BioCPR - A tool for correlation plots

BioCPR is an open-source application written in the R programming language. To run BioCPR in a personal system the primary requirment is the latest version of R and R-studio. This supplementary file provides information on how install R, R-studio and run BioCPR.

Step 1: Installing R

Download and install R by visiting https://cran.r-project.org/ and select an installer for your operating system.



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load and Install R

- Precompiled binary distributions of the base system and contributed packages, Windows and Mac users most likely want one of these versions of R:
 - Download R for Linux
 - Download R for (Mac) US 2
 Download R for Windows

R is part of many Linux distributions, you should check with your Linux package management system in addition to the link above.

Source Code for all Platforms

Windows and Mac users most likely want to download the precompiled binaries listed in the upper box, not the source code. The sources have to be compiled before you can use them. If you do not know what this means, you probably do not want to do it!

- The latest release (2021-05-18, Camp Pontanezen) R-4.1.0.tar.gz, read what's new in the latest version
- Sources of R alpha and beta releases (daily snapshots, created only in time periods before a planned release).
- Daily snapshots of current patched and development versions are <u>available here</u>. Please read about <u>new features and bug fixes</u> before filing corresponding feature requests or bug reports.
- Source code of older versions of R is available here.
- Contributed extension <u>packages</u>

Ouestions About R

 If you have questions about R like how to download and install the software, or what the license terms are, please read our answers to frequently asked questions before you send an email.

What are R and CRAN?

R is 'GNU S', a freely available language and environment for statistical computing and graphics which provides a wide variety of statistical and graphical techniques: linear and nonlinear modelling, statistical tests, time series analysis, classification, clustering, etc. Please consult the <u>R project homepage</u> for further information.

CRAN is a network of ftp and web servers around the world that store identical, up-to-date, versions of code and documentation for R. Please use the CRAN mirror nearest to you to

Supplementary Figure A1: The download page on the official R website (https://cran.r-project.org/).

Step 2: Installing Rstudio

Download RStudio from https://www.rstudio.com/products/rstudio/download/ for your operating system. RStudio Desktop is sufficient for personal users.

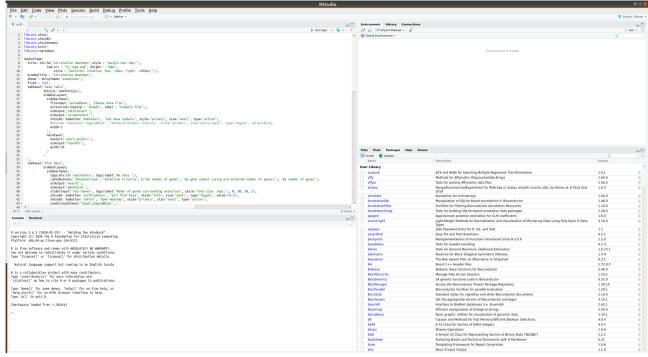
| | RStudio Desktop Open Source License Free | RStudio Desktop Pro Commercial License \$995 /year | RStudio Server Open Source License Free | RStudio Server Pro Commercial License \$4,975 /year (5 Named Users) |
|---|--|--|---|---|
| | Learn more | Learn more | Learn more | Evaluation Learn more |
| Integrated Tools for R | ~ | ~ | ~ | ~ |
| Priority Support | | ~ | | ~ |
| Access via Web Browser | | | ~ | ~ |
| RStudio Professional Drivers | | ~ | | ~ |
| Connect to RStudio Server Pro remotely | | ~ | | |
| Enterprise Security | | | | ~ |
| Project Sharing | | | | ~ |
| Manage Multiple R Sessions & Versions | | | | ~ |
| Admin Dashboard | | | | ~ |
| Load Balancing | | | | ~ |
| Auditing and Monitoring | | | | ~ |
| Data Connectivity | | | | ~ |
| Launcher | | | | ~ |
| Tutorial API | | | | ~ |
| License | AGPL | Commercial | AGPL | Commercial |

Supplementary Figure A2: The download page on the official RStudio website (https://rstudio.com/products/rstudio/download/)

Note: A detailed instruction on installing R and Rstudio on various operating system can be found in the following tutorial; https://www.datacamp.com/community/tutorials/installing-R-windows-mac-ubuntu

Step 3: Checking and setting working directory

Check if R and Rstudio are properly working and set the working directory to the folder all the BioCPR files are located (if the git repository is already downloaded).



Supplementary Figure A3: RStudio startup page

Step 4: Installing required libraries

Install the required libraries for running BioCPR.

You can install the stable version from CRAN and it will install all the necessary depencies for running the tool. Please find the commands below;

```
install.packages("heatmapFlex")
install.packages("convertid")
install.packages("readmoRe")
install.packages("coreheat")
```

If you prefer to install the latest development version, there are 4 custom in-built libraries that needs to be installed in addition to libraries from CRAN and BioConductor. Please find the commands below;

```
.libPaths()
libLocation <- .libPaths()[1]

## CRAN packages ##

cranPackages <- c("shinyBS", "shinythemes", "knitr", "rmarkdown", "shinyjs", "plyr", "RColorBrewer",
"R.utils", "gdata", "data.table", "foreach", "ggplot2", "scales", "curl", "openssl", "httr", "Rcurl", "XML",
"WGCNA", "DT", "devtools")

sapply(cranPackages, install.packages, lib = libLocation)

## Bioconductor packages ##
```

bioConPackages <- c("AnnotationDbi", "biomaRt", "org.Hs.eg.db", "org.Mm.eg.db", "Heatplus", "genefilter", "impute", "preprocessCore", "GO.db")

```
if (!requireNamespace("BiocManager", quietly = TRUE))
install.packages("BiocManager")
BiocManager::install(bioConPackages)
```

```
install_github("vfey/heatmapFlex")
install_github("vfey/convertid")
install_github("vfey/readmoRe")
install_github("vfey/coreheat")
```

Step 5: Run BioCPR

- Download the <u>git repository</u> and extract the files from the zip file.
- Set the working directory in Rstudio to the BioCPR folder downloaded from github and open the ui.R file in Rstudio.
- After opening the file in Rstudio, A "run App" button should appear and clicking the button will open the BioCPR app in a new screen.

```
| Sun App | Sun
```

Supplementary Figure A4: ui.R file in the RStudio interface with the "Run App" button highlighted.

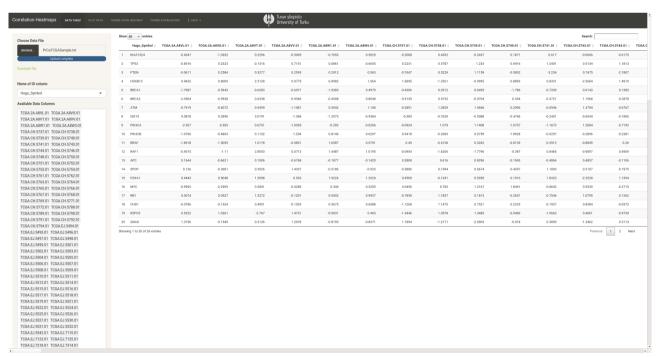
Step 6: Using BioCPR

1. Upon running, the BioCPR interface is initiated. The BioCPR interface is depicted in supplementary figure A5. You can choose the browse button in the app to upload the sample dataset provided in the git bundle.



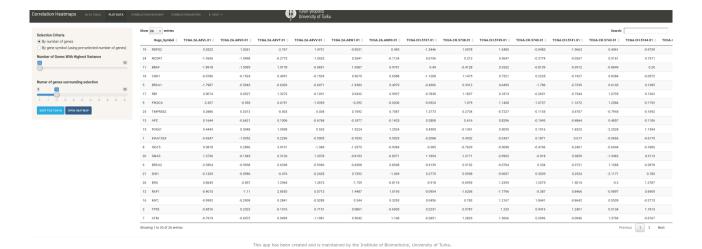
Supplementary Figure A5: Screenshot of startup page of BioCPR tool.

2. Upon uploading the file, the app shows the contents of the file on the right side of the window and the option to select the ID column on the left along with the sample names in the file.

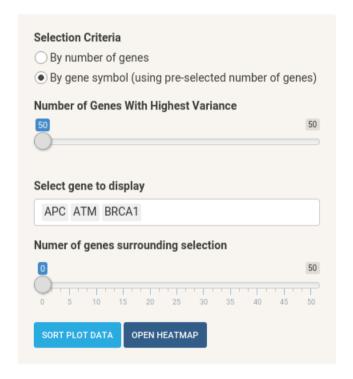


Supplementary Figure A6: Screenshot of startup page of BioCPR tool after loading the dataset.

3. After reviewing the dataset, we move on to the "PLOT DATA" tab where we select the data to the plotted in the heatmap (Figure A7). We can select either by limiting the number of genes with highest variance between them or by using gene symbol to select required genes while also including genes surrounding them using the "Number of genes surrounding selection" option (Figure A8).

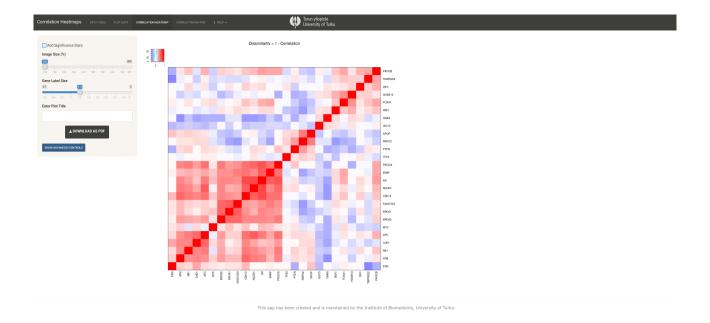


Supplementary Figure A7: Screenshot of "PLOT DATA" tab where we select the genes for plotting.

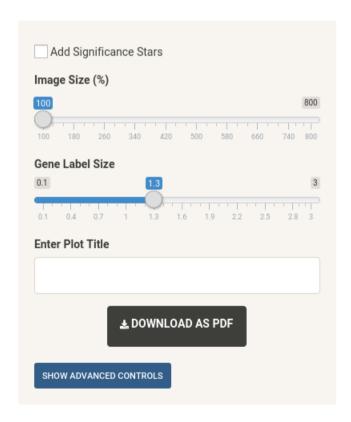


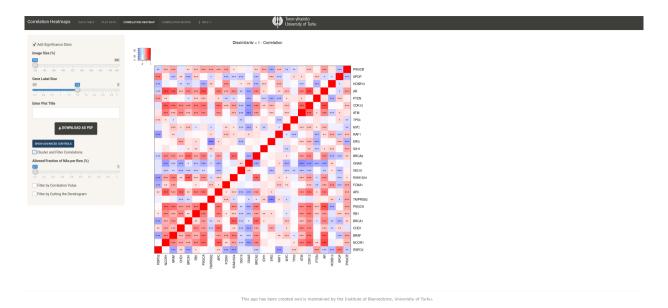
Supplementary Figure A8: Screenshot of selection criteria under the plot data tab.

4. After selecting the required genes you want to plot, you then move on to the "CORRELATION HEATMAP" tab. Here you can see a visualization of the heatmap of the data selected in the "PLOT DATA" tab (Figure A9). Here you can notice heatmap on the right and the options to edit the heatmap on the left (figure A10).



Supplementary Figure A9: Screenshot of correlation heatmaps generated from data selected in "PLOT DATA" tab.

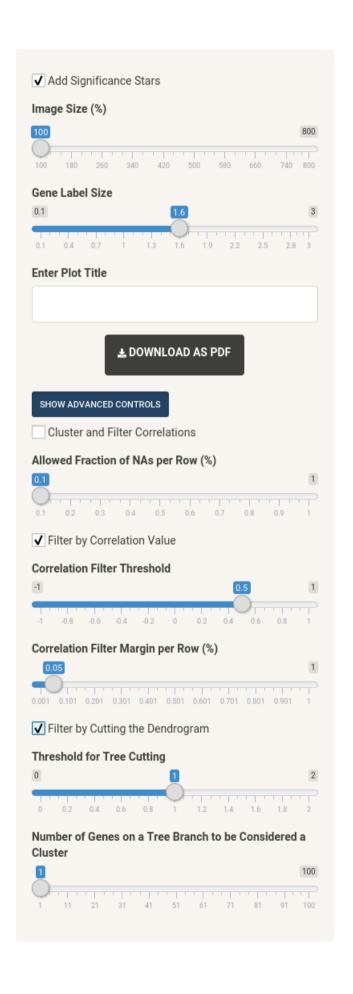




Supplementary Figure A11: Screenshot of the heatmap with significance stars added.

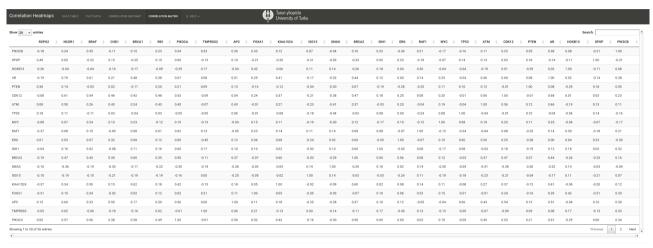
Using options tab, various parameters of the heatmap can be altered including:

- 1. Adding significance stars to the heatmap (Figure A11).
- 2. Adjusting the image size
- 3. Adjusting the gene label size
- 4. Advanced options such as removing clustering, filtering by correlation value, cutting the heatmap into dendrogram trees (Figure A12)



Supplementary Figure A12: Screenshot of the advanced options for editing heatmap.

- 5. After editing the heatmap as per our requirements, we can enter a title for our plot and download the edited heatmap using the download button.
- 6. We can also view the correlation matrix that was used to create the heatmap in the "CORRELATION MATRIX" tab (Figure A13).



This app has been created and is maintained by the Institute of Biomedicine, University of Turku.

Supplementary Figure A13: Screenshot of the "CORRELATION MATRIX" tab.

7. Finally the help tab, provides a information on frequently asked questions and the format of the data required.

R session information:

| — Session info | | |
|--------------------------------------|--|--|
| | | |
| | | |
| setting value | | |
| version R version 3.5.2 (2018-12-20) | | |
| os Ubuntu 18.04.1 LTS | | |
| system x86_64, linux-gnu | | |
| ui RStudio | | |
| language (EN) | | |
| collate en_US.UTF-8 | | |

```
ctype en_US.UTF-8
     Europe/Helsinki
date 2019-01-08
- Packages
           * version date
                            lib source
package
           1.4.1 2016-10-29 [1] CRAN (R 3.5.1)
acepack
annotate
          1.58.0 2018-11-14 [1] Bioconductor
AnnotationDbi * 1.42.1 2018-11-14 [1] Bioconductor
assertthat 0.2.0 2017-04-11 [1] CRAN (R 3.5.1)
backports 1.1.2 2017-12-13 [1] CRAN (R 3.5.1)
base64enc 0.1-4 2018-11-14 [1] local
bindr
          0.1.1 2018-03-13 [1] CRAN (R 3.5.1)
bindrcpp
          0.2.2 2018-03-29 [1] CRAN (R 3.5.1)
Biobase
           * 2.40.0 2018-11-14 [1] Bioconductor
BiocGenerics * 0.26.0 2018-11-14 [1] Bioconductor
biomaRt * 2.36.1 2018-11-14 [1] Bioconductor
bit
         1.1-14 2018-05-29 [1] CRAN (R 3.5.1)
bit64
         0.9-7 2017-05-08 [1] CRAN (R 3.5.1)
          1.0-6 2013-08-17 [1] CRAN (R 3.5.1)
bitops
blob
          1.1.1 2018-03-25 [1] CRAN (R 3.5.1)
checkmate 1.8.5 2017-10-24 [1] CRAN (R 3.5.1)
cli
         1.0.1 2018-09-25 [1] CRAN (R 3.5.1)
         * 0.4.1 2018-06-23 [1] CRAN (R 3.5.2)
clipr
         * 2.0.7-1 2018-04-09 [1] CRAN (R 3.5.1)
cluster
codetools 0.2-16 2018-12-24 [4] CRAN (R 3.5.2)
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```
colorspace 1.4-0 2018-11-05 [1] R-Forge (R 3.5.1)
           * 0.1-5 2018-11-14 [2] local
coreheat
           1.3.4 2017-09-16 [1] CRAN (R 3.5.1)
crayon
data.table * 1.11.8 2018-09-30 [1] CRAN (R 3.5.1)
          1.0.0 2018-05-02 [1] CRAN (R 3.5.1)
              1.0-8 2016-11-19 [1] CRAN (R 3.5.1)
          * 0.6.18 2018-10-10 [1] CRAN (R 3.5.1)
           1.0.14 2018-09-21 [2] R-Forge (R 3.5.1)
doParallel
           0.7.8 2018-11-10 [1] CRAN (R 3.5.1)
dplyr
                 2018-11-05 [2] CRAN (R 3.5.1)
DT
         * 0.5
dynamicTreeCut 1.63-1 2016-03-11 [2] CRAN (R 3.5.1)
evaluate
            0.12 2018-10-09 [1] CRAN (R 3.5.1)
fastcluster 1.1.25 2018-06-07 [1] CRAN (R 3.5.1)
fit.models 0.5-14 2017-04-06 [2] CRAN (R 3.5.1)
foreach
           * 1.5.1 2018-08-27 [1] R-Forge (R 3.5.1)
           0.8-71 2018-07-20 [4] CRAN (R 3.5.1)
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genefilter
          * 1.62.0 2018-11-14 [1] Bioconductor
ggplot2
           * 3.1.0 2018-10-25 [1] CRAN (R 3.5.1)
glue
          1.3.0 2018-07-17 [1] CRAN (R 3.5.1)
GO.db
            3.6.0 2018-11-14 [1] Bioconductor
            2.3
                  2017-09-09 [1] CRAN (R 3.5.1)
gridExtra
           0.2.0 2016-02-26 [1] CRAN (R 3.5.1)
gtable
           3.8.1 2018-06-26 [1] CRAN (R 3.5.1)
gtools
heatmapGen * 0.1-0 2018-11-14 [2] local
heatmapGen2 * 0.1-3 2018-11-14 [2] local
Heatplus * 2.26.0 2018-11-14 [1] Bioconductor
```

```
Hmisc
            4.1-1 2018-01-03 [1] CRAN (R 3.5.1)
           0.4.2 2018-03-10 [1] CRAN (R 3.5.1)
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htmlTable
            1.12 2018-05-26 [1] CRAN (R 3.5.1)
htmltools
            0.3.6 2017-04-28 [1] CRAN (R 3.5.1)
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                    2018-09-30 [1] CRAN (R 3.5.1)
           1.4.5 2018-07-19 [1] CRAN (R 3.5.1)
          1.3.1 2017-08-20 [1] CRAN (R 3.5.1)
httr
           1.54.0 2018-11-14 [2] Bioconductor
impute
           * 2.14.12 2018-09-20 [1] Bioconductor
IRanges
           1.0.11 2018-07-05 [1] R-Forge (R 3.5.1)
iterators
jsonlite
           1.5
                 2017-06-01 [1] CRAN (R 3.5.1)
         * 1.20 2018-02-20 [1] CRAN (R 3.5.1)
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          0.7.5 2018-09-18 [1] CRAN (R 3.5.1)
later
lattice
          0.20-38 2018-11-04 [4] CRAN (R 3.5.1)
latticeExtra 0.6-28 2016-02-09 [1] CRAN (R 3.5.1)
           0.2.1 2017-10-29 [1] CRAN (R 3.5.1)
lazyeval
                  2014-11-22 [1] CRAN (R 3.5.1)
magrittr
            1.5
markdown
            0.8.1 2018-11-14 [1] local
MASS
            7.3-51.1 2018-11-01 [4] CRAN (R 3.5.1)
Matrix
            1.2-15 2018-11-01 [4] CRAN (R 3.5.1)
matrixStats 0.54.0 2018-07-23 [1] CRAN (R 3.5.1)
            * 0.4-10 2018-11-14 [2] local
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memoise
            1.1.0 2017-04-21 [1] CRAN (R 3.5.1)
            0.6.1 2018-11-14 [1] local
mime
            0.5.0 2018-06-12 [1] CRAN (R 3.5.1)
munsell
             1.0-8 2018-05-31 [1] CRAN (R 3.5.1)
mvtnorm
nnet
           7.3-12 2016-02-02 [4] CRAN (R 3.5.0)
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      1.3.0 2018-07-14 [1] CRAN (R 3.5.1)
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       * 1.8.4 2016-06-08 [1] CRAN (R 3.5.1)
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progress 1.2.0 2018-06-14 [1] CRAN (R 3.5.1)
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purrr 0.2.5 2018-05-29 [1] CRAN (R 3.5.1)
R.methodsS3 * 1.7.1 2016-02-16 [1] CRAN (R 3.5.1)
R.oo
         * 1.22.0 2018-04-22 [1] CRAN (R 3.5.1)
R.utils
         * 2.7.0 2018-08-27 [1] CRAN (R 3.5.1)
          2.3.0 2018-10-04 [1] CRAN (R 3.5.1)
RColorBrewer * 1.1-2 2014-12-07 [1] CRAN (R 3.5.1)
           1.0.0 2018-11-07 [1] CRAN (R 3.5.1)
Rcpp
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          0.3.0.1 2018-10-25 [1] CRAN (R 3.5.1)
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robustbase 0.93-3 2018-09-21 [1] CRAN (R 3.5.1)
          4.1-13 2018-02-23 [4] CRAN (R 3.5.0)
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          1.4-7 2018-11-15 [1] CRAN (R 3.5.1)
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RSQLite 2.1.1 2018-05-06 [1] CRAN (R 3.5.1)
```

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                 2018-10-02 [1] CRAN (R 3.5.1)
$4Vectors * 0.18.3 2018-11-14 [1] Bioconductor
       * 1.0.0 2018-08-09 [1] CRAN (R 3.5.1)
scales
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shiny * 1.2.0 2018-11-02 [1] CRAN (R 3.5.1)
           * 0.61 2015-03-31 [1] CRAN (R 3.5.1)
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           2.2.0 2018-07-25 [1] CRAN (R 3.5.1)
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