teaching_material_book

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r paste(Sys.time())

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Preface

Who is this for

This tutorial is aimed at data scientists / analysts who want to write their first teal application. We expect you to already have some R experience and that you know what you want to achieve with the application. You do not need to know how to create a shiny application. The teal package aims to make the creation of (teal-specific) shiny applications relatively intuitive without the need to understand how shiny works exactly. Shiny knowledge is useful once you want to either modify an application, or write your own teal module from scratch.

What will you learn

How is it structured

All chapters in this book follow a common structure: First, the learning goals of a chapter is defined. We then go through the step-by-step process of using specific teal modules. In each chapter we combine what we learned into one code chunk with which we can create a functional teal application. As every teal app requires data, the app created in the introduction will be the fundamental building block of the apps in the subsequent chapters. In the toy apps created in each chapter you should be able to understand every line of code. The code for a more developed app (but built for the same purpose) and a deployed instance of that app is linked at the end of each chapter. What you learned in this chapter should give you a good understanding of how the more developed application works.

Where can you find more

1 Introduction

At the end of this chapter you should be able to

- understand the structure and utility of teal apps
- understand the utility of teal apps
- determine whether teal apps are useful for you

1.1 Structure of teal apps

teal is a shiny-based modular framework. It provides an interactive user interface allowing users to customize tables and visualizations facilitating the exploration and analysis of clinical trial data. Dynamic analysis and display control, data filtering and code reproducibility are included in each module.

teal modules are laid out in three panels.

- The Left Panel, referred to as the encoding panel, includes elements used to adjust the output interactively.
- The Center Panel, referred to as the output panel, where the tables and visualizations are displayed.
- The Right Panel, referred to as the data filter panel, includes elements used to subset the data interactively.

The layout is pre-set within each module. However, user can decide overall tab order and nesting. See **?@fig-layout** for the typical teal app structure.

1.1.1 Modules

teal modules are modular building blocks for your teal app. They implement specific app functionalities, such as outlier exploration, data visualizations, and survival analysis. Although modules can be created from scratch, many teal modules have already been released and can be found in the following packages:

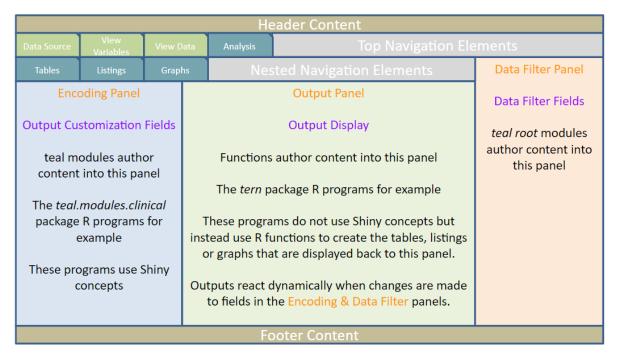


Figure 1.1: Typical teal app layout

- teal.modules.general: general modules for exploring relational/independent/CDISC data
- teal.modules.clinical: modules specific to CDISC data and clinical trial reporting
- $\bullet \ \ teal.modules.hermes: \ modules \ for \ analyzing \ \underline{MultiAssayExperiment \ objects}$

are shiny modules built within the teal framework that specify analysis to be performed.

2 Data app

The App created at the end of this chapter will be the basic building block of all subsequent chapters in this book. The structure of the app code will remain consistent throughout the book — make sure you understand each line of code of the *toy* application at the end.

At the end of this chapter you should be able to

- understand the code structure of teal apps
- understand the workflow to create a teal app
- install and load the relevant packages
- write your own app that displays data

2.1 Setup

In this tutorial we will be working with realistic looking but simulated data.

```
if (!require("remotes")) install.packages("remotes")

# install teal
remotes::install_github("insightsengineering/teal@*release")

# install teal.modules.general
remotes::install_github("insightsengineering/teal.modules.general@*release")

# install tern mmrm (required for teal.modules.clinical)
remotes::install_github("insightsengineering/tern.mmrm@*release")

# install teal.modules.clinical
remotes::install_github("insightsengineering/teal.modules.clinical@*release")

# install teal.modules.hermes
remotes::install_github("insightsengineering/teal.modules.hermes@*release")
```

```
library(random.cdisc.data) # this package provides the data used
  library(teal.data)
Loading required package: shiny
  library(scda)
There are no scda.XXXX libraries installed, like scda.2021.
Please install an scda database to take full advantage of the scda package.
  library(teal.modules.general)
Loading required package: ggmosaic
Loading required package: ggplot2
Loading required package: shinyTree
Loading required package: teal
Loading required package: teal.transform
Loading required package: magrittr
You are using teal version 0.12.0
  library(sparkline)
  library(teal)
  library(tidyverse)
```

-- Attaching packages ----- tidyverse 1.3.2 --

```
v dplyr
v tibble 3.1.8
                            1.0.10
v tidyr
         1.2.1
                   v stringr 1.4.1
         2.1.3
                   v forcats 0.5.2
v readr
         0.3.5
v purrr
-- Conflicts ----- tidyverse conflicts() --
x tidyr::extract()
                    masks magrittr::extract()
x dplyr::filter()
                    masks stats::filter()
x dplyr::lag()
                    masks stats::lag()
x tidyr::replace_na() masks random.cdisc.data::replace_na()
x purrr::set_names() masks magrittr::set_names()
```

We first load two synthetic data sets. random.cdisc.data::cadsl is a Subject Level Analysis Dataset (ADSL) with one record (row) per subject. The unique identifier per subject is stored in the variable USUBJID. According to CDISC the main purpose of ADSL is to provide a "(...) source for denominators for populations of interest, stratification variables, and other important subject subgroups".

We can now load the three data-sets:

Data Structure for Adverse Event Analysis

Data-set name	Explanation	Purpose
ADSL ADAE ADTTE	Analysis data-set subject level Analysis data-set for adverse events Analysis data-set for time to event	Filtering Adverse events Efficacy

```
ADSL <- random.cdisc.data::cadsl
ADAE <- random.cdisc.data::cadae
ADTTE <- random.cdisc.data::cadtte
```

Let's look at the first three rows of the data-frames.

```
map(list(ADSL, ADAE, ADTTE), head, 3)
```

[[1]]

```
# A tibble: 3 x 56
            STUDYID USUBJID
                                                                                                                                                  SUBJID SITEID
                                                                                                                                                                                                                                                               AGE AGEU SEX
                                                                                                                                                                                                                                                                                                                                                                           RACE ETHNIC COUNTRY DTHFL
             <chr>
                                                                 <chr>
                                                                                                                                                   <chr> <chr> <int> <fct> <fct > <f
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         <fct>
1 AB12345 AB12345-CH~ id-128 CHN-3
                                                                                                                                                                                                                                                                     32 YEARS M
                                                                                                                                                                                                                                                                                                                                                                          ASIAN HISPA~ CHN
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Y
2 AB12345 AB12345-CH~ id-262 CHN-15
                                                                                                                                                                                                                                                                     35 YEARS M
                                                                                                                                                                                                                                                                                                                                                                          BLAC~ NOT H~ CHN
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        N
3 AB12345 AB12345-RU~ id-378 RUS-3
                                                                                                                                                                                                                                                                     30 YEARS F
                                                                                                                                                                                                                                                                                                                                                                          ASIAN NOT H~ RUS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        N
```

- # ... with 45 more variables: INVID <chr>, INVNAM <chr>, ARM <fct>,
- # ARMCD <fct>, ACTARM <fct>, ACTARMCD <fct>, TRT01P <fct>, TRT01A <fct>,
- # TRTO2P <fct>, TRTO2A <fct>, REGION1 <fct>, STRATA1 <fct>, STRATA2 <fct>,
- # BMRKR1 <dbl>, BMRKR2 <fct>, ITTFL <fct>, SAFFL <fct>, BMEASIFL <fct>,
- # BEP01FL <fct>, AEWITHFL <fct>, RANDDT <date>, TRTSDTM <dttm>,
- # TRTEDTM <dttm>, TRT01SDTM <dttm>, TRT01EDTM <dttm>, TRT02SDTM <dttm>,
- # TRTO2EDTM <dttm>, APO1SDTM <dttm>, APO1EDTM <dttm>, APO2SDTM <dttm>, ...

[[2]]

- # A tibble: 3 x 92
 - STUDYID USUBJID SUBJID SITEID AGE AGEU SEX RACE ETHNIC COUNTRY DTHFL chr> chr> <a href="ht

- # ... with 81 more variables: INVID <chr>, INVNAM <chr>, ARM <fct>,
- # ARMCD <fct>, ACTARM <fct>, ACTARMCD <fct>, TRT01P <fct>, TRT01A <fct>,
- # TRTO2P <fct>, TRTO2A <fct>, REGION1 <fct>, STRATA1 <fct>, STRATA2 <fct>,
- # BMRKR1 <dbl>, BMRKR2 <fct>, ITTFL <fct>, SAFFL <fct>, BMEASIFL <fct>,
- # BEP01FL <fct>, AEWITHFL <fct>, RANDDT <date>, TRTSDTM <dttm>,
- # TRTEDTM <dttm>, TRT01SDTM <dttm>, TRT01EDTM <dttm>, TRT02SDTM <dttm>,
- # TRTO2EDTM <dttm>, AP01SDTM <dttm>, AP01EDTM <dttm>, AP02SDTM <dttm>, ...

[[3]]

- # A tibble: 3 x 68
- STUDYID USUBJID SUBJID SITEID AGE AGEU SEX RACE ETHNIC COUNTRY DTHFL ><a href="https://doi.org/

- # ... with 57 more variables: INVID <chr>, INVNAM <chr>, ARM <fct>,
- # ARMCD <fct>, ACTARM <fct>, ACTARMCD <fct>, TRT01P <fct>, TRT01A <fct>,
- # TRTO2P <fct>, TRTO2A <fct>, REGION1 <fct>, STRATA1 <fct>, STRATA2 <fct>,
- # BMRKR1 <dbl>, BMRKR2 <fct>, ITTFL <fct>, SAFFL <fct>, BMEASIFL <fct>,
- # BEP01FL <fct>, AEWITHFL <fct>, RANDDT <date>, TRTSDTM <dttm>,
- # TRTEDTM <dttm>, TRT01SDTM <dttm>, TRT01EDTM <dttm>, TRT02SDTM <dttm>,
- # TRTO2EDTM <dttm>, AP01SDTM <dttm>, AP01EDTM <dttm>, AP02SDTM <dttm>, ...

2.2 Your first app

As is generally the case when you start a project, you should have an approximate idea of what you would like to achieve. In this case, the aim is to create an app that visualizes variables of three connected study data-sets.

2.3 Code structure

First, you want to define the project metadata. This can include the author name, the study name, the molecule name, etc. then, we have a section where we can define parameters for the app. This can be simple parameter values but they can also be short code snippets that would otherwise often be repeated throughout the application, e.g. pre-selected variables of interest.

After that, we define the app using the init() function. This function takes data, modules, and html input and returns a list (usually named app) containing the user interface (the ui object) and back end (the server object) of the application. In the final step we instantiate the app by calling shinyApp(app\$ui, app\$server).

2.4 init()

The most crucial function of the teal package(s) is the teal::init() function, which is structured as follows:

```
init(
  data,
  modules,
  title = NULL,
  filter = list(),
  header = tags$p("Add Title Here"),
  footer = tags$p("Add Footer Here"),
  id = character(0)
)
```

The init() function sets up a shiny app that consists of teal modules. Let's go through the arguments:

data

In the data argument we define one or more data-frames for the application. If more than one data-frames are specified, they should be combined as a list, e.g. data = list(ADSL,

ADTR). For ADaM data-sets, the package teal.data provides helper functions with which the data argument of init() can be specified, — the cdisc_data() and the cdisc_dataset() functions. They allow the teal app to know the merge key variables of the data-sets. The cdisc_datafunction returns an S6 object.

keys are pre-specified variables in ADaM data to merge data-sets. Each type of data-set requires specific keys to be specified. The package teal.data automatically chooses the correct key variables based on the dataname for the following data-sets: ADSL, ADAE, ADEG, ADTTE, ADAETTE, ADCM, ADEX, ADLB, ADMH, ADQS, ADRS, ADSAFTTE, ADVS, ADDV, ADSUB, ADHY.

If you want to work with other ADaM data-sets you must specify the keys manually. ADTR is a Tumor Results Analysis data-set where there is one record (row) per subject. As of now, this is not part of the default cdisc keys in the teal.data package, so we would have to specify them manually:

```
ADTR <- random.cdisc.data::cadtr

cdisc_data(
  cdisc_dataset("ADSL", ADSL),
  cdisc_dataset("ADTR", ADTR, keys = c("STUDYID", "USUBJID", "PARAMCD", "AVISIT"))
)</pre>
```

Note that all keys have to be specified.

pre-processing

Generally, it is recommended to collect all data pre-processing steps in a separate R-script, which can then directly be applied to the 'cdisc_data()' output. One such pre-processing steps could be the creation of a new variable in the ADSL data-set which splits patients into two age groups where patients younger than 40 are AGEGR = 0 and patients older than 40 are AGEGR = 1.

```
# you would write this in a separate file directly, without the writeLines
writeLines(
  text = "
ADSL <- ADSL %>%
  dplyr::mutate(AGEGR = if_else(AGE < 40, 0, 1, NA_real_))",
  con = "ADSL_preproc.R"
)</pre>
```

The resulting R file looks something like this:

```
ADSL <- ADSL %>%
            dplyr::mutate(AGEGR = if_else(AGE < 40, 0, 1, NA_real_))</pre>
      ADSL_processed <- ADSL %>%
            cdisc_dataset("ADSL", .) %>%
            mutate_dataset(script = "ADSL_preproc.R")
       # check if the new variable exists:
      ADSL processed$get raw data()$AGEGR
       \begin{smallmatrix} [1] \end{smallmatrix} 0 \hspace{.1cm} 0 \hspace{.1cm} 0 \hspace{.1cm} 0 \hspace{.1cm} 0 \hspace{.1cm} 1 \hspace{.1cm} 1 \hspace{.1cm} 0 \hspace{.1cm} 0 \hspace{.1cm} 1 \hspace{.1cm} 1 \hspace{.1cm} 0 \hspace{.1cm} 0 \hspace{.1cm} 1 \hspace{.1cm} 0 \hspace{.
   [75] 0 0 1 0 0 0 0 0 0 0 0 0 0 0 0 1 0 0 0 0 1 0 0 0 0 1 0 0 0 0 0 1 0 0 1 0 1
[112] 0 0 0 1 0 1 0 0 1 0 1 0 0 0 0 0 0 0 1 0 1 0 0 0 0 1 1 0 1 0 0 0 0 1 1 0 0 0 0
[149] 0 1 1 0 1 0 0 1 0 0 0 1 0 1 0 0 1 1 0 0 0 1 0 0 0 0 0 0 0 0 0 0 0 0 0 0
In a real world case we would not necessarily want to separately save the pre-processed
ADSL in our environment. Instead, we would apply the pre-processing within the data
argument of the init() function, e.g., like this:
       app <- teal::init(</pre>
            data = cdisc data(
                  cdisc_dataset("ADSL", ADSL) %>%
                       mutate_dataset(script = "ADSL_preproc.R"),
                  cdisc_dataset("ADAE", ADAE),
                  cdisc_dataset("ADTTE", ADTTE)
                  # <<additional dataset code>>
            # , <<additional init arguments code>>
```

2.4.1 modules

header

footer

id

Combining everything above, our init() function looks like this:

```
app <- teal::init(
   data = cdisc_data(
      cdisc_dataset("ADSL", ADSL) %>%
        mutate_dataset(script = "ADSL_preproc.R"),
      cdisc_dataset("ADAE", ADAE),
      cdisc_dataset("ADTTE", ADTTE)
   ),
   header = "My first application",
   footer = "footer"
)
```

2.5 Toy app

We have now gone through all steps to create your first teal app.

```
# data app
## ---- load packages data app ----
library(random.cdisc.data) # this package provides the data used
library(teal.data)
library(scda)
library(teal.modules.general)
library(sparkline)
library(teal)
library(tidyverse)
## ---- parameter data app ----
author <- "Stefan Thoma"
molecule <- "hexadromedar"</pre>
## ---- load data data app ----
ADSL <- random.cdisc.data::cadsl
ADAE <- random.cdisc.data::cadae
ADTTE <- random.cdisc.data::cadtte
```

```
## ---- init data app ----
app <- teal::init(
   data = cdisc_data(
        cdisc_dataset("ADSL", ADSL) %>%
        mutate_dataset(script = "ADSL_preproc.R"),
        cdisc_dataset("ADAE", ADAE),
        cdisc_dataset("ADTTE", ADTTE)
   ),
   header = "My first application",
   footer = "footer"
)

## ---- run data app ----
shinyApp(app$ui, app$server)
?teal.modules.clinical::tm_g_forest_tte
```

The output should look something like this:

2.6 Production app

My first application

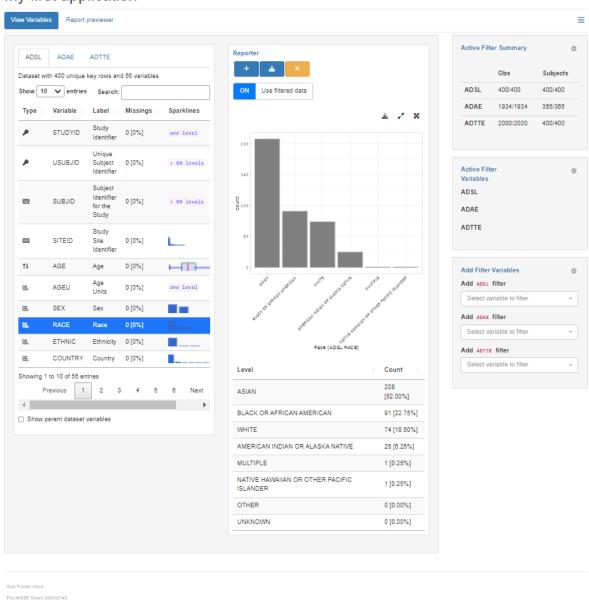


Figure 2.1: Data App preview

3 efficacy

Stefan Thoma

```
http://docs.roche.com/\#/agile-R/2022\_01\_28/teal/sample\_apps/sample-app-efficacy/2022\_01\_28/teal/sample-apps/sample-apps-efficacy/2022\_01\_28/teal/sample-apps/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/sample-apps-efficacy/2022\_01\_28/teal/
        sessionInfo()
R version 4.2.1 (2022-06-23)
Platform: x86_64-pc-linux-gnu (64-bit)
Running under: Ubuntu 20.04.5 LTS
Matrix products: default
                         /usr/lib/x86_64-linux-gnu/blas/libblas.so.3.9.0
LAPACK: /usr/lib/x86_64-linux-gnu/lapack/liblapack.so.3.9.0
locale:
    [1] LC_CTYPE=en_US.UTF-8
                                                                                                      LC_NUMERIC=C
    [3] LC_TIME=en_US.UTF-8
                                                                                                      LC_COLLATE=en_US.UTF-8
    [5] LC_MONETARY=en_US.UTF-8
                                                                                                      LC_MESSAGES=en_US.UTF-8
    [7] LC_PAPER=en_US.UTF-8
                                                                                                      LC_NAME=C
                                                                                                      LC_TELEPHONE=C
    [9] LC_ADDRESS=C
 [11] LC_MEASUREMENT=en_US.UTF-8 LC_IDENTIFICATION=C
attached base packages:
                                            graphics grDevices utils
[1] stats
                                                                                                                                            datasets methods
                                                                                                                                                                                                            base
loaded via a namespace (and not attached):
    [1] compiler_4.2.1 magrittr_2.0.3 fastmap_1.1.0
                                                                                                                                                                          cli_3.4.1
    [5] tools_4.2.1
                                                                  htmltools_0.5.3 rstudioapi_0.14 yaml_2.3.6
    [9] stringi_1.7.8
                                                                   rmarkdown_2.17 knitr_1.40
                                                                                                                                                                          stringr_1.4.1
 [13] xfun_0.34
                                                                   digest_0.6.30
                                                                                                                       jsonlite_1.8.2 rlang_1.0.6
 [17] evaluate_0.17
```

.libPaths()

- [1] "/home/thomas7/R/x86_64-pc-linux-gnu-library/4.2"
- [2] "/usr/local/lib/R/site-library"
- [3] "/usr/lib/R/site-library"
- [4] "/usr/lib/R/library"

4 Summary

In summary, this book has no content whatsoever.

1 + 1

[1] 2

References