

## PATATO: A Python Photoacoustic Analysis Toolkit

- <sup>2</sup> Thomas R. Else <sup>1,2</sup>, Janek Gröhl<sup>1,2</sup>, Lina Hacker<sup>1,2</sup>, and Sarah E.
- **3** Bohndiek<sup>1,2¶</sup>
- 1 CRUK Cambridge Institute, University of Cambridge, UK 2 Department of Physics, University of
- $_{\scriptscriptstyle{5}}$  Cambridge, UK  $\P$  Corresponding author

#### DOI: 10.xxxxx/draft

#### **Software**

- Review 🗗
- Repository 2
- Archive ♂

# Editor: Open Journals ♂ Reviewers:

@openjournals

**Submitted:** 01 January 1970 **Published:** unpublished

#### License

Authors of papers retain copyright and release the work under a Creative Commons Attribution 4.0, International License (CC BY 4.0),

## Summary

13

PATATO is a toolkit for processing and analyzing tomographic photoacoustic data. It offers fast implementations of commonly-used methods, including pre-processing, reconstruction, and time-series data analysis, via a convenient command-line interface or Python API. The toolkit utilizes JAX for GPU-accelerated pre-processing and backprojection, and NumPy for less intensive algorithms and easy integration with other pipelines. PATATO can take in data from many different sources, and outputs can be saved in HDF5 format for further analysis and easy transfer to other languages like R or MATLAB. The toolkit is open-source, hosted on GitHub and PyPi, and welcomes contributions, bug-reports, and feedback. Detailed examples and an API reference is available at <a href="https://patato.readthedocs.io/en/latest/">https://patato.readthedocs.io/en/latest/</a>.

## Statement of Need

As a relatively new field, photoacoustic imaging (PAI) often lacks standardization in image processing methods. Despite the numerous algorithms published in literature for reconstruction, spectral analysis, and post-processing, access to the underlying code is limited, making it challenging to use the most current and effective methods. This lack of transparency also hinders reproducibility and slows the adoption of new methods within the community. While other tools for photoacoustic image reconstruction and analysis have been developed, they all lack open-source code for GPU acceleration, and have limited documentation, making it difficult to extend their functionality. PATATO offers an open-source solution with accelerated reconstruction algorithms, end-to-end analysis, an easy-to-use interface, open data format integration, and an extensible structure.

Like many other imaging techniques, PAI requires multiple processing steps, many of which are computationally intensive. A typical pipeline for generating PA images includes filtering and interpolation of time-series data, delay-and-sum reconstruction, and linear spectral unmixing. The first two steps are particularly computationally intensive and require GPU implementations for efficient processing of large datasets, which can sometimes consist of thousands of frames. Additionally, the resulting images are highly sensitive to the parameters used, such as the cut-off frequency for filters, highlighting the need for efficient and transparent implementations of these algorithms to ensure consistent biological conclusions across datasets. In PATATO, these computationally intensive algorithms are implemented using JAX, a high-performance numerical computing toolkit, enabling GPU acceleration and consistent implementations between CPU and GPU. Additionally, PATATO stores all reconstruction parameters in the HDF5 file, making it easy to reproduce the precise analysis.



## Example Datasets

- Some example datasets are included in PATATO to demonstrate its key features and to
- allow benchmarking of different algorithms. Two devices were used to collect this data, a
- 42 circular-array pre-clinical system (MSOT inVision, iThera Medical) and a handheld clinical PA
- 43 system (MSOT Acuity, iThera Medical). The properties of the two systems are summarised
- below for reference.

	Pre-Clinical	Clinical
Transducer Number	256	256
Sampling Frequency (MHz)	40	40
Transducer Coverage (degrees)	125	270
Wavelengths (nm)	660 to 1300	660 to 1300
<b>Example Datasets</b>	Phantom	Phantom scan and time-series
		scans: Oxygen-Enhanced (OE),
		and Dynamic Contrast-Enhanced
		(DCE)

#### 45 Phantom Data

Phantom imaging data is provided for both the clinical and pre-clinical imaging systems.

#### Oxygen Enhanced and Dynamic Contrast Enhanced Data

- Examplar data is provided for oxygen enhanced (OE) and dynamic contrast enhanced (DCE)
- 49 imaging. In the OE experiment, a mouse was imaged continuously under inhaled anaesthesia
- 50 (1-2% isofluorane) while breathing air. After five minutes, the breathing gas was changed
- to 100% oxygen. Consequently, there is an increase in the blood oxygenation, which can be
- measured using PAI. In the DCE experiment, a bolus of the PA contrast agent indocyanine
- green (ICG) was injected while imaging. The results of the analysis procedure provided by
- PATATO are shown below. These datasets can be accessed directly from PATATO, and this
- <sub>55</sub> analysis can be reproduced, as described in the documentation.



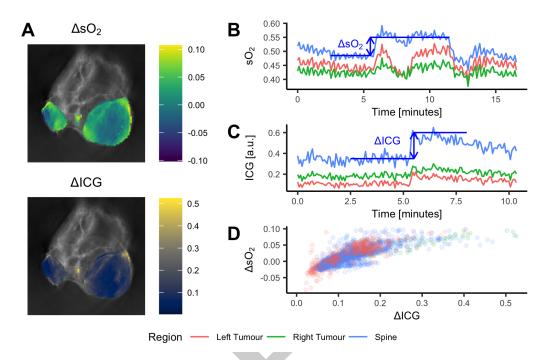


Figure 1: An example of the time-series data analysis methods applied to one scan. A: example images of the increase in blood oxygenation after the oxygen challenge  $(\Delta sO_2)$  and increase in unmixed ICG signal after the ICG bolus injection  $(\Delta ICG)$ . B and C: Plots of the mean tumour  $sO_2$  and ICG respectively over time. Also showing the calculation procedure for the corresponding  $\Delta sO_2$  and  $\Delta ICG$ . D: Plot of the pixel-wise values showing a strong positive correlation between  $\Delta sO_2$  and  $\Delta ICG$ .

#### Mouse Imaging Methods

All animal procedures were conducted in accordance with project (PE12C2B96) and personal licenses (I33984279) issued under the United Kingdom Animals (Scientific Procedures) Act, 1986, and were approved locally under compliance form number CFSB2022. PC3 cell suspensions were injected into both flanks of nude (BALB/C) mice,  $1.5 \times 10^6$  cells in  $100\mu l$  containing a 1:1 mixture of PBS and matrigel. Mice were imaged and culled once the tumours had reached a size of approximately 1cm in diameter. Detailed methods for this procedure have been published previously (Tomaszewski et al., 2018).

## Acknowledgements

We'd like to thank a number of members of our lab for their helpful feedback, particularly Mariam-Eleni Oraiopoulou, Ellie Bunce and Thierry Lefebvre.

#### References

Tomaszewski, M. R., Gehrung, M., Joseph, J., Gonzalez, I. Q., Disselhorst, J. A., & Bohndiek, S. E. (2018). Oxygen-enhanced and dynamic contrast-enhanced optoacoustic tomography provide surrogate biomarkers of tumour vascular function, hypoxia and necrosis. *Cancer Research*, 78, canres.1033.2018. https://doi.org/10.1158/0008-5472.CAN-18-1033