

A compact Raman Imaging System for Bladder Tissue Analysis

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New modalities in detection and management of bladder tumours for early bladder cancer diagnosis still depend of invasive and time consuming procedures, such as the extraction of biopsies, following a pathological analysis of the samples [1,2]. There is a significant need for non-invasive and rapid methods for in vivo bladder cancer diagnostic. A large number of in vivo and in vitro experiments have demonstrated the ability of Raman spectroscopy to diagnose oncological diseases by providing label-free biochemical information from tissue samples [3-5]. Recent in vivo investigations have shown promising results, but were limited by long acquisition times and high auto-fluorescence backgrounds, due to the nature of the samples [6,7]. A more in-depth characterization of bladder biopsies is needed and new computational methods to preprocess and analyze the measured Raman signals are required [8].

Therefore, this work is aimed to develop a small and compact Raman fiber probe based imaging system to characterize large fresh bladder tissue biopsies. Moreover, we present a comprehensive study on the evaluation of background correction strategies for Raman spectra of highly fluorescent bladder tissues. We compare theoretically and experimentally an instrumental approach, i.e. shifted Excitation difference Raman spectroscopy (SERDS), with a computational approach, i.e. extended multiplicative scattering correction (EMSC), for background correction.

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References

- [1] O. W. Darren, B. K. Beharry, D. Wetherell, N. Papa, M. Weerakoon, A. Sliwinski, M. Brausi, D. Bolton, N. Lawrentschuk, "New Technologies for Diagnosis of Non-Muscle-Invasive Bladder Cancer (NMIBC) and its Management", *J Integr Oncol* 3 (2014)
- [2] R. Colombo, R. Naspro, P. Bellinzoni, F. Fabbri, G. Guazzoni, V. Scattoni, A. Losa, P. Rigatti, "Photodynamic diagnosis for follow-up of carcinoma in situ of the bladder", *Ther Clin Risk Manag* 3, 1003-1007 (2007).
- [3] J. T. Motz, M. Hunter, L. H. Galindo, J. A. Gardecki, J. R. Kramer, R. R. Dasari, M. S. Feld, "Optical fiber probe for biomedical Raman spectroscopy", *Appl. Opt.* 43, 543-553 (2004).
- [4] W. Wang, J. Zhao, M. Short, H. Zeng, "Real-time in vivo cancer diagnosis using Raman spectroscopy", *J Biophotonics* 8, 527-545 (2015).
- [5] K. Kong, C. Kendall, N. Stone, I. Notingher, "Raman spectroscopy for medical diagnostics- From in-vitro biofluid assays to in-vivo cancer detection", *Adv, Drug Deliv. Rev.* 89, 121-134 (2015).
- [6] E. Cordero, F. Korinth, C. Stiebing, C. Krafft, I. Schie, J. Popp, "Evaluation of Shifted Excitation Raman Difference Spectroscopy and Comparison to Computational Background Correction Methods Applied to Biochemical Raman Spectra", *Sensors* 17 (2017).
- [7] R. Gautam, S. Vanga, F. Ariese, S. Umapathy, "Review of multidimensional data processing approaches for Raman and infrared spectroscopy", *EPJ Tech. Instrum.* 2 (2015).
- [8] P. J. Cadush, M. M. Hlaing, S. A. Wade, S. L. Mxarthur, P. R. Stoddart, "Improved methods for fluorescence background subtraction from Raman spectra", *Mater. Sci.* 44, 1587-1595 (2013).