

¹ PICNIC: an open-source Python library for ² preprocessing of dynamic Positron Emission ³ Tomography (PET) brain imaging data

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Software

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⁹ Summary

¹⁰ PICNIC (Pipeline Initializing Container for Neuro-Imaging Computations) is an open-source
¹¹ Python-based coding library that includes modular wrappers for most standard preprocessing
¹² steps for quantification of Positron Emission Tomography (PET) brain imaging data. PICNIC
¹³ uniquely allows full, transparent modular control over the selection of strategy/software package
¹⁴ and associated parameters for each preprocessing step throughout the pipeline. Therefore,
¹⁵ the user is provided with the flexibility to select different methods depending on the target of
¹⁶ interest and/or radiotracer properties, study population, or other preferences. These settings
¹⁷ can be saved as a text-based “input deck” file to support consistent preprocessing across scans,
¹⁸ participants, and projects. Furthermore, PICNIC supports freezing software versions within
¹⁹ Docker containers, ensuring robust reproducibility.

²⁰ PICNIC does not require Brain Imaging Data Structure (BIDS) source-level data, but instead
²¹ takes reconstructed PET static or dynamic data in industry standard imaging filetypes (e.g.,
²² DICOM, Nifti, ANALYZE), and converts them to be BIDS-compliant ([Knudsen et al., 2020](#)).
²³ PICNIC employs the most commonly used brain imaging software packages, including Advanced
²⁴ Normalization Tools ([Avants et al., 2008](#)), Analysis of Function NeuroImages ([Cox, 1996](#)),
²⁵ FSL ([Jenkinson et al., 2012](#)), FreeSurfer ([Fischl, 2012](#)), dcm2niix ([Li et al., 2016](#)), and SPM
²⁶ ([Laboratory, 2020](#)). PICNIC was designed with PET in mind, but includes steps involving data
²⁷ from multiple imaging modalities, most prominently structural magnetic resonance imaging
²⁸ (sMRI). PICNIC preprocessing capabilities include image reorientation (to force all images
²⁹ into a standard orientation regardless of scanner defaults), brain extraction (to perform
³⁰ skull-stripping of associated sMRI images), tissue segmentation (to classify sMRI voxels
³¹ within the brain boundary as grey matter vs. white matter vs. cerebrospinal fluid), automatic
³² anatomical parcellation (to identify regions of interest using any available atlas/parcellation),
³³ post-reconstruction rigid-body motion-correction (to correct inter-frame participant motion over
³⁴ the duration of the scan), PET-sMRI co-registration (with 20 different PET-sMRI rigid-body
³⁵ transformations that are automatically ranked based on mutual information), and extraction of
³⁶ time-activity-curves (curves that represent the PET signal over the duration of the scan in
³⁷ delineated regions of interest or in each brain voxel).

³⁸ PICNIC can be run from the command line or through the provided graphical user interface.
³⁹ An html summary output with interactive quality control features allows the user to inspect,
⁴⁰ comment, and approve/disapprove each module for each participant upon completion. Pre-
⁴¹ loaded templates have been provided for commonly used workflows. The user can add, remove
⁴² or edit pre-processing steps to fully customize their pipeline with regards to their dataset.

43 PICNIC currently supports over 100,000 module/parameter combinations.

44 PICNIC is designed for in vivo brain investigations by expert PET researchers, and has already
45 been applied to preprocess data for recent scientific publications from our Lab (Bartlett,
46 Patil, et al., 2023, 2024; Bartlett, Zanderigo, Stanley, et al., 2023; Bartlett, Zanderigo,
47 Ogden, et al., 2023; Bartlett, Herzog, et al., 2024; Bartlett, Sublette, et al., 2024; Graves
48 et al., 2024; Herzog et al., 2024, 2025; Mann et al., 2022; Matheson et al., 2024; Miller et
49 al., 2022) (<https://www.columbiapsychiatry.org/research-labs/brain-imaging-lab>). PICNIC's
50 modular nature enables extensive customization of preprocessing for a variety of brain PET
51 studies, and its design may allow future extension to preprocessing of data from imaging
52 modalities other than PET. The source code for PICNIC is available at the following link:
53 <https://github.com/bil-mind-nyspi/PICNIC>.

54 Statement of need

55 PET is a valuable tool used in research and clinical settings to noninvasively image the
56 human brain in vivo. PET can quantify up to nanomolar levels of specific components of
57 brain metabolic and neurochemical processes, thus providing information on the distribution
58 of specific biological targets, such as receptors and enzymes, or on the uptake of specific
59 compounds into the brain, like glucose and polyunsaturated fatty acids. Rigorous preprocessing
60 of the raw data captured by the PET scanner (Norgaard et al., 2019) is key to obtaining the
61 most accurate estimates of the distribution of a biological target or uptake of a substance
62 using PET. Although a few software packages already exist (Funck et al., 2018; Karjalainen
63 et al., 2020; Routier et al., 2021) that perform most of the required preprocessing steps, a
64 fully open-source library that can modularly and flexibly combine appropriate preprocessing
65 strategies depending on the study design at hand is still missing.

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