dpcReport: POC tests analysis framework

Michał Burdukiewicz 1 , Piotr Sobczyk 2 , Paweł Mackiewicz 1 and Stefan Rödiger 3

¹University of Wrocław, Department of Genomics, Poland

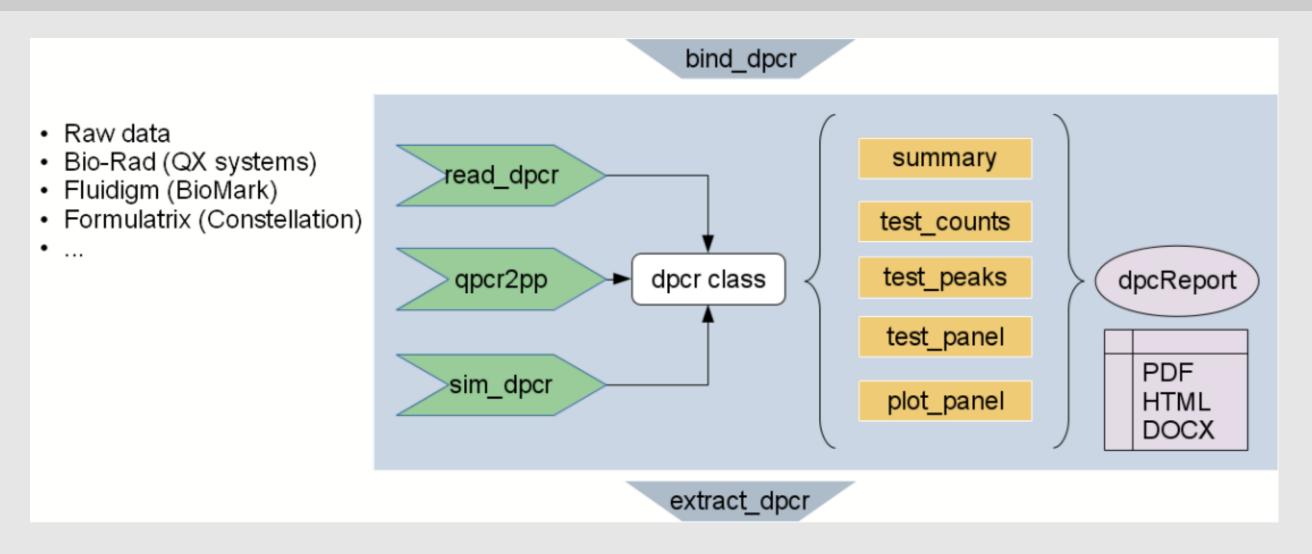
²Wrocław University of Technology, Department of Mathematics, Poland

³Faculty of Natural Sciences, Brandenburg University of Technology Cottbus–Senftenberg, Germany

Introduction

Point-of-Care tests brings the medical examination to the patient. The hardware necessary for such diagnosis should be completed with a suitably portable analysis framework. We introduce dpcReport, a versatile open source cross-platform software for analysis of digital PCR (dPCR) experiments. It might be accessed as a web server from any modern personal computer or even a smartphone, greatly enhancing its portability.

Workflow



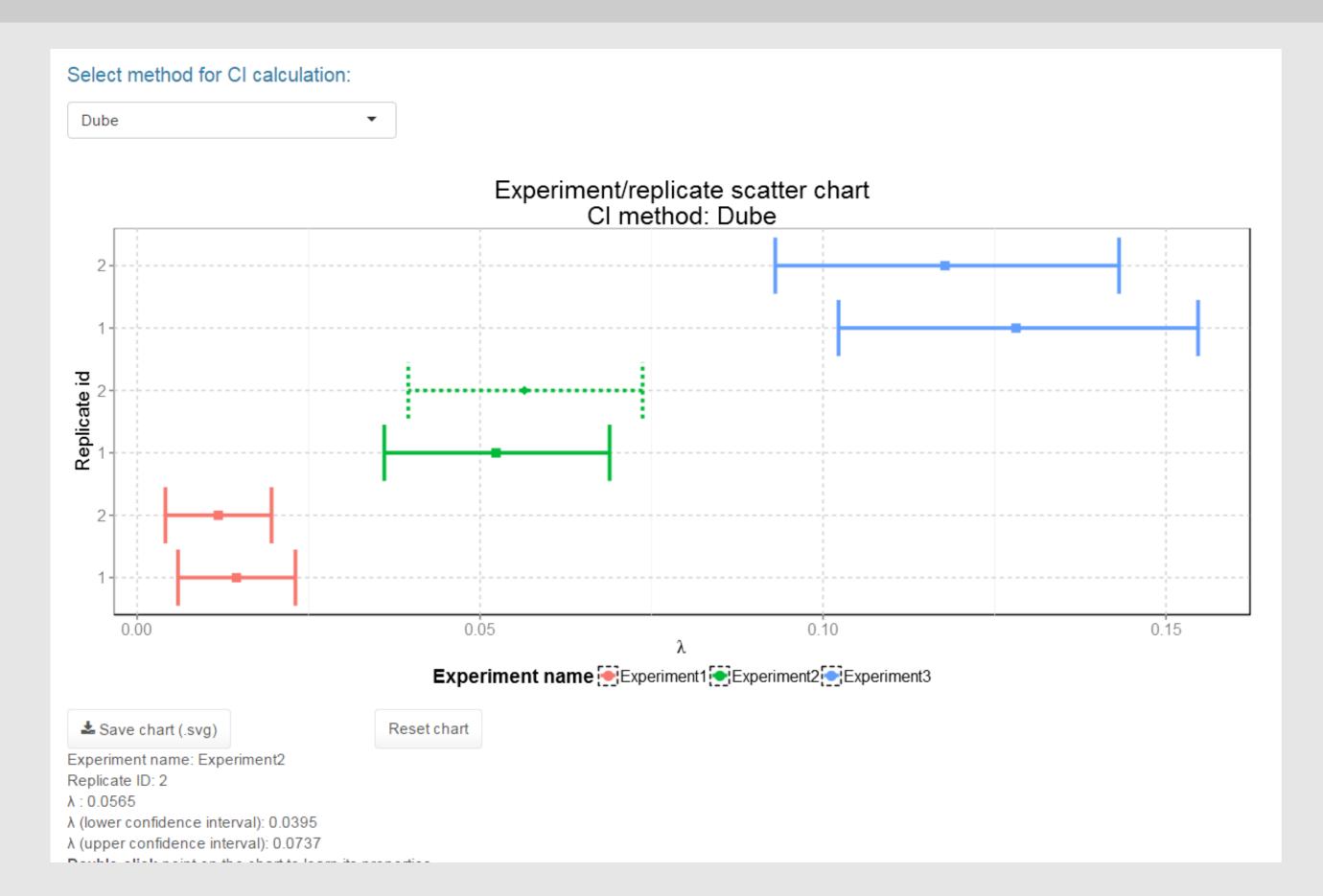
The aim of dpcReport is an exhaustive data analysis integrated into a single framework. Even the most basic data pre-processing (as renaming runs) does not require additional software, because it is already covered by dpcReport. Such approach is particulary suitable for POC framework, where access to the external software might be limited.

Input data

Vendor	System
Bio-Rad	QX100 & QX200
Fluidigm	BioMark
Formulatrix Constellation Digital PCR	

Results from systems described in the table above may be directly used as the input in the dpcReport worflow. Moreover, additional data formats can be processed with the functionality provided by the R environment Rödiger et al. (2015). Broad importing functionalities allow an integration of information from many dPCR systems.

Data summary



The summaries of input data are available as interactive tables (allowing filtering and selecting specified runs) and interactive charts (as presented on the figure above, where replicate 2 of experiment 2 is selected).

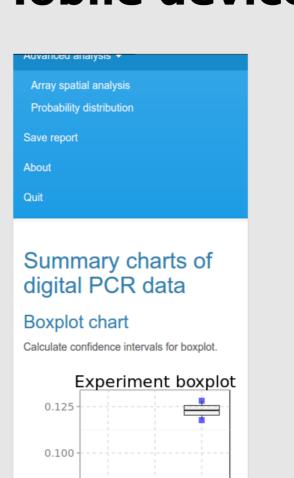
Statistical analysis

Array based dPCR experiments provide information about spatial distribution of partitions. Procedures belonging to spatial statistics verify if the status (positive, negative) of partition depends on its location. To address such questions, we implemented a Complete Spatial Randomness test (Baddeley and Turner, 2005) for dPCR arrays.

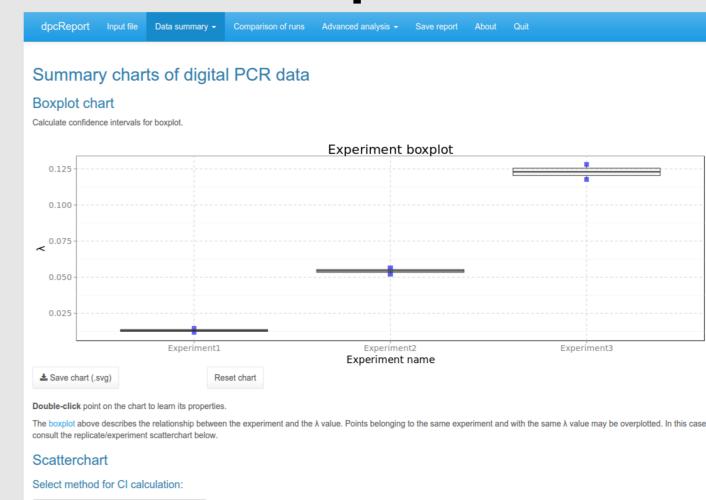
dpcReport uses two methods of comparing dPCR experiments, able to simultaneously compare λ values (mean number of template molecules per partition) of multiple runs. They are based respectively on Generalized Linear Models (Bretz et al., 2010) and on the uniformly most powerful (UMP) ratio test (Fay, 2010).

Graphical User Interface

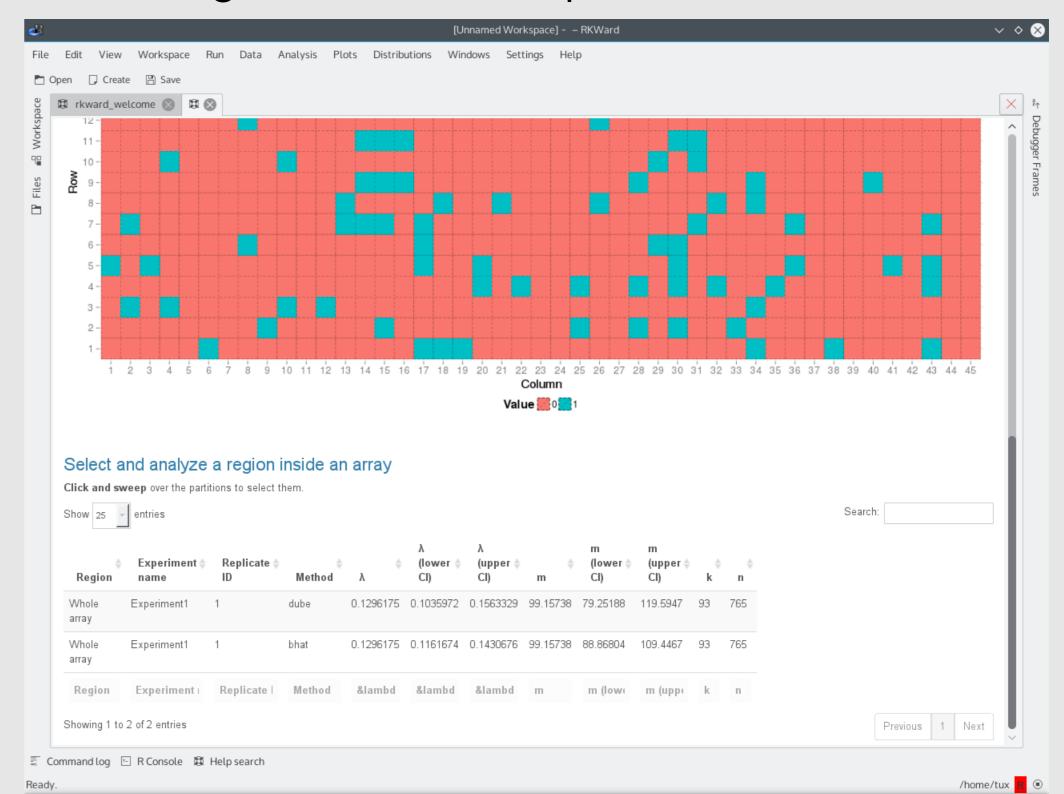
Mobile device



Desktop device



The dpcReport may be used as a web application accessible from every modern web browser, including web browsers for portable devices.



Additionally, dpcReport is accessible through R IDE/GUI such as RKWard Rödiger et al. (2012) or **RStudio**. Moreover, it may be downloaded and installed as a standalone application.

Analysis output

The structure of the output is fully modular and will contain only areas of the analysis chosen by the investigator. It is also possible to save only selected charts or tables.

The framework is based on the sophisticated statistical computing environment $\bf R$. An important option of dpcReport is an export of the $\bf R$ source code used for the report generation is provided to the user. This code can be used for recreating the analysis in the R environment or prototyping more complicated workflows.

Summary

The phylosophy of point-of-care testing require frameworks that can be used directly at the site of patient care. The portable and self-contained dpcReport fulfills such requirements and might be easily integrate into POCT routines involving dPCR assays.

Availability

dpcReport (standalone version for Microsoft Windows $^{(R)}$):

http://sourceforge.net/projects/dpcreport/



dpcReport (web server):

http://www.smorfland.uni.wroc.pl/dpcReport



dpcR package: https://github.com/michbur/dpcR



Bibliography

Baddeley, A. and Turner, R. (2005). spatstat: An R package for analyzing spatial point patterns. Journal of Statistical Software, 12(6):1-42. Bretz, F., Hothorn, T., and Westfall, P. (2010). Multiple Comparisons Using R. Chapman & Hall/CRC Press, Boca Raton, Florida, USA.

Fay, M. (2010). Two-sided exact tests and matching confidence intervals for discrete data. Proceedings of the National Academy of Sciences of the United States of America, 2(1):53–58.

Rödiger, S., Burdukiewicz, M., Blagodatskikh, K. A., and Schierack, P. (2015). R as an Environment for the Reproducible Analysis of DNA Amplification Experiments. The R Journal, 7(2):127–150.

Rödiger, S., Friedrichsmeier, T., Kapat, P., and Michalke, M. (2012). RKWard: A Comprehensive Graphical User Interface and Integrated Development Environment for Statistical Analysis with R. Journal of Statistical Software, 49(9):1-34.