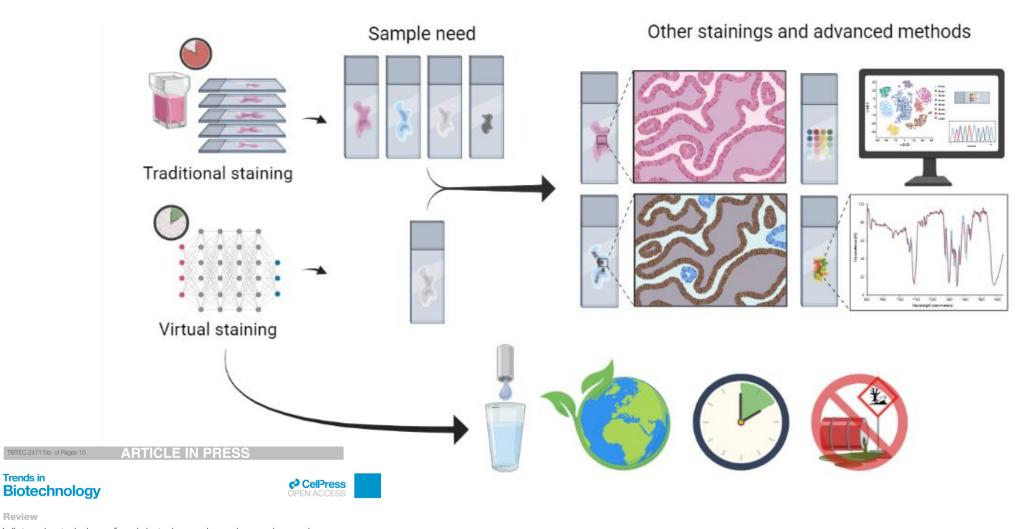


Umair Khan,¹ Sonja Koivukoski,² Mira Valkonen,³ Leena Latonen,^{2,4} and Pekka Ruusuvuori^{1,3,5,6,*}

assessment of histological feasibility

Article

Potential advantages of virtual staining



Review

Trends in

Virtual staining for histology by deep learning

Leena Latonen ⁽⁰⁾, ^{1,*, (0)} Sonja Koivukoski, ¹ Umair Khan, ² and Pekka Ruusuvuori ⁽⁰⁾ ^{2,(0)}

Translating research results for the benefit of society and economy

From innovations to startups, from novel methods to clinical trials



Societal impact

- Two start-ups
 - Quva oy est. 2010, Louhi Health Data est. 2023
- IPR:
 - Two international patents, several (~10) invention reports
- Participating in four ongoing clinical trials where AI-systems are built for diagnostics, patient characterization and treatment outcome prediction



LUMI AI Factory



Speeding up development of Al-based decision support systems for biomedical imaging



LUMI AI Factory – our wishlist for speeding up development cycle

- Massive scale means massive data transfer and computing it takes time
 - Extreme-scale access projects for longer periods
 - Continued funding support
- Regulation and data access
 - Speed-up in handling ethical/data access/tissue processing permissions
 - For example, our timeline for tackling permissions >2yrs
 - Storage of diagnostic data 2 years? Longer who covers the cost?



Thank you!



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