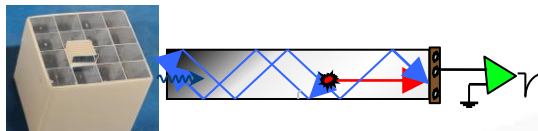


Medical Physics

Vision/Research lines/Social Impact



L1. Sensors/ Signal processing

IOP Publishing | Institute of Physics and Engineering in Medicine
Phys. Med. Biol. 61 (2016) 4679–4698
Physics in Medicine & Biology
doi:10.1088/0031-9155/61/12/4679

A new method for depth of interaction determination in PET detectors

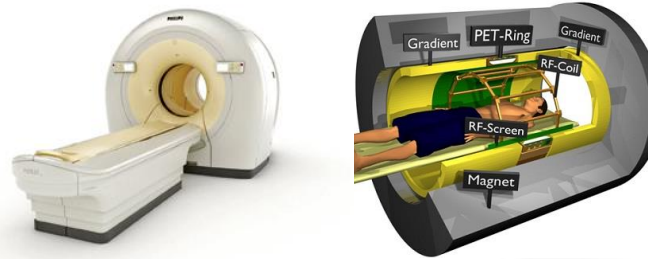
M Pizzichemi¹, G Stringhini^{1,2}, T Niknejad¹, Z Liu¹, P Lecoq², S Tavernier^{3,4}, J Varela¹, M Paganoni¹ and E Auffray²

¹ University of Milano-Bicocca, Milano, Italy

² CERN, Geneva, Switzerland

³ Laboratório de Instrumentação e Física Experimental de Partículas, Portugal

⁴ Vrije Universiteit Brussel, Ixelles, Belgium



L2. Imaging systems/ multimodal image processing and decision support systems

European Journal of Nuclear Medicine and Molecular Imaging (2019) 46:2673–2699
https://doi.org/10.1007/s00259-019-04414-4

REVIEW ARTICLE

AI-based applications in hybrid imaging: how to build smart and truly multi-parametric decision models for radiomics

Isabella Castiglioni¹ · Francesca Gallivanone¹ · Paolo Soda² · Michele Avanzo³ · Joseph Stancanella^{3,4} · Marco Aiello³ · Matteo Interlenghi¹ · Marco Salvatore³



L3. Imaging infrastructures automatic classification and decision support systems

REVIEW ARTICLE

Phase-contrast X-ray imaging of breast

JANI KEVILÄINEN¹, ALBERTO BRAVIN², MANUEL FERNÁNDEZ¹, MIKKO TENHUNEN¹, PEKKA VIRKKUNEN⁴ & PEKKA SUORTTI³

¹Department of Physics and ²Department of Radiology, HUCH Cancer Center, Helsinki University Central Hospital (HUCH), Helsinki, Finland, ³Bio-medical Beamline ID17 and ⁴High Brilliance Beamline ID2, European Synchrotron Radiation Facility (ESRF), Grenoble, France and ⁵Department of Physics, University of Helsinki, Helsinki, Finland

ACTA RADIOLOGICA

Combination of advanced imaging sensors, image processing and imaging infrastructure with newly available AI algorithms can significantly impact in medicine

Empirical medicine



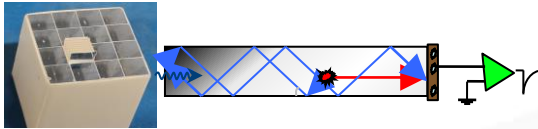
Stratified medicine



Personalised (or individualised) medicine



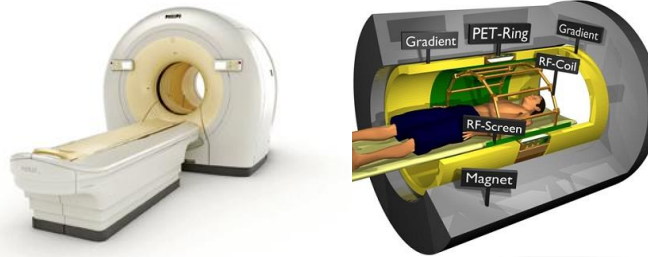
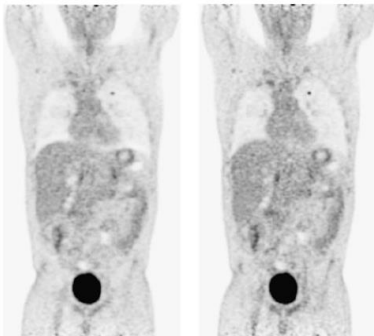
Medical Physics Research Lines and impact



L1. Sensors/ Signal processing

Better timing
Better spatial localization

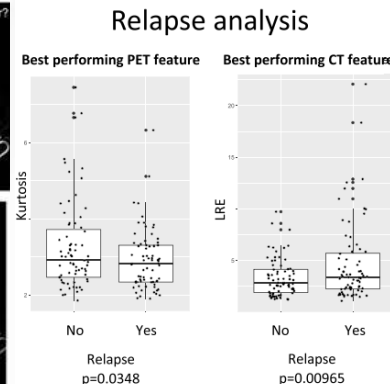
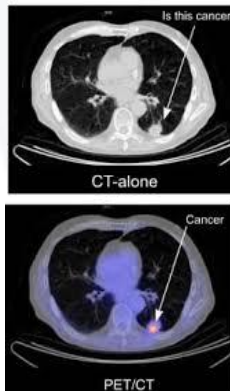
Improved lesion detectability



L2. Imaging systems/
multimodal image processing and
decision support systems

Association between complex
image content and clinical outcome

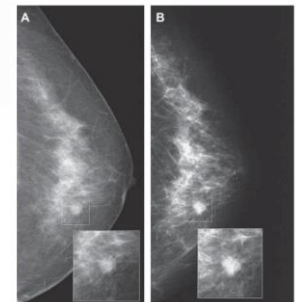
Improved medical
diagnosis/prognosis and decision



L3. Imaging infrastructures
automatic classification

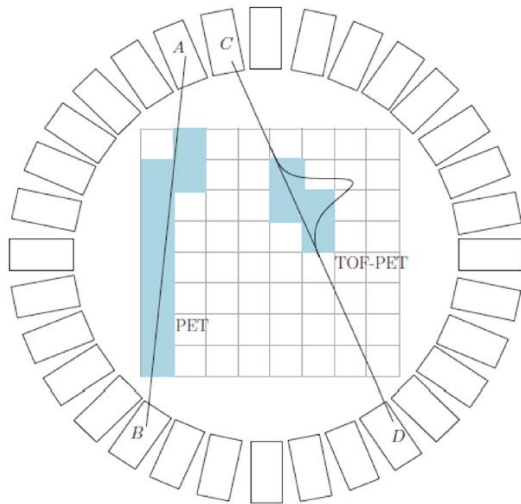
Better signal
Less noise

Improved lesion
detectability/
classification



L1. Sensors/ Signal processing

Better timing
PET with Time of flight (TOF) info



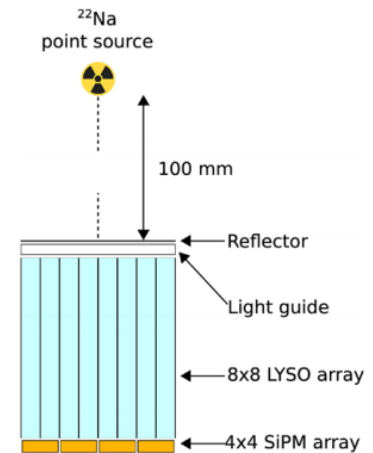
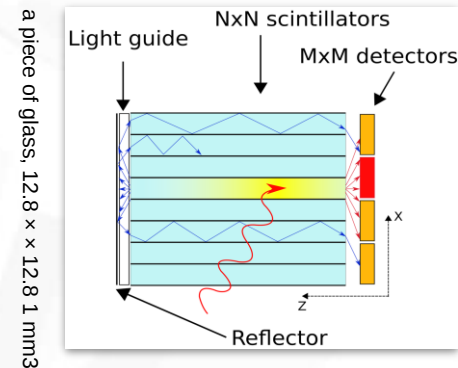
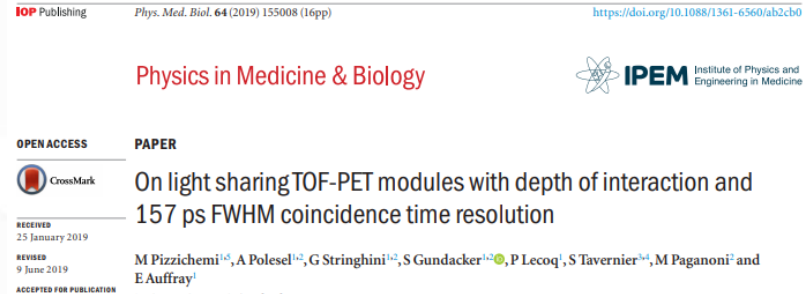
Compute the difference in time of arrival of gammas Δt

$$\Delta x = c \frac{\Delta t}{2}$$

$$SNR_{TOF} \sim \sqrt{\frac{D}{\Delta x}} \cdot SNR_{CONV}$$

Time resolution (ns)	Δx (cm)	TOF NEC gain	TOF SNR gain
0.1	1.5	26.7	5.2
0.3	4.5	8.9	3.0
0.6	9.0	4.4	2.1
1.2	18.0	2.2	1.5
2.7	40.0	1.0	1.0

Pushing coincidence time resolution down to 150 ps and including Depth of Interaction (DOI) info

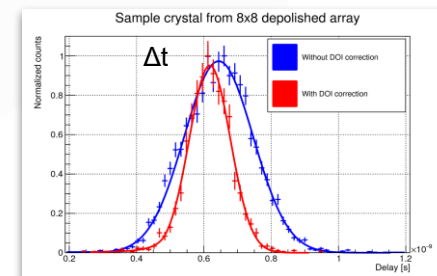


Very good timing properties

d=3mm
d=0.25mm

Array type	Crystals dim. [mm ³]	DOI res. FWHM [mm]
8x8	1.5x1.5x15	3.7 ± 0.1
8x8	1.5x1.5x15	2.8 ± 0.1

Improved lesion detectability (>25)



L1. Sensors/ Signal processing

Improving accuracy in crystal identification
(including DOI info)

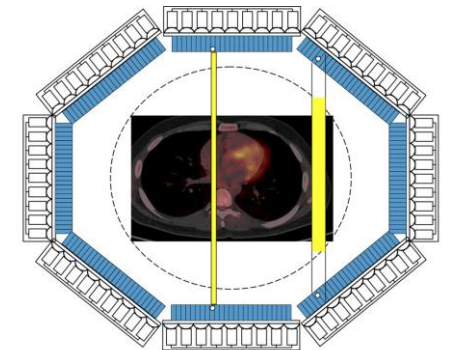
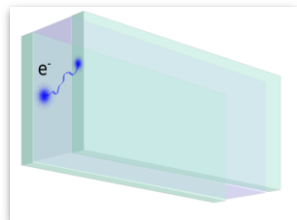
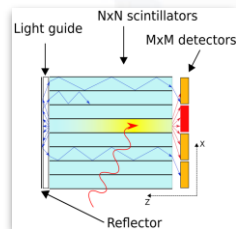


Illustration of the parallax or depth-of-interaction error associated with annihilations occurring off



Faster light
production



DOI
measurement

Time resolution (ns)	Δx (cm)	TOF NEC gain	TOF SNR gain
0.1	1.5	26.7	5.2
0.3	4.5	8.9	3.0
0.6	9.0	4.4	2.1
1.2	18.0	2.2	1.5
2.7	40.0	1.0	1.0

Pushing coincidence time resolution down to 10 ps
with accurate DOI info

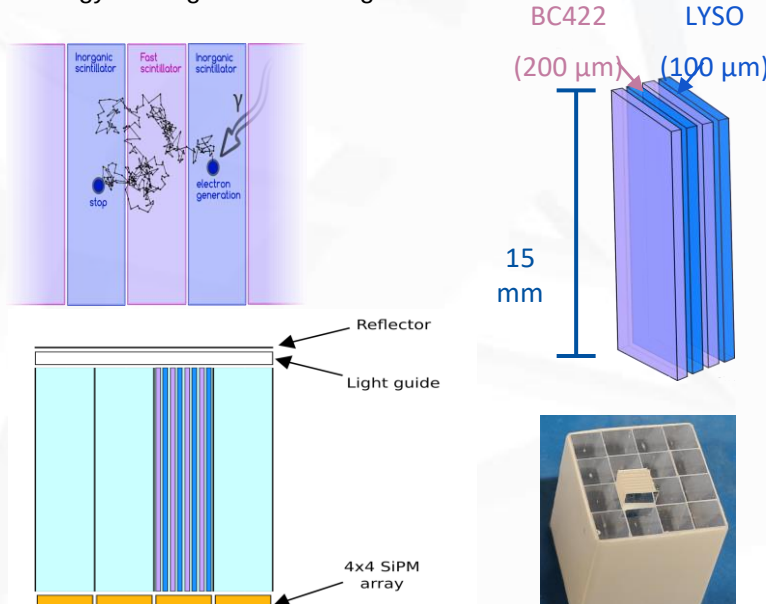


Depth of Interaction information encoding in a sampling scintillator detector geometry

Marco Pizzichemi, Rosana Martinez Turtos, Stefan Gundacker, Andrea Polesel, Marco Paganoni, Paul Lecoq,
and Etienne Auffray

Heterostructure in a crystal array

Energy sharing between inorganic and fast scintillator



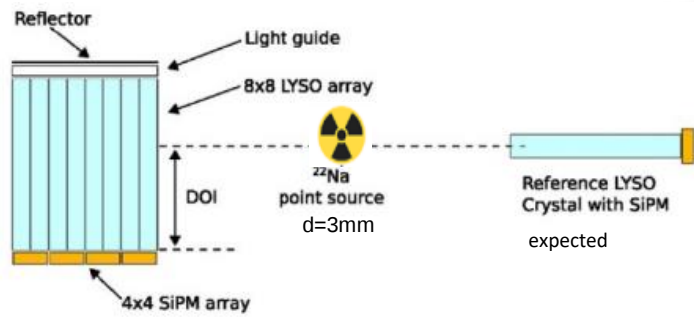
Excellent timing properties



Improved lesion detectability and
quasi-direct image reconstruction

L1. Sensors/ Signal processing

use of AI to improve sensor DOI resolution



	15	23	31	39	47	55	63
	14	22	30	38	46	54	62
5	13	21	29	37	45	53	
4	12	20	28	36	44	52	
3	11	19	27	35	43	51	
2	10	18	26	34	42	50	
1	9	17	25	33	41	49	
0	8	16	24	32	40	48	

(a) 8x8 crystal array

	15	23	31	39	47	55	63
	14	22	30	38	46	54	62
5	13	21	29	37	45	53	
4	12	20	28	36	44	52	
3	11	19	27	35	43	51	
2	10	18	26	34	42	50	
1	9	17	25	33	41	49	
0	8	16	24	32	40	48	

(a) 8x8 crystal array

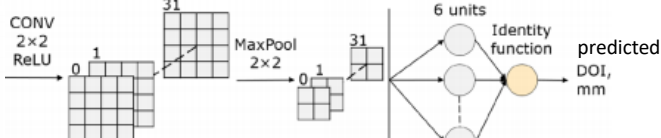
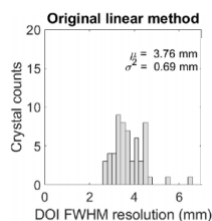
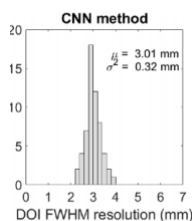


Figure 4. The architecture of the CNN. The network receives 16 integrated charge values arranged in a 4×4 matrix as an input (in green). The ReLU is employed as an activation function on the convolutional (CONV) and the FC layer. The output layer (in yellow) uses the identity function.



(a)



(d)

IOP Publishing Phys. Med. Biol. 65 (2020) 175017 <https://doi.org/10.1088/1361-6560/ab96fc>

Physics in Medicine & Biology

IPeM Institute of Physics and Engineering in Medicine

CrossMark

OPEN ACCESS

PAPER

Improving depth-of-interaction resolution in pixellated PET detectors using neural networks

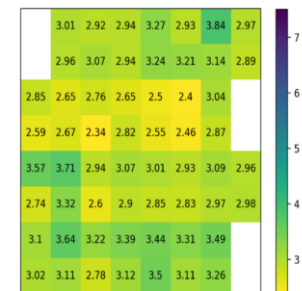
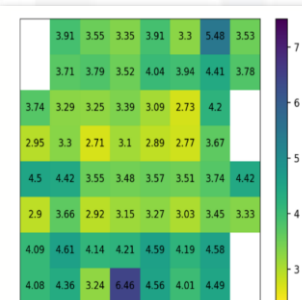
Artem Zatcepin^{1,2}, Marco Pizzichemi^{1,3}, Andrea Polesel^{1,3}, Marco Paganoni¹, Etienne Auffray⁴, Sibylle I Ziegler^{1,4} and Negar Omidvari^{1,5}

¹ Department of Nuclear Medicine, Klinikum rechts der Isar, Technical University of Munich, Ismaninger Str. 22, 81675 Munich, Germany
² CERN, Geneva, Switzerland
³ University of Milano-Bicocca, Milan, Italy
⁴ Department of Nuclear Medicine, University Hospital of LMU Munich, Marchioninstr. 15, 81377 Munich, Germany
⁵ Current address: Department of Nuclear Medicine, University Hospital of LMU Munich, Marchioninstr. 15, 81377 Munich, Germany
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RECEIVED 20 February 2020
 REVISED 24 May 2020
 ACCEPTED FOR PUBLICATION 22 June 2020
 PUBLISHED 28 August 2020

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d=3mm

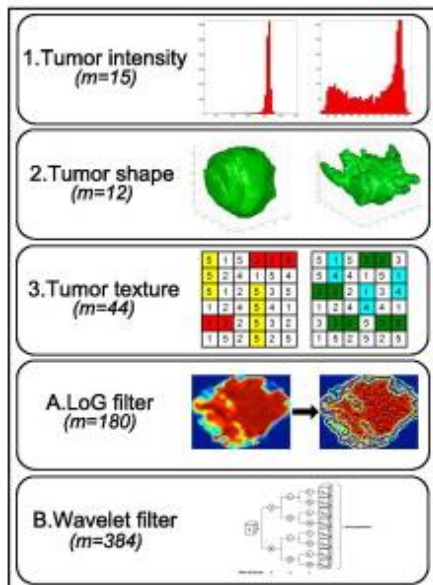


Significant improvement in LYSO DOI resolution, especially for edge crystals (err from 3.76 to 3 mm)

Beneficial impact on image quality, especially for organ dedicated PET scanners

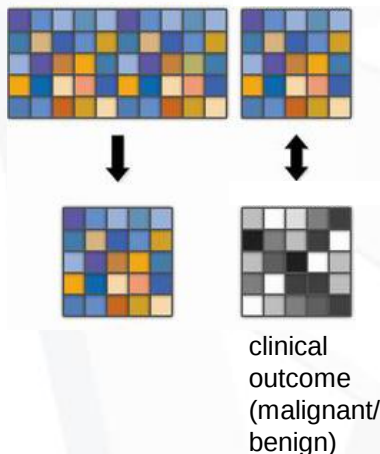
L2. Imaging systems/ multimodal image processing and decision support systems⁷

Selecting combination of structural and functional features of local disease from expert-domain (developing representation methods - radiomics)



Using such features to train machine learning to predict disease clinical outcome (developing machine learning predicting models)

Explainable AI output for clinicians



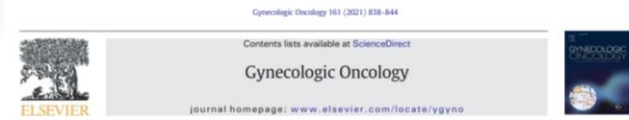
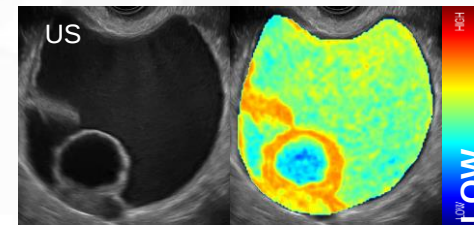
Journal of Ultrasound
<https://doi.org/10.1007/s40477-020-00503-5>

ORIGINAL PAPER



The Adoption of Radiomics and machine learning improves the diagnostic processes of women with Ovarian Masses (the AROMA pilot study)

Valentina Chiappa¹ · Giorgio Bogani¹ · Matteo Interlenghi² · Christian Salvatore³ · Francesca Bertolina¹ · Giuseppe Sarpietro¹ · Mauro Signorelli¹ · Isabella Castiglioni⁴ · Francesco Raspagliesi¹



Using rADioMics and machine learning with ultrasonography for the differential diagnosis of myometrial tumors (the ADMIRAL pilot study). Radiomics and differential diagnosis of myometrial tumors

V. Chiappa^{A,*}, M. Interlenghi^B, C. Salvatore^B, F. Bertolina^A, G. Bogani^A, A. Ditto^A, F. Martinelli^A, I. Castiglioni^{C,1}, F. Raspagliesi^{A,1}

^A Gynecologic Oncology, Fondazione IRCCS Istituto Nazionale Tumori di Milano, Italy

^B Derynace Technologies S.p.A., Milano, Italy

^C Dipartimento di Fisica G. Occhialini, University of Milan-Bicocca, Milano, Italy

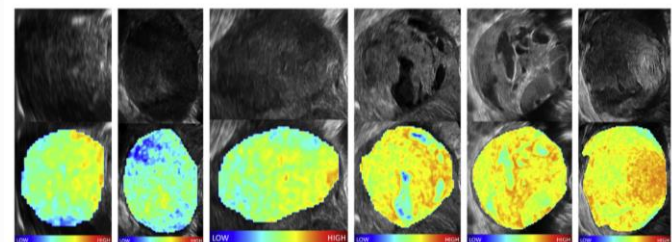
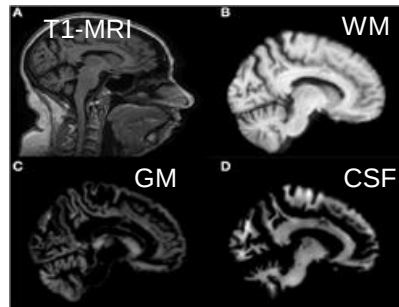


Fig. 2. Local spatial distribution of the feature Entropy evaluated on the Gray-Level Co-occurrence Matrices: Sarcoma (2 a,b,c) and Myoma (2 d,e,f).

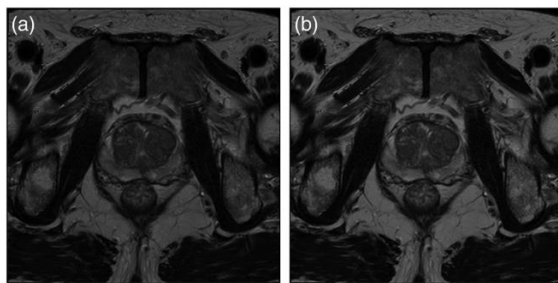
New disease biomarkers from imaging for diagnosis

L2. Imaging systems/ multimodal image processing and decision support systems⁸

Selecting features from expert-domain (developing tissue segmentation methods)



Developing signal/image pre-processing from expert-domain (developing image harmonization methods)



No image normalization

Nyul normalization

Table 3 Evaluating the effect of pre-processing on classification and Segmentation tasks in terms of validation accuracy. Bolded values indicate the improved results.

Task	Image type	Criteria	With pre-processing (%)	No pre-processing (%)
Classification	MR	Accuracy	73.30	68.74
	CT	Accuracy	82.28	77.72
Segmentation	MR	Mean Abs. Err.	2.73	47.64
		Dice	98.64	81.74
	CT	Mean Abs. Err.	3.68	19.99
		Dice	98.25	95.25

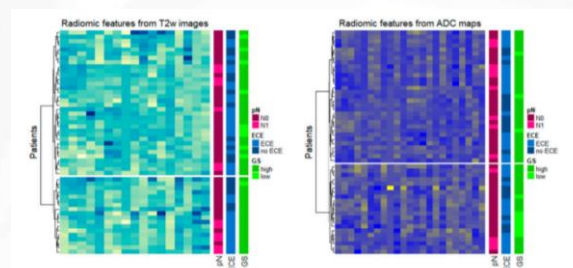
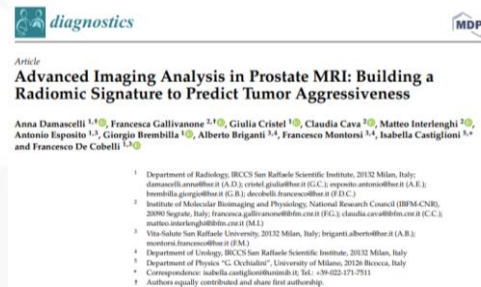
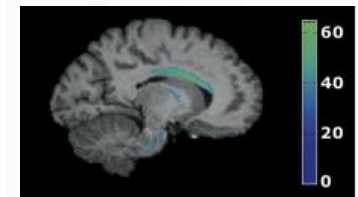
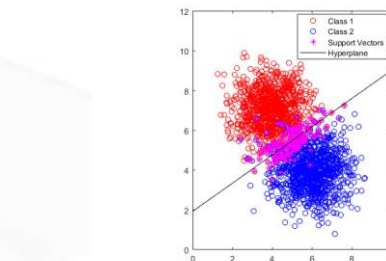
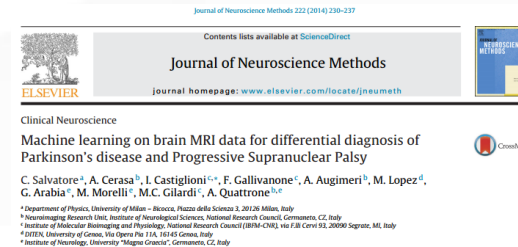
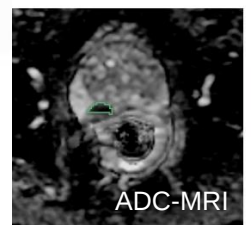


Figure 2. Radiomic patterns obtained on T2w images (left) and ADC maps (right); the dendrogram at the left side represents the patient's grouping obtained from the clustering procedure; annotation at the right side represent the distribution of Gleason Grade Group (GGG), extracapsular extension (ECE), and nodal stage (pN) status in the groups.

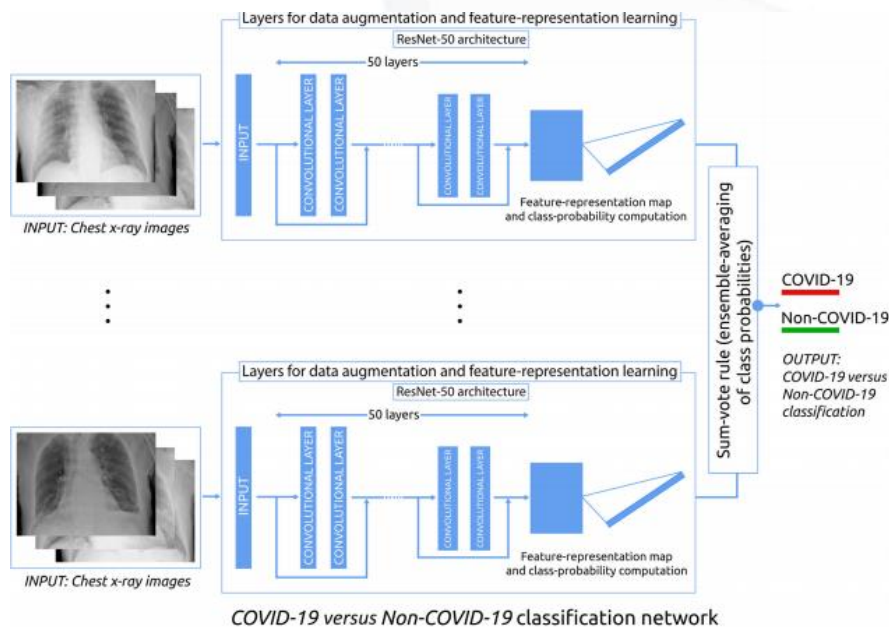
region growing segmentation



New disease biomarkers from imaging for prognosis

L2. Imaging systems/ multimodal image processing and decision support systems⁹

Developing Computer Aided Diagnosis (CAD)/ Decision Support Systems (DSS) for intensive/urgent/critical care



Castiglioni et al. *European Radiology Experimental* (2021) 5:7
<https://doi.org/10.1186/s41747-020-00203-z>

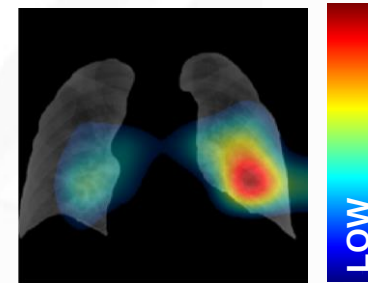
European Radiology
Experimental

ORIGINAL ARTICLE

Open Access

Machine learning applied on chest x-ray can aid in the diagnosis of COVID-19: a first experience from Lombardy, Italy

Isabella Castiglioni^{1,2,†}, Davide Ippolito^{3,†}, Matteo Interleghi³, Caterina Beatrice Monti⁴, Christian Salvatore^{1,6,*}, Simone Schiaffino⁷, Annalisa Polidori⁸, Davide Gandola⁷, Cristina Mesa^{8,9} and Francesco Sardaneli^{8,7}



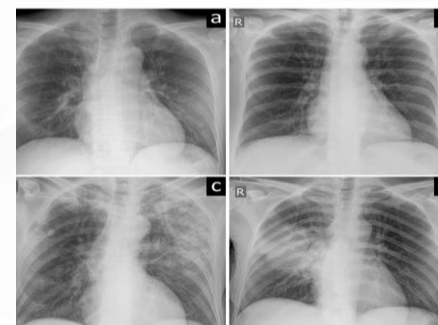
diagnostics

MDPI

Article Artificial Intelligence Applied to Chest X-ray for Differential Diagnosis of COVID-19 Pneumonia

Christian Salvatore^{1,2,*}, Matteo Interleghi^{3,†}, Caterina B. Monti^{4,†}, Davide Ippolito³, Davide Capra³, Andrea Cuzzi⁵, Simone Schiaffino^{7,8}, Annalisa Polidori⁸, Davide Gandola⁷, Marco Ali^{9,10}, Isabella Castiglioni^{7,8,*}, Cristina Mesa^{8,9} and Francesco Sardaneli^{8,7}

1. Department of Science, Technology and Society, Scuola Universitaria IU30, Istituto Universitario di Studi Superiori, Piazza della Vittoria 15, 27100 Pavia, Italy; salvatore@diagnostica.it
 2. DeepFlow Technologies S.R.L., via Conservatore 12, 20122 Milano, Italy; interleghi@deepflowtech.com (M.I.); polidori@deepflowtech.com (A.P.)
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 5. Unit of Radiology, IRCCS Policlinico San Donato, Via Mantovani, 50, 20097 San Donato Milanese, Italy; schiaffino@asstmonza.it
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 7. Department of Physics, Università degli Studi di Milano-Bicocca, Piazza della Scienza 3, 20126 Milano, Italy; francesco.sardaneli@unimi.it
 8. Department of Biomedical Imaging and Physiology, Consiglio Nazionale delle Ricerche, Via Fratelli Cervi 93, 20090 Segrate, Italy
 9. School of Medicine and Surgery, Università degli Studi di Milano-Bicocca, Piazza dell'Ateneo Nuovo 1, 20126 Milano, Italy; cristina.mesa@unimi.it
 10. Fondazione Scintec, Università degli Studi di Milano-Bicocca, Palazzina Cicchione—Via Pergolesi 35, 20090 Monza, Italy
- * Correspondence: isabella.castiglioni@unimi.it
- † C. Salvatore, M. Interleghi, and C. B. Monti contributed equally to this paper.



Covid-19 pneumonia

no pneumonia

other viral pneumonia

Bacterial pneumonia

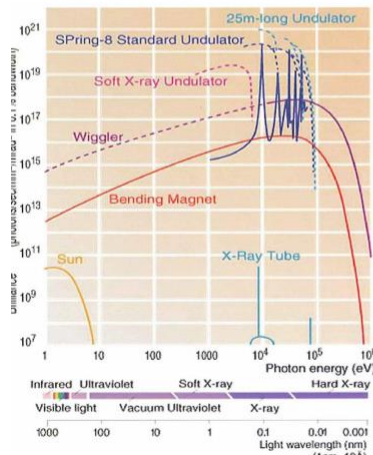
Fast automatic diagnosis and differential diagnosis

L3. Imaging infrastructures and automatic classification

X-ray Phase Contrast Imaging (PCI) at Biomedical beamline (ID17) of the European Synchrotron (ESRF, France)

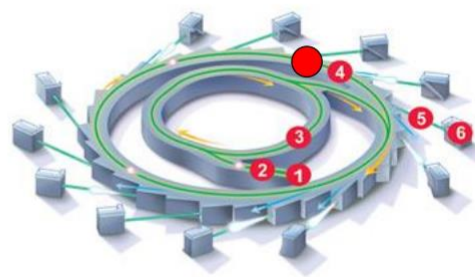
- image contrast derives from the perturbations of the X-ray wave-front induced by the presence of an object along its propagation path, in addition to the x-ray absorption exploited in conventional radiography.
- superior image contrast and sensitivity with respect to standard X-ray attenuation (soft tissues)

SR is a very intense source of radiation from Infra Red to hard X-rays

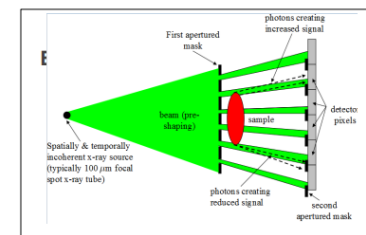
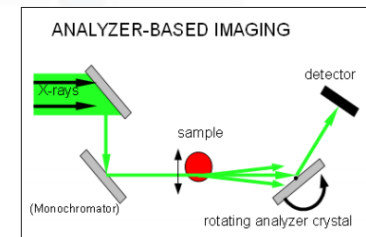
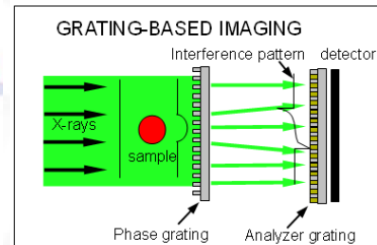
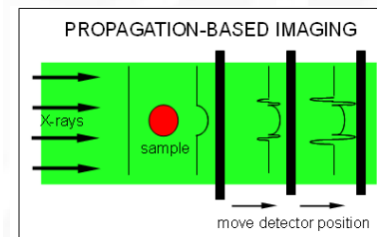


SR is produced by electrons with relativistic energies circulating in a ring

Synchrotron machine scheme



1 injector 2 transfer line 3 booster
4 storage ring 5 beamline 6 exp. station



The detection systems include a portfolio of detectors to perform multiscale imaging, with pixel size from $0.65 \times 0.65 \mu\text{m}^2$ (PCO.Edge.5.5 + 10X optics) to $75 \times 75 \mu\text{m}^2$ (Eiger2-CdTe single photon counting)

High-contrast and sensitive images at quasi-histological resolution

L3. Imaging infrastructures and automatic classification

X-ray PCI for clinical diagnosis from CT to histology image resolution

IOP PUBLISHING
J. Phys. D: Appl. Phys. 46 (2013) 494007 (12pp)

JOURNAL OF PHYSICS D: APPLIED PHYSICS
doi:10.1088/0022-3727/46/49/494007

Phase-contrast x-ray imaging of the breast: recent developments towards clinics

P Coan^{1,2}, A Bravin³ and G Tromba⁴

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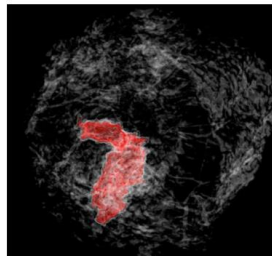
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High-resolution, low-dose phase contrast X-ray tomography for 3D diagnosis of human breast cancers

Yunzhe Zhao¹, Emmanuel Brun^{1,2}, Paola Coan^{1,2}, Zhifeng Huang³, Aniko Sztróky⁴, Paul Claude Diemoz¹, Susanne Liebhardt¹, Alberto Mittone¹, Sergei Gasikov¹, Jianwei Miao^{1,2}, and Alberto Bravin^{1,2}



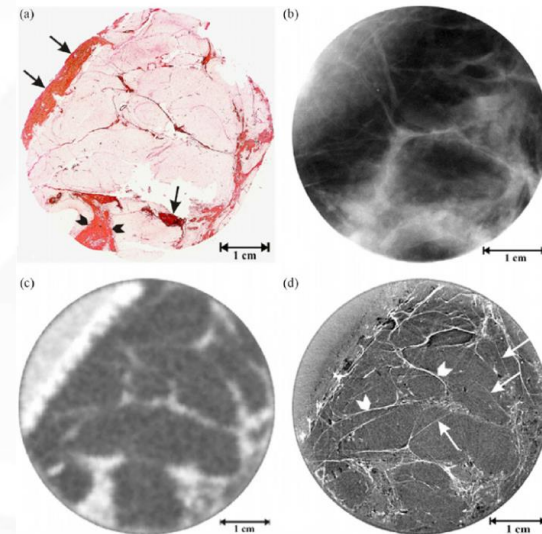
PNAS | November 6, 2012 | vol. 109 | no. 45 | 18293

Phase contrast CT
+ iterative reconstruction method

Dose = 2.0 ± 0.1 mGy
25 times dose saving vs clinical
breast CT at same resolution

histology eusina image

MG image



CT image

PCI image

Radiology

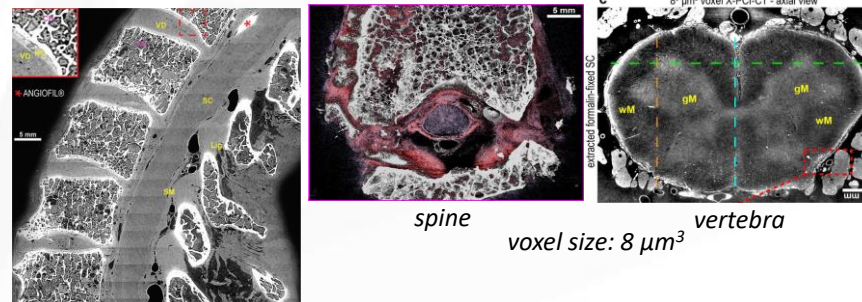
ORIGINAL RESEARCH • MUSCULOSKELETAL IMAGING

High-Spatial-Resolution Three-dimensional Imaging of Human Spinal Cord and Column Anatomy with Postmortem X-ray Phase-Contrast Micro-CT

Giacomo E. Barbone, MSc • Alberto Bravin, PhD • Alberto Mittone, PhD¹ • Sergio Grossi, MD • Jens Rieke, MD • Guido Cavaletti, MD • Valentin Djonov, MD • Paola Coan, PhD

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Supported by the Deutsche Forschungsgemeinschaft (Cluster of Excellence)-Munich Center for Advanced Photonics (EXE158), International Max Planck Research School of Advanced Photon Science (IMPRS-APS), and European COST Action CA16122 in the form of a networking STSM Travel Grant.



spine

voxel size: $8 \mu\text{m}^3$

vertebra

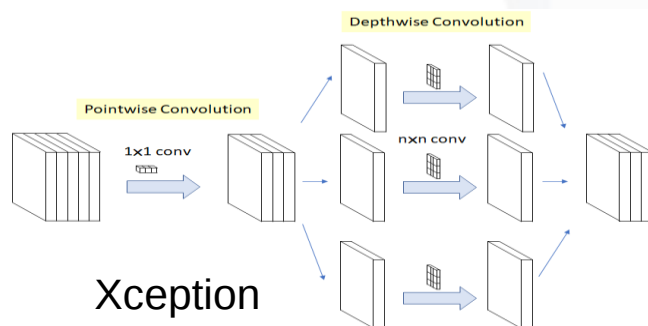
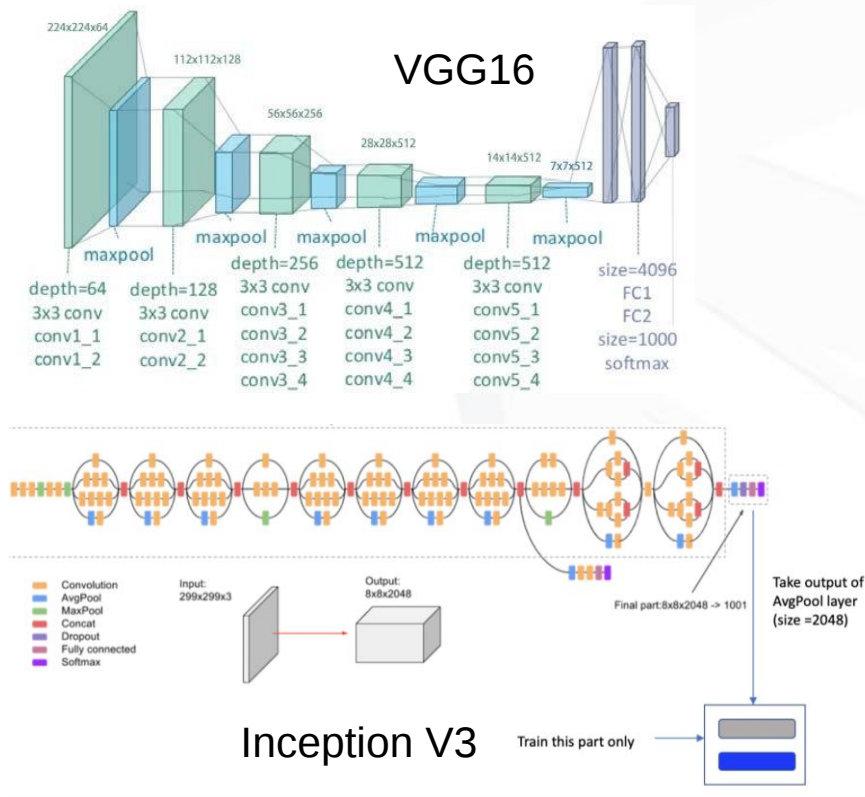
bony vertebral bodies and soft-matter structures within spinal cords, vertebral disks and spinal fat and muscles, voxel size: $46 \mu\text{m}^3$



Better lesion detectability and interpretation

L3. Imaging infrastructures and automatic classification

X-ray PCI combined with AI to speed and improve image reading



scientific reports

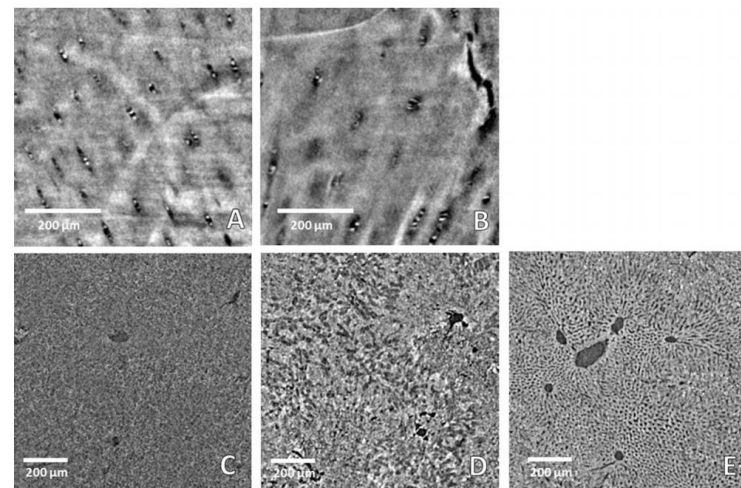
OPEN

Convolutional neuronal networks combined with X-ray phase-contrast imaging for a fast and observer-independent discrimination of cartilage and liver diseases stages

Johannes Stroebe¹, Annie Horng^{1,2}, Marco Armbruster², Alberto Mittone^{3,4}, Maximilian Reiser², Alberto Bravin² & Paola Coan^{1,2,3,5}

sagittal PCI CT images of a healthy cartilage sample

osteoarthritic cartilage sample with a small crack of the tissue visible on the right side



Healthy liver

fibrotic liver

fat liver

automatic diagnostic classification

