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#### ▶ To cite this version:

Levy Batista, Yaël Kolasa, Thierry Bastogne. i-Cellulo: a SaaS platform for the automatic statistical analysis of cell impedance signals. 8th International Meeting on Statistical Methods in Biopharmacy, SMB 2017, Sep 2017, Paris, France. hal-01670076

HAL Id: hal-01670076

https://hal.science/hal-01670076

Submitted on 21 Dec 2017

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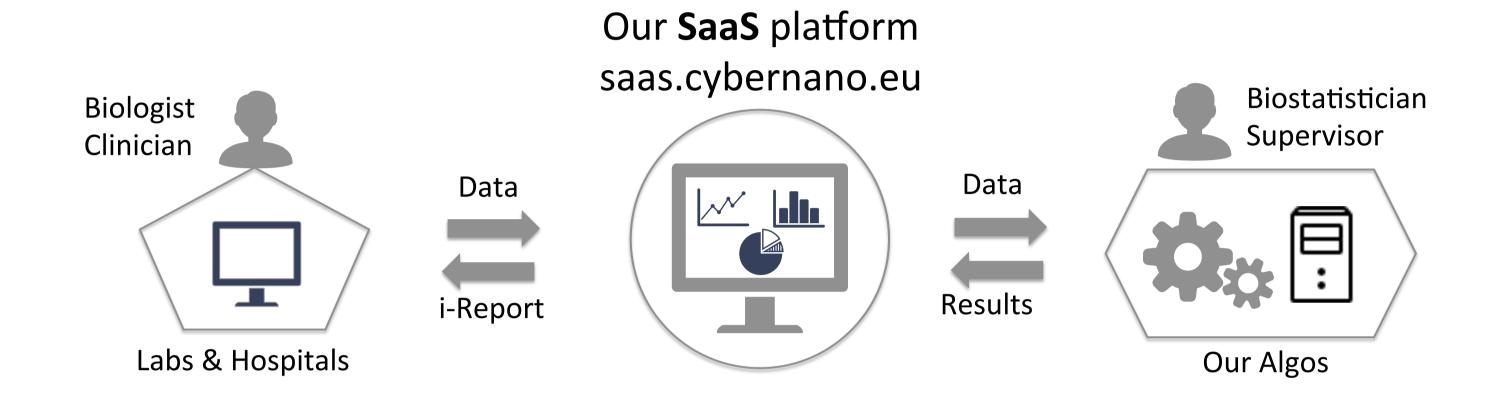




#### i-Cellulo: a SaaS platform for the automatic statistical analysis of cell impedance signals

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<u>Objectives.</u> Label-free methods such as **cell impedance assays** are *in vitro* tests increasingly used in drug development. An indirect difficulty with those technologies is the large amount of kinetic responses to be processed. Our objective is to **automate the processing and analysis** of those data with a web computational server available to all biologists and able to perform multivariate tests, response profile clustering and dynamic AC50 estimation.



# Methods

The proposed solution relies on a SaaS platform in which R-language algorithms have been implemented for the on-line processing of cell impedance signals. Three generic statistical problems are addressed: clustering of response profiles to screen compounds, multivariate testing to compare their activity and AC50 estimation to determine their concentration effects. ANOVA, Kruskall-Wallis and Tuckey's range tests have been implemented for the multivariate testing. Hierarchical clustering based on Singular Spectrum Analysis were used for the non-supervised response profile classification. A Hill's model structure and a maximum likelihood estimator were adopted for the AC50 calculation.

### Results

Hundreds of tests were carried out on real in vitro signals to assess the practical relevance of i-Cellulo for the fast analysis and characterization of anti-cancer activities in early steps of drug development. Results clearly show the ability of this web-based solution to correctly discriminate, classify, compare and rank the anti-cancer responses of tested compounds compared to gold standards.

## Conclusion

With the advent of real-time cell measurement technologies in preclinical tests, new services for the analysis of **high-content data** are needed. **i–Cellulo** is a solution to that challenge and allows biologists to speed up their data analysis and facilitate interpretation of their results.

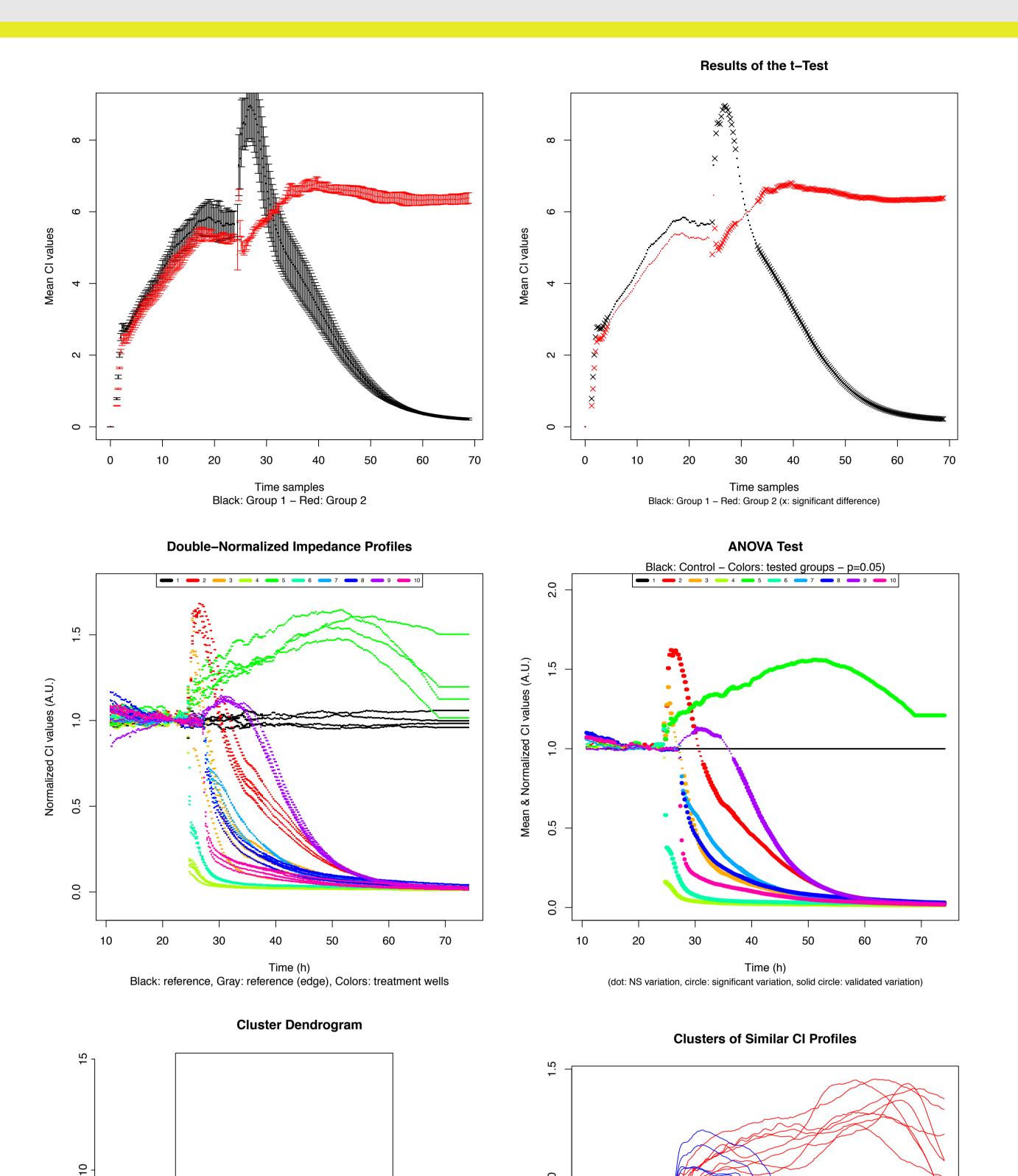


Fig.1: Comparing 2 groups of cell impedance profiles. Left: original CI responses to be compared. Right: t-Test results

Fig.2: Multiple test of CI responses. Left: groups of CI responses to be compared. Right: ANOVA results

Fig.3: Clustering Cell Index Profiles for Screening Applications. Left: dendrogram of culture wells. Right: 3 clusters of response.

Wells

AC50a = 4.68 uM - AC50r = 4.2 uM

Time variation of the IC50 values

9

0 20 40 60 80 100 120

Time (h)

Time (h)

Concentration (uM)

Measured values (red, Proceded values (black), Relative AC50 (green)

(Red): Absolute IC50, (Black); Relative IC50, Solid circles: confident IC50 values

Fig.4: Estimation of the AC50 profiles. Left: AC50 values estimation based on the Hill's model. Right: time profil of the AC50 estimates