

A novel concept for optical concentration determination of hemoglobin derivatives in non-hemolyzed human blood

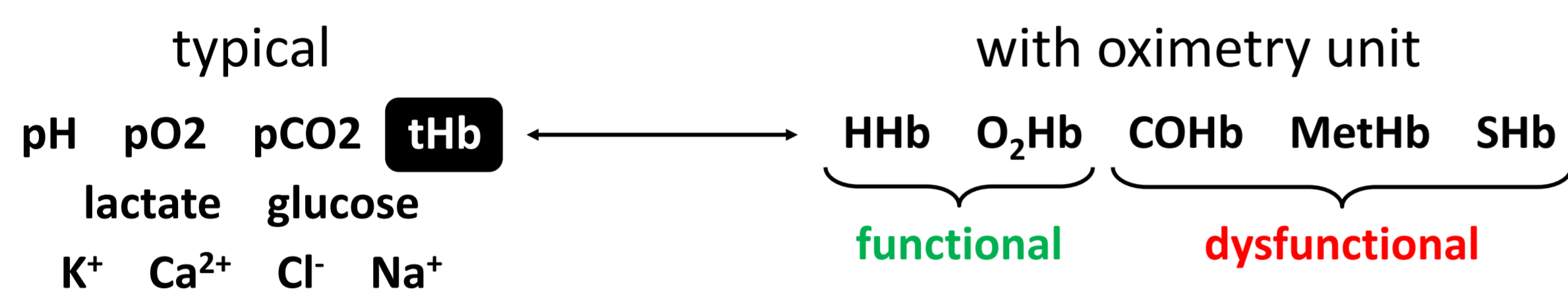
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Introduction

- Knowing the proportions of hemoglobin derivatives in human blood is important in clinical diagnostics due to possible dysfunctional forms.

Parameters in blood gas analysis



- Conventionally, hemolysis is necessary in combination with optical measurements to eliminate light scattering effects by red blood cells.
- A novel approach based on optical measurements in non-hemolyzed whole blood and mathematical modeling is currently being developed.
- The nondestructive approach results in a more flexible analysis process and reduces maintenance costs without the need for consumables.

Objectives

- Design of a measurement setup to determine the optical properties of non-hemolyzed human blood.
- Development of an approach for modeling the dependencies of the optical properties on different condition variables.

Current status

- Case to protect the setup against unwanted external influences, mainly temperature fluctuations.
- Linear slide to simplify calibration process and to increase the reproducibility of measurements.
- Micro bench part for adjusting unscattered transmission intensity allows easy insertion of further modifications (e.g. filters).

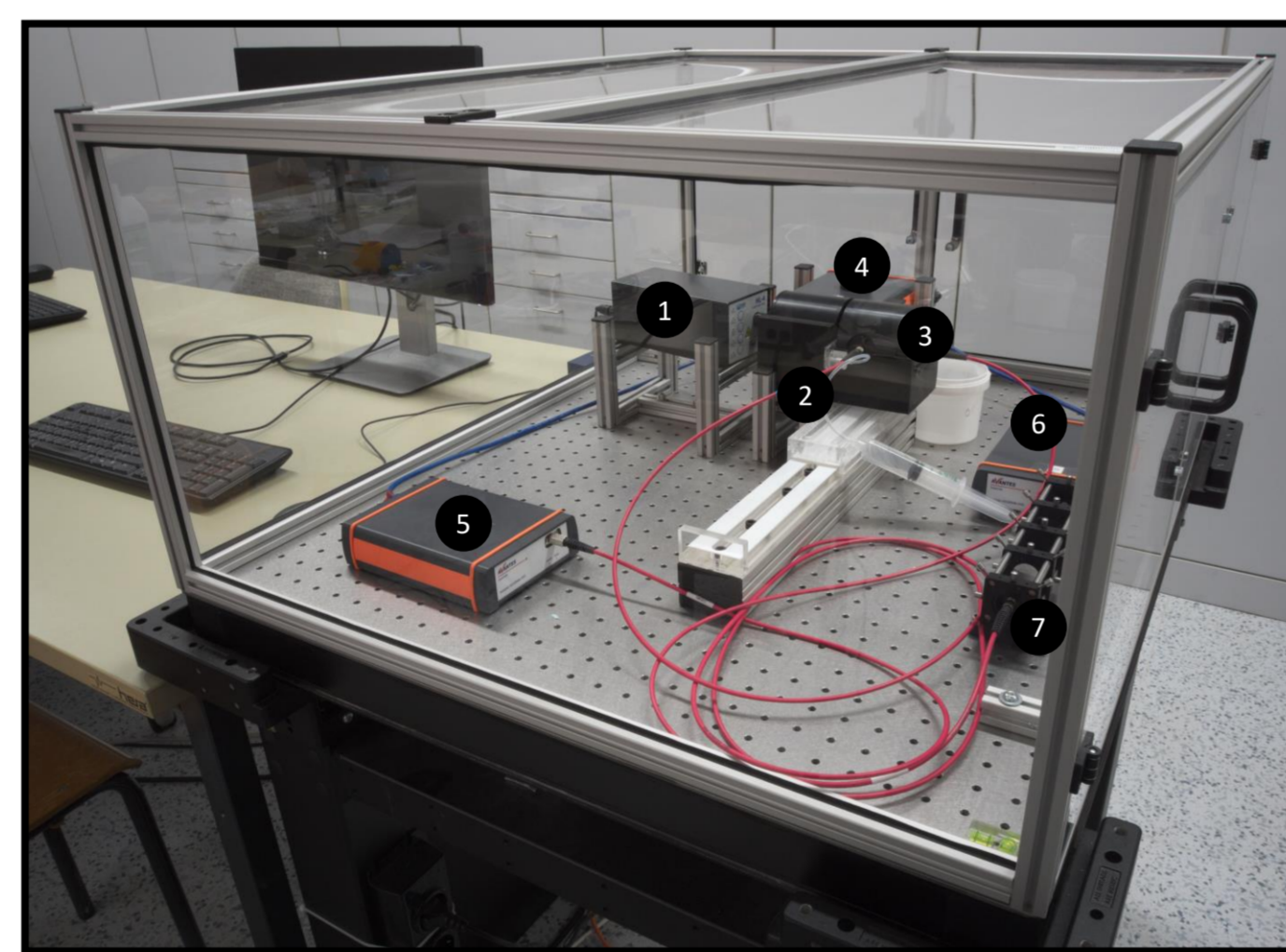


Fig. 3 – Current measurement setup without fluidic circuit components like valves and pump.

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|-------------------------------|-------------------------------|
| 1 Light source | 5 Spectrometer T _d |
| 2 Flow cell | 6 Spectrometer T _u |
| 3 Integrating sphere | 7 Micro bench |
| 4 Spectrometer R _d | |

Outlook

- Evaluation of the measured data with literature data.
- Creation of a database of μ_a , μ_s and g at different condition variables.
- Construction of regression model for the determination of functional and dysfunctional hemoglobin concentrations.

Design concept

- Double-integrating sphere setup with tungsten-halogen light source.
- Measurement of diffuse reflection R_d , diffuse transmission T_d and unscattered transmission T_u in the spectral range from 400 to 1000 nm.
- From R_d , T_d and T_u the absorption coefficient μ_a , scattering coefficient μ_s and anisotropy factor g are calculated using the Inverse Adding-Doubling (IAD) method [1, 2].

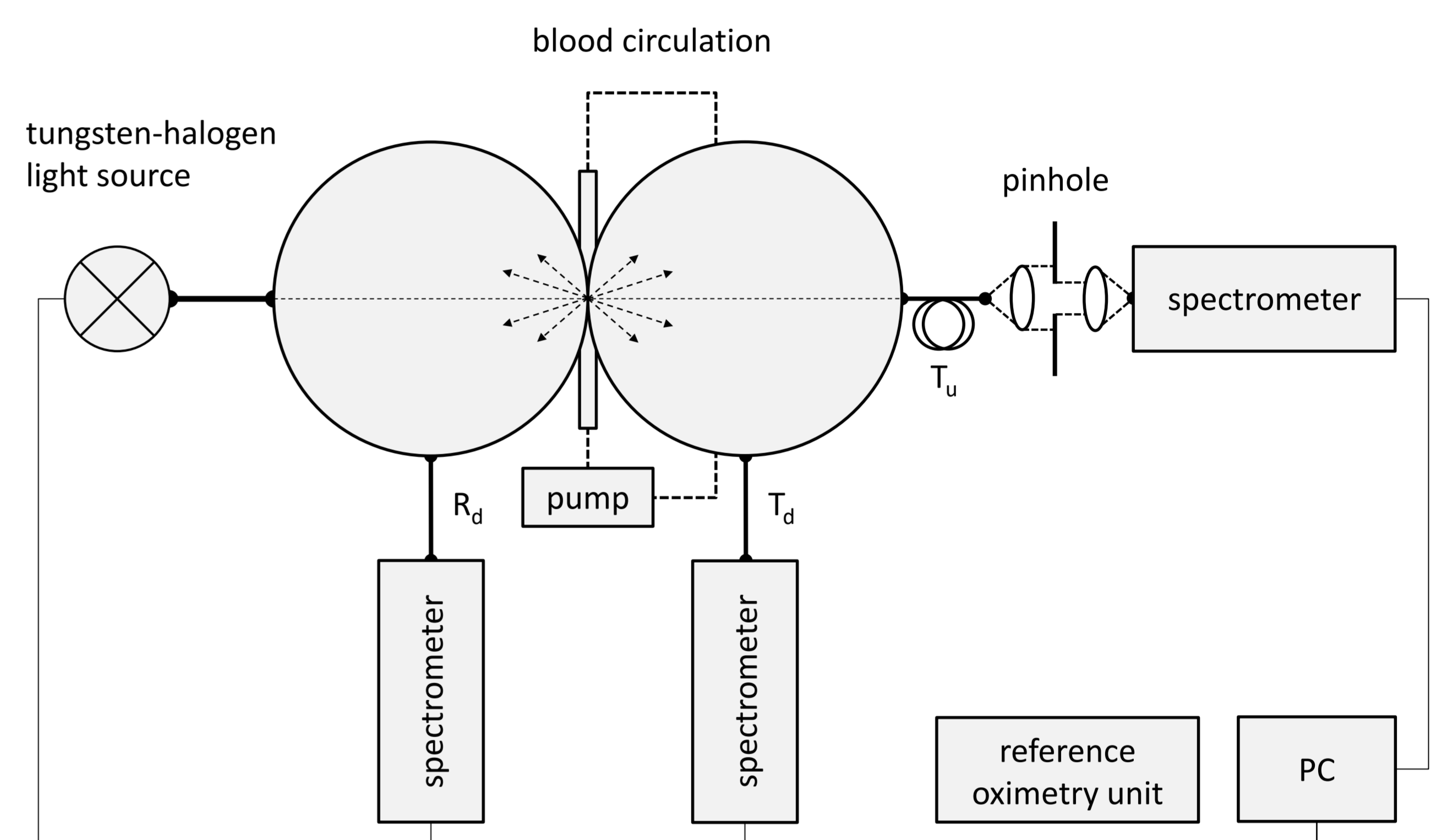


Fig. 1 – A tungsten-halogen light source irradiates a blood sample which continuously flows through a 100 µm optical path length cuvette that is placed between two integrating spheres. Back-scattered and transmitted radiation intensities are measured by spectrometers.

- Determination of optical parameters at sequential variation of condition variables (hematocrit, osmolarity, hemoglobin derivatives and more) through optical measurements on non-hemolyzed human blood [3].
- Shear rate and temperature will be kept at constant values.
- Selection of spectral ranges with good correlation and low cross sensitivity to other variables.
- Construction of a regression model, which combines suitable spectral changes at different optical parameters.

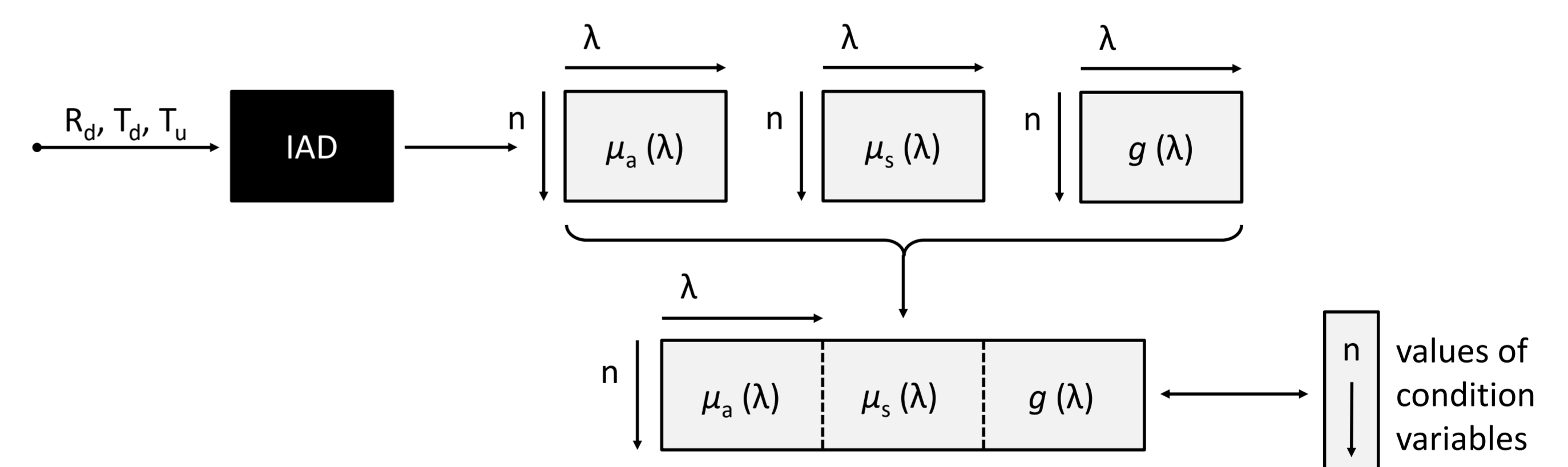


Fig. 2 – Changes in all three optical parameters μ_a , μ_s and g are used to build the regression model.

- Relative quantities provide a well transferable model, e.g. to different measurement geometries.

References

- [1] Scott Prahl, "Everything I think you should know about Inverse Adding-Doubling", March 2011.
- [2] John W. Pickering, Christian J. M. Moes, H. J. C. M. Sterenborg, Scott A. Prahl, and Martin J. C. van Gemert, "Two integrating spheres with an intervening scattering sample", Journal of the Optical Society of America, Vol. 9, No. 4, April 1992.
- [3] André Roggan, Moritz Friebel, Klaus Dörschel, Andreas Hahn, and Gerhard Müller, "Optical properties of circulating human blood in the wavelength range 400-2500 nm", Journal of Biomedical Optics, Vol. 4, No. 1, January 1999.
- [4] Moritz Friebel, André Roggan, Gerhard Müller, Martina Meinke, "Determination of optical properties of human blood in the spectral range 250 to 1100 nm using Monte Carlo simulations with hematocrit-dependent effective scattering phase functions", Journal of Biomedical Optics 11(3), May/June 2006.



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