

Developing an Integrated Multimodal Platform for the POETIC Breast Cancer Trial

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*: These authors contributed equally

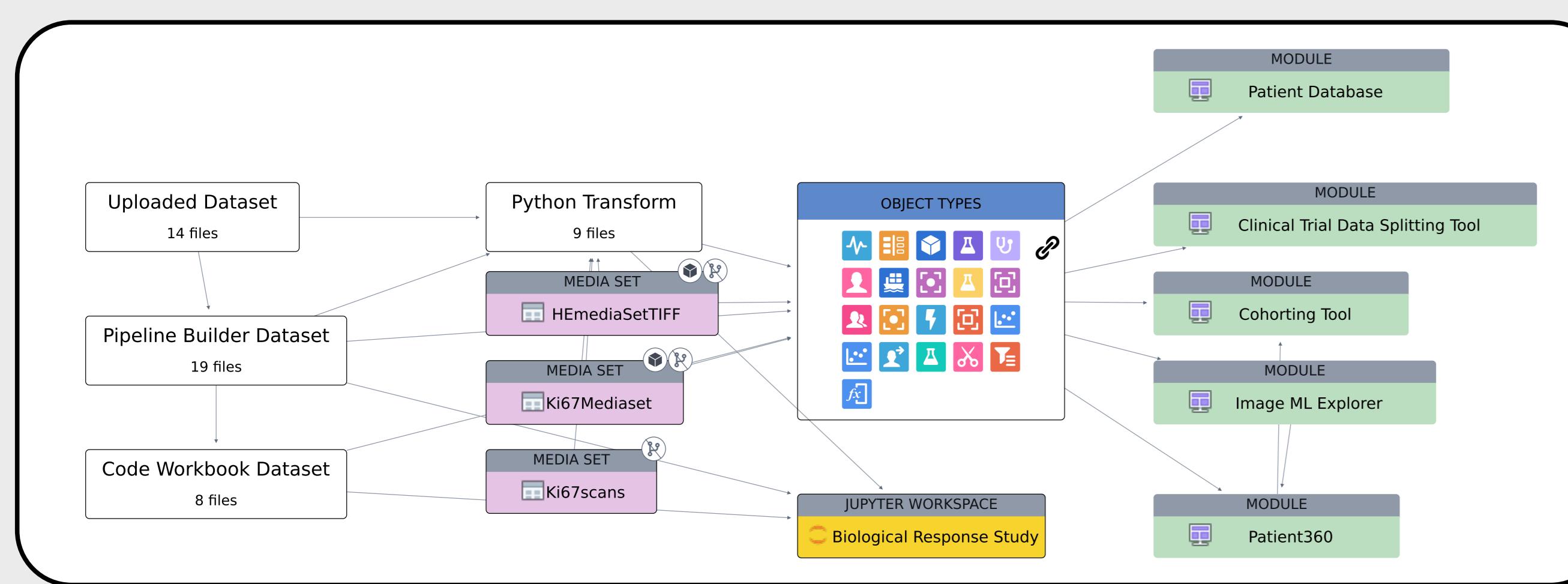
Peri-Operative Endocrine Therapy for Individualised Cared

The POETIC trial (NCT02338310) evaluated peri-operative endocrine therapy in hormone receptor-positive early breast cancer. POETIC represents a unique translational resource in the UK, covering ~ 4,500 patients and combining detailed clinicopathological annotations with extensive multi-omics profiling.

Here we present POETIC-AI, a data integration study* where we have used clinical, molecular (including transcriptomics, genomics, and spatially resolved omics), and digitised histopathology data from the POETIC trial.

To demonstrate the capabilities of POETIC-AI we present a cross-modality study that, using orthogonal measurements of the proliferative marker Ki67, recapitulates proliferative changes as a biological response to intervention.

Building the Graph-based POETIC-AI Interoperable Ecosystem

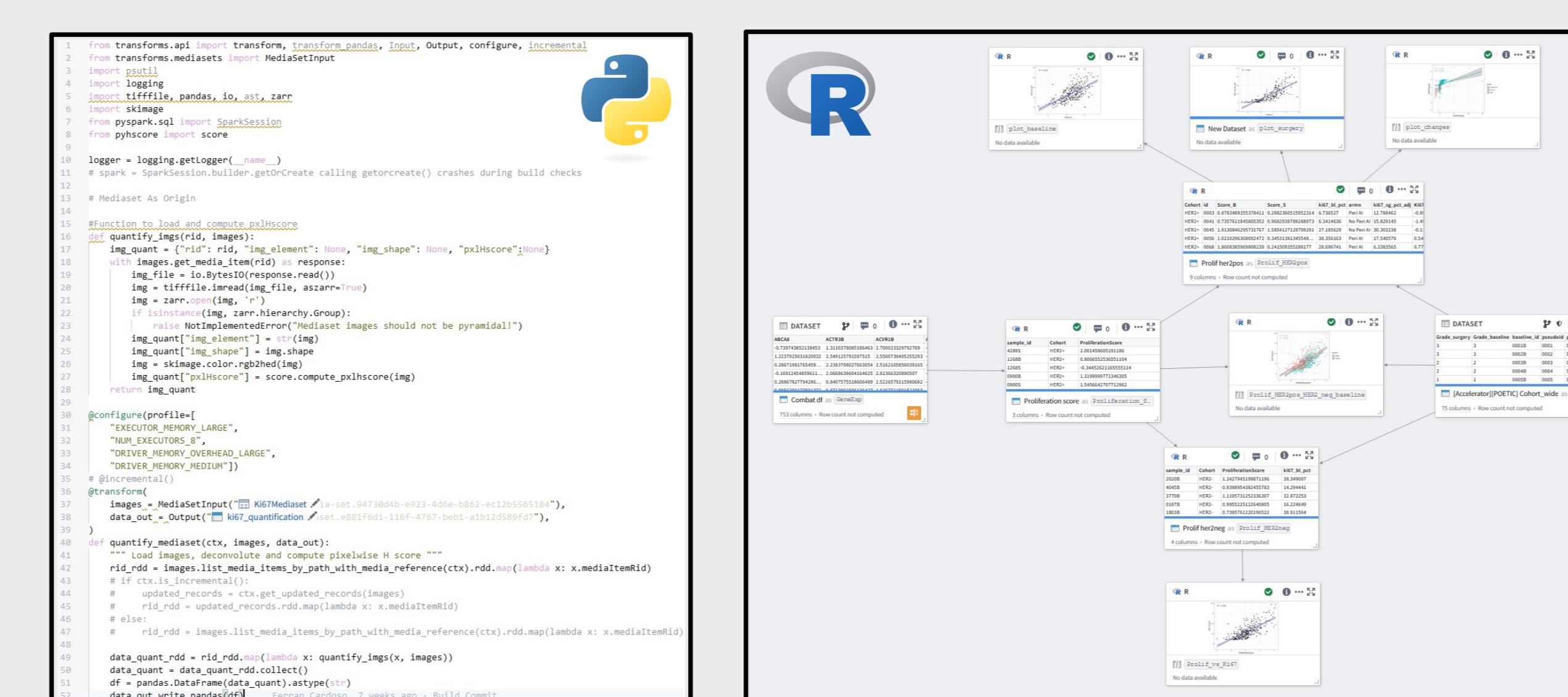
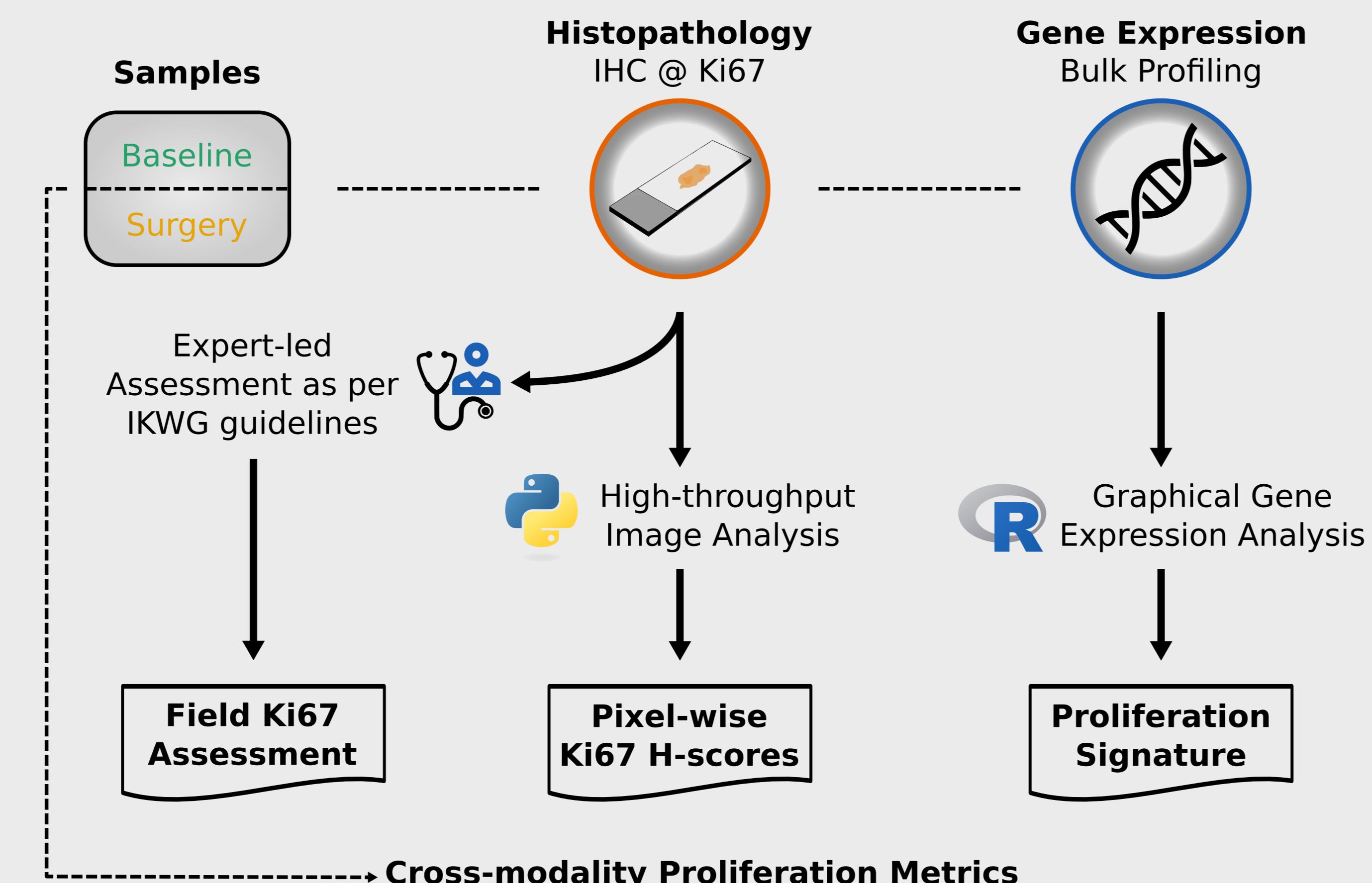


Leveraging the commercial solution Foundry, in POETIC-AI we have developed and implemented analysis, visualisation, and exploration tools across the multiple data modalities. We have built this interconnected platform around a graph-based ontology that unifies clinical records, molecular profiles, and pathology scans within a single interoperable ecosystem with built-in versioning and access control.

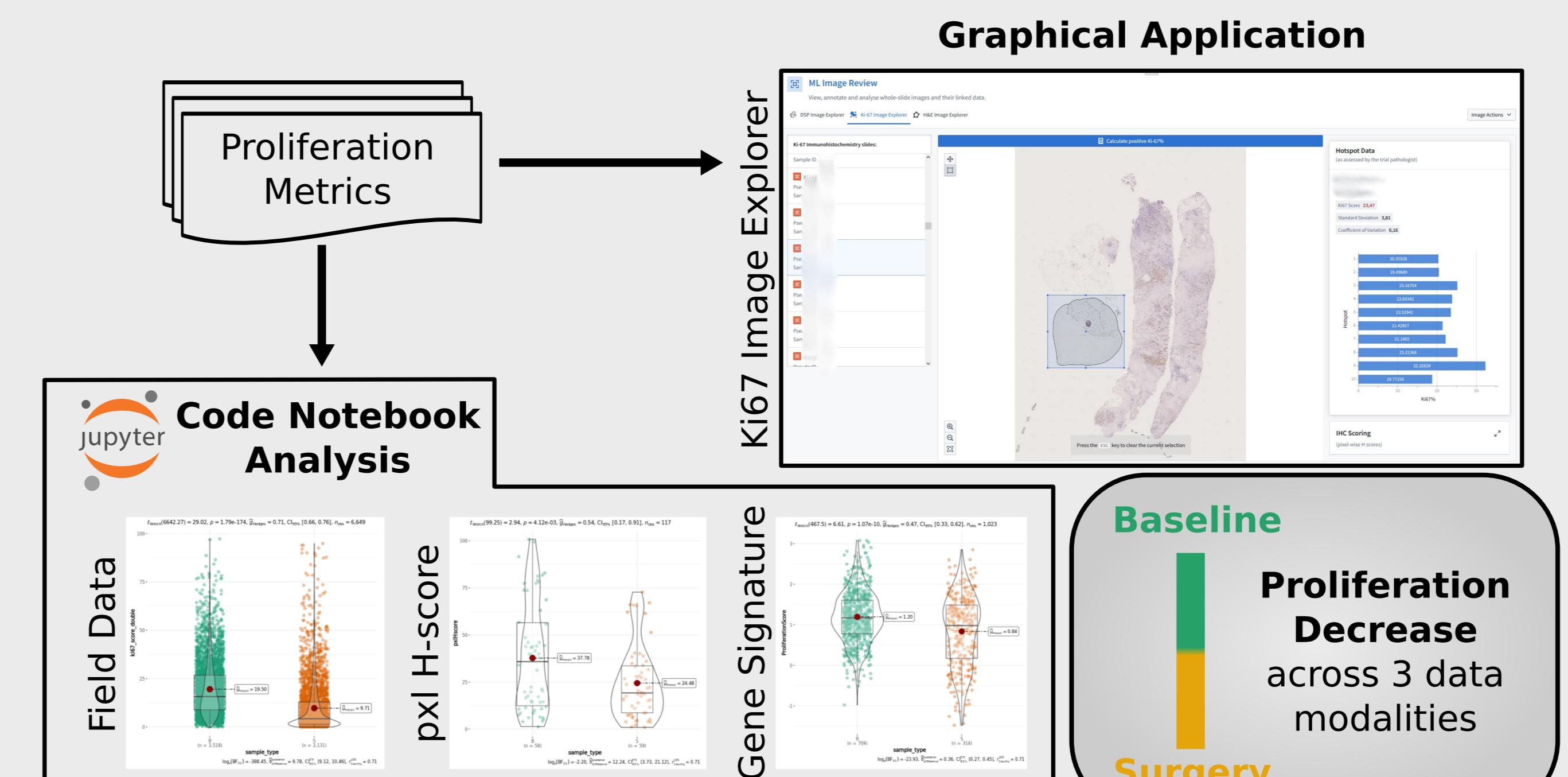
Within this environment we can map relationships among entities such as patient, sample, assays, and derived molecular features. In addition to uploaded datasets and data transform operations, our custom built applications and analyses (e.g. Biological Response Study) are also integrated within the broader ontology. This architecture supports dynamic data linkage, hierarchical querying, and real-time visualisation of multimodal relationships.

Recapitulating the Early Biological Response to Endocrine Therapy [1/2]

Within POETIC-AI, we have implemented analytical pipelines in two complementary forms: interactive graphical workflows and server-executed code-based pipelines. The graphical R-based workflows offer an accessible interface for data exploration and reproducible analysis directly linked to ontology-defined entities, while the Python-based pipelines operate on dedicated compute infrastructure for scalable, distributed processing of large datasets such as whole-slide images. To demonstrate the need for both these forms we have aimed to quantify proliferative changes across modalities to recapitulate an early biological response to intervention. A graphical workflow was used to compute proliferation signatures based on gene expression data from bulk profiling, while a code-based image analysis pipeline is used to quantify IHC targeting the proliferation marker Ki67.

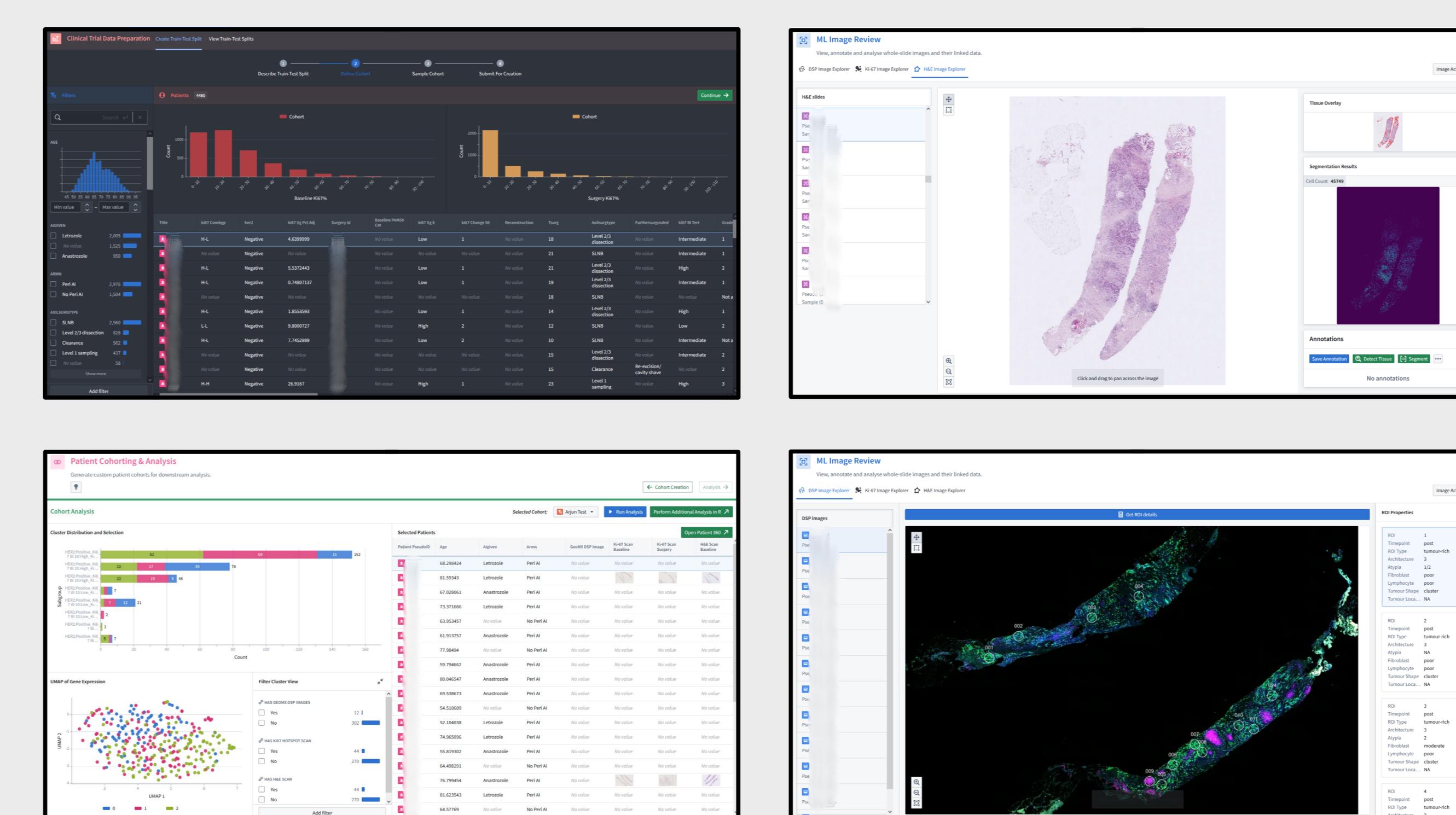


Recapitulating the Early Biological Response to Endocrine Therapy [2/2]



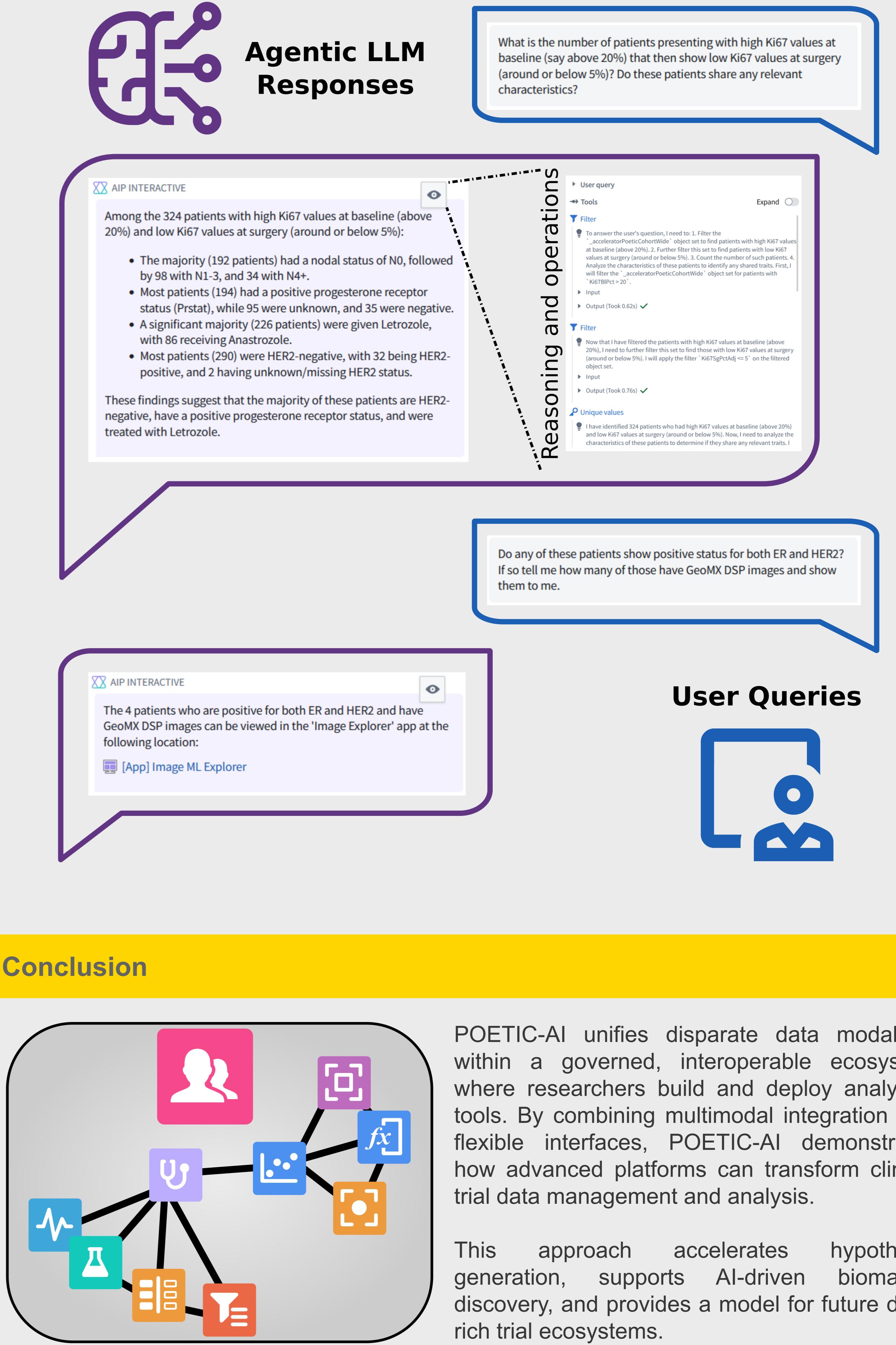
Incorporating field-spot assessments by expert trial pathologists following IKWG guidelines, this multimodal framework enables consistent evaluation of early biological response across molecular and histopathological modalities; recapitulating a decrease in proliferative index between baseline and surgical samples and surfacing the results as a graphical application.

Building Graphical Applications for Accessible Analysis and Exploration



To facilitate accessibility, we have developed a series of graphical applications to support zero-code visualisation, exploration and analysis. These can be used to create cohorts and design splits (for downstream machine learning tasks), perform compositional analysis of the transcriptomic profiles of patients, and visualise histological and spatial modalities; linking digitised images with relevant metadata within POETIC-AI.

Demonstrating the Natural Language Interface Via an Agentic LLM



POETIC-AI unifies disparate data modalities within a governed, interoperable ecosystem where researchers build and deploy analytical tools. By combining multimodal integration with flexible interfaces, POETIC-AI demonstrates how advanced platforms can transform clinical trial data management and analysis.

This approach accelerates hypothesis generation, supports AI-driven biomarker discovery, and provides a model for future data-rich trial ecosystems.

References

- Smith, I. et al. Long-term outcome and prognostic value of Ki67 after perioperative endocrine therapy in postmenopausal women with FCR confirms no conflicts of interest to declare hormone-sensitive early breast cancer (POETIC). *The Lancet Oncology* 11, 1443-1454 (2020). 10.1016/S1470-2045(20)30458-7

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Disclosure Statement

FCR confirms no conflicts of interest to declare

Acknowledgments

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