

# Title

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## Abstract

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**Keywords:** key1 · key2

**Mathematics Subject Classification:** If · Needed

## 1 Introduction

Problem: field of biology, surge of highly multiplexed experimental protocols capable of recovering thousands of cells from which several measurements are obtained. This poses a challenge to analyse this multidimensional data. To do so, people recur to Machine Learning (ML) techniques, in order to investigate the data for structures that have biological relevance.

An ML method that fits the data can detect data structures that can then be challenged for biological relevance, thus helping researchers in their investigation.

Examples (Kara Davis's paper, and Garry Nolan's in general)

Feature selection is one of the desirable features. You run an experiment for several markers to classify cell types, you want to know which are the most relevant that should be retained in a follow up experiment (eventually with lower multiplexing power) to optimise the cell classification.

As cells are often classified by gating (describe gating), RF classification is an automatic routine that mimics gating. Although at present no automatic routine outperforms expert gating, thanks to the similarity with gating RF findings can be integrated, interpreted, verified with more ease from researchers.

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The metrics in output must also be well-defined in unbalanced data, where often rare populations are very important in the investigation.

For these reasons we aim to provide further explainability to the RF classifier by redefining feature importance. Must be theoretically sounding -> proper definition Be explainable -> plots that provide better understanding of the impact of feature to the class prediction in the model, even for non machine learning experts. Useful statistic -> on which researchers can base their feature selection decision to select for the top predicting features, tailored for specific classes of interest.

[RECYCLE] In particular, the recursive thresholding in Random Forest (RF) mirrors cytometry gating to classify data, still underperformed by automatic solution that have difficulties in dealing with biological variability while taking previous field knowledge into account.

Background, Context, State-of-Art

Why we did this

What we did-Paper structure

## 2 Formal Definition

We seek to define a new measure for feature importance that do this as opposed to the what we call Global Importance

## 3 Application to dermatology data

Explain model: heatmap and expression figures Feature importance profile across classes -> high average for globally important features; large standard deviation for features with different impact among classes Comparison with global importance

### 3.1 Detailed model explanation

### 3.2 Feature selection

Feature importance distribution within each class Comparison with features selected via global importance

## 4 Discussion

Recap problem and what we achieve

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