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Automated detection of microsatellite status in early colon cancer (CC) using artificial intelligence (AI) integrated infrared (IR) imaging on unstained samples from the AIO ColoPredictPlus 2.0 (CPP) registry study

F. Großerüschkamp¹, S.M. Schörner¹, A-L. Kraeft², D. Schuhmacher³, C. Sternemann⁴, H. Jütte⁴, I. Feder⁴, S. Wisser¹, C. Lugnier², O. Overheu², D. Arnold⁵, C. Teschendorf⁶, L. Mueller⁷, W. Uhl⁸, N. Timmesfeld⁹, A. Mosig³, A. Reinacher-Schick², K. Gerwert¹, A. Tannapfel⁴

¹Center for Protein Diagnostics (PRODI), Dept. of Biophysics, Ruhr-Universität Bochum, Bochum, Germany; ²Dept. of Hematology, Oncology and Palliative Care, St. Josef-Hospital, Ruhr-University Bochum, Bochum, Germany; ³Center for Protein Diagnostics (PRODI), Dept. of Bioinformatics, Ruhr-Universität Bochum, Bochum, Germany; ⁴Institut für Pathologie, Georgius Agricola Stiftung Ruhr - Institut für Pathologie - Ruhr-Universität Bochum, Bochum, Germany; ⁵Oncology, Haematology, Palliative Care Dept., Asklepios Tumorzentrum Hamburg AK Altona, Hamburg, Germany; ⁶Internal Medicine, Medizinische Klinik St. Josefs-Hospital, Dortmund, Germany; ⁷Onkologie UnterEms Leer Emden Papenburg, Onkologische Schwerpunktpraxis Leer-Emden, Leer, Germany; ⁸St. Josef Hospital - Department of Surgery, Ruhr-University Bochum, Bochum, Germany; ⁹Medical Informatics, Biometry And Epidemiology, Ruhr-Universität Bochum, Bochum, Germany

Background: Label-free Quantum Cascade Laser (QCL) based IR imaging combined with AI provides spatially and molecularly resolved alterations in unstained cancer tissue thin sections. For example, molecular alterations such as the microsatellite status (MS) can be classified. To verify the method exemplarily for MS classification, tissue samples from the prospective multicentre AIO CPP registry study were analysed (Nöpel-Dünnebacke et al., ESMO 2020).

Methods: IR images of tissue thin sections taken in 30 min with QCL IR microscopes are classified by AI (convolutional neural networks, CNN). An in-house developed segmenting CNN (U-Net) localizes tumour regions and a second CNN (VGG-Net) classifies the MS. Endpoints were area under curve of receiver operating characteristic (AUC-ROC) and area under precision recall curve (AUPRC).

Results: The multicentre clinical cohort includes 491 pts. (tumour-free 100 / tumour 391). Baseline characteristics including BRAF mutation were equally distributed among test cohorts (Table). The U-Net was verified on 491 pts. (train n=294, test n=100, validation n=97) resulting in an AUC-ROC of 0.99 for the validation dataset. The MS classifier was verified on 391 pts. (train n=245, test n=73, validation n=73) presently reaching an AUC-ROC of 0.83 and an AUPRC of 0.64. Further significant improvement is expected during longer training phase.

Conclusions: QCL IR imaging combined with AI can automatically classify unstained tumour tissue accurately in 30 min with an AUC-ROC of 0.99. Further, it provides concurrently molecular tumour classification, as shown here for the MS. Based on the morphological and molecular alterations encoded in the IR images, AI models will be extended to issues such as prognosis and response prediction to facilitate precision oncology.

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Table: 3850 Cohort details

	Tumour detector (tumour tumour-free)			MS classifier					
	Train	Test	Validation	Train	Test	Validation	Train	Test	Validation
N	240 54	75 25	76 21	71	174	19	54	12	61
Age mean	68 68	70 72	73 72	76	68	72	69	77	68
Sex f/m in %	48/52 54/46	52/48 32/68	53/47 52/48	78/22	41/59	68/32	30/70	75/25	48/53
UICC I (%)	0	0	1 (1)	0	1 (0)	0	0	0	0
II (%)	46 (19)	16 (21)	41 (54)	32 (45)	32 (30)	7 (13)	13 (24)	6 (50)	13 (21)
III (%)	194 (81)	59 (79)	34 (45)	39 (55)	141 (70)	12 (87)	41 (76)	6 (50)	48 (77)
Location left (%)	96 (40)	31 (41)	30 (40)	8 (11)	93 (53)	2 (11)	26 (48)	2 (17)	26 (43)
right (%)	141 (59)	44 (59)	46 (60)	63 (89)	80 (46)	17 (89)	28 (52)	10 (83)	33 (54)
other (%)	3 (1)	0	0	0	1 (1)	0	0	0	2 (3)

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