Radiation Oncology (RadOnc) Tools

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$March\ 27,\ 2017$

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1 Introduction

The *RadOnc* package provides a number of tools for the import and analysis of dose-volume histogram (DVH) data used routinely in Radiation Oncology clinical practice and research. Supported formats for data import currently include:

- Varian's Aria/Eclipse platform (v.10 and v.11)
- CadPlan (for historical/archival data)
- TomoTherapy
- RayStation
- Elekta's Monaco platform
- DICOM-RT files

The functionality contained herein also enables visualization of dosimetric and volumetric data, and statistical comparison among multiple DVHs and three-dimensional structures. In order to use these tools, you must first load the RadOnc package:

> library(RadOnc)

It is assumed that the reader is already familiar with DVH analysis. If this is not the case, consult the relevant literature for a thorough treatment of the subject (1).

Throughout this vignette, we will be exploring actual data for 2 patients, each possessing a set of 10 structures (including organs at risk and treatment planning volumes). We will also demonstrate rudimentary three-dimensional structural processing and a number of other features.

For additional reference and details, please consult our published report on the RadOnc package(2).

2 Changes for RadOnc in current release

- Added ability to display gridlines when plotting DVH graphs.
- Added ability to extract median dose as a dosimetric parameter.
- Added ability to check for arbitrary dose constraints (e.g. V20Gy < 15(%)).
- New function subset() to extract a desired subset of a DVH.list object based upon patient identifiers, structure name, and/or dosimetric constraints.
- Extended read.DVH() function to import Elekta Monaco files.

Please refer to NEWS.Rd file for a more extensive revision history.

> news(package="RadOnc")

3 DVH Analysis

3.1 DVH file import

The read.DVH() function is designed to take an input text file and output a list of DVH data objects containing all relevant data. Supported file types currently include Varian's Aria/Eclipse platform (v.10 and v.11), CadPlan (for archival/historical data), TomoTherapy, RayStation, and Elekta's Monaco platform. Other treatment planning systems are not currently supported however will be included in future releases.

For Varian-specific file types, data must be exported directly from the treatment planning system and should include all DVH structures of interest. In Eclipse, this is accomplished via the "Export DVH in Tabular Format..." option, accessed by right-clicking over DVHs in Plan Evaluation mode. Exported files will adhere to the following form (an example file, 50 lines of which are shown here, is contained within this release of the *RadOnc* package):

Patient Name : Doe, Jane (111111111)

Patient ID : 1111111111

Comment : DVHs for one plan
Date : 05.24.2013 00:00:00

Type : Cumulative Dose Volume Histogram

Description : The cumulative DVH displays the percentage (relative)

or volume (absolute) of structures that receive a dose

equal to or greater than a given dose.

Plan: PLAN_NAME

Prescribed dose [cGy]: 5500.0

% for dose (%): 100.0

Structure: LIVER

Approval Status: Unapproved

Plan: PLAN_NAME
Course: COURSE_1
Volume [cc]: 1635.9
Dose Cover.[%]: 100.0
Sampling Cover.[%]: 100.0
Min Dose [cGy]: 42.7
Max Dose [cGy]: 5634.2
Mean Dose [cGy]: 707.0
Modal Dose [cGy]: 99.5

STD [cGy]: 917.2

Equiv. Sphere Diam. [cm]: N/A

Conformity Index: N/A Gradient Measure [cm]: N/A

Median Dose [cGy]: 276.4

```
Dose [cGy]
              Relative dose [%] Ratio of Total Structure Volume [%]
                                                          100
         0
         5
                       0.0909091
                                                          100
        10
                                                          100
                        0.181818
        15
                        0.272727
                                                          100
        20
                                                          100
                        0.363636
        25
                        0.454545
                                                          100
        30
                        0.545455
                                                          100
        35
                        0.636364
                                                          100
        40
                        0.727273
                                                          100
        45
                        0.818182
                                                     99.9247
        50
                        0.909091
                                                     99.2638
        55
                                                     98.1752
        60
                         1.09091
                                                     96.8538
        65
                         1.18182
                                                     95.3989
        70
                         1.27273
                                                     93.8907
        75
                                                     92.3371
                         1.36364
        80
                                                     90.7697
                         1.45455
                                                     89.2061
        85
                         1.54545
        90
                         1.63636
                                                     87.6496
```

. .

This DVH data may be imported using the read.DVH() function, with an example shown here:

```
> read.DVH(file="Jane_Doe.dvh", type="aria10", verbose=TRUE)
```

Reading DVH file ('/private/var/folders/ll/krvs9rds693bhfr0tm22jnb40000gn/T/RtmphdLpkH/R Patient: Doe, Jane (111111111)
Plan: PLAN_NAME

Dose: 5500cGy (at 100% isodose line)

..Importing structure: LIVER [volume: 1635.9cc, dose: 42.7 - 5634.2cGy]

..Importing structure: LEFT_KIDNEY [volume: 195.7cc, dose: 75.8 - 3846.8cGy]

..Importing structure: STOMACH [volume: 695.2cc, dose: 59 - 5353.2cGy]

..Importing structure: DUODENUM [volume: 34.2cc, dose: 2707.8 - 5620.1cGy]

..Importing structure: RIGHT_KIDNEY [volume: 223.9cc, dose: 102.4 - 4201.9cGy]

..Importing structure: CTV [volume: 146.7cc, dose: 5168.6 - 5646.9cGy] ..Importing structure: PTV [volume: 239.4cc, dose: 4749.8 - 5664.7cGy]

..Importing structure: SMALL_BOWEL [volume: 232.2cc, dose: 59.6 - 4934.1cGy]

..Importing structure: CORD [volume: 64.9cc, dose: 0 - 3442.8cGy]

.. Importing structure: BODY [volume: 25507.5cc, dose: 0 - 5664.7cGy]

Multiple files can be imported simultaneously, generating a list of DVH.list objects:

```
> DVHs <- read.DVH(file=c("Jane_Doe.dvh", "John_Doe.dvh"), type="aria10")
Reading DVH file ('/private/var/folders/11/krvs9rds693bhfr0tm22jnb40000gn/T/RtmphdLpkH/R
 Patient: Doe, Jane (111111111)
 Plan: PLAN_NAME
 Dose: 5500cGy (at 100% isodose line)
  .. Importing structure: LIVER [volume: 1635.9cc, dose: 42.7 - 5634.2cGy]
  .. Importing structure: LEFT_KIDNEY [volume: 195.7cc, dose: 75.8 - 3846.8cGy]
  .. Importing structure: STOMACH [volume: 695.2cc, dose: 59 - 5353.2cGy]
  .. Importing structure: DUODENUM [volume: 34.2cc, dose: 2707.8 - 5620.1cGy]
  ..Importing structure: RIGHT_KIDNEY [volume: 223.9cc, dose: 102.4 - 4201.9cGy]
  .. Importing structure: CTV [volume: 146.7cc, dose: 5168.6 - 5646.9cGy]
  .. Importing structure: PTV [volume: 239.4cc, dose: 4749.8 - 5664.7cGy]
  ..Importing structure: SMALL_BOWEL [volume: 232.2cc, dose: 59.6 - 4934.1cGy]
  .. Importing structure: CORD [volume: 64.9cc, dose: 0 - 3442.8cGy]
  .. Importing structure: BODY [volume: 25507.5cc, dose: 0 - 5664.7cGy]
Reading DVH file ('/private/var/folders/11/krvs9rds693bhfr0tm22jnb40000gn/T/RtmphdLpkH/R
 Patient: Doe, John (555555555)
 Plan: PLAN_NAME
 Dose: 5500cGy (at 100% isodose line)
  .. Importing structure: LIVER [volume: 1366.8cc, dose: 0 - 92.9cGy]
  ..Importing structure: SMALL_BOWEL [volume: 206.2cc, dose: 0 - 99.8cGy]
  .. Importing structure: DUODENUM [volume: 93.1cc, dose: 0 - 102.4cGy]
  ..Importing structure: STOMACH [volume: 303.6cc, dose: 0 - 101.3cGy]
  .. Importing structure: CTV [volume: 88.4cc, dose: 96.8 - 102.6cGy]
  ..Importing structure: PTV [volume: 155.7cc, dose: 84.1 - 102.6cGy]
  .. Importing structure: BODY [volume: 17666.2cc, dose: 0 - 102.6cGy]
  .. Importing structure: LEFT_KIDNEY [volume: 154.2cc, dose: 0 - 44.4cGy]
  ..Importing structure: RIGHT_KIDNEY [volume: 155.1cc, dose: 0 - 98.5cGy]
  ..Importing structure: CORD [volume: 40.7cc, dose: 0 - 55.5cGy]
```

[1] "List containing 20 DVH objects (LIVER, LEFT_KIDNEY, STOMACH, DUODENUM, RIGHT_KIDNEY

3.2 DVH calculation from dose grid

Alternatively, read.DVH(..., method="dicom") can be employed to extract DVH data from a DICOM-RT dataset including a dose grid (calculated from the treatment planning system) and one or more structure set(s). The DVH import implicitly calls the calculate.DVH() function without the user having to manually specify dose calculation, however the calculate.DVH() function can also be called explicitly to calculate or recalculate DVHs using specific input parameters (generally the user will not need to call this function explicitly). In any of the above cases, DVH calculation from dose grid data utilizes the method previously described by Straube et al. with trilinear interpolation of dosimetric information (3). Please note that this is a computationally intensive process and has not yet been optimized in the current release (future releases will include speed-ups of this dose calculation).

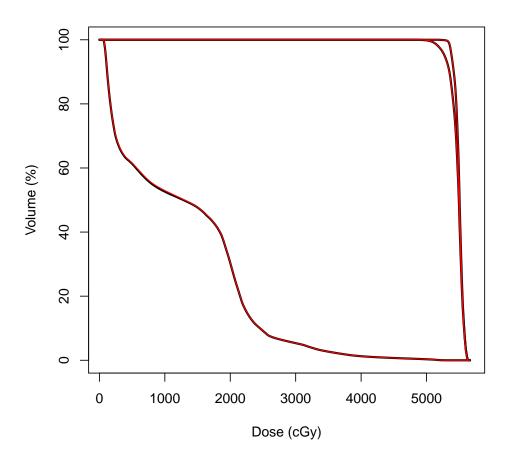


Figure 1: Comparison of DVH data for patient Jane Doe, calculated from dose grid data using the *RadOnc* package (black) or otherwise imported as pre-calculated DVH from the treatment planning system (red).

3.3 DVH list manipulation

The read.DVH() function returns a DVH list that can be manipulated in multiple ways. Subsets of DVH lists can be obtained using the subset() function or [] modifier, and any number of DVH lists can be combined using the c() function. Additionally, single DVH objects can be directly accessed using the [[]] modifier, and individual elements of a DVH list may be directly replaced with other DVH objects using the [[<- function.

```
> janedoe[1:4]
[1] "List containing 4 DVH objects (LIVER, LEFT_KIDNEY, STOMACH, DUODENUM)"
> c(janedoe["PTV"], johndoe[c("CTV", "DUODENUM")])
[1] "List containing 3 DVH objects (PTV, CTV, DUODENUM)"
> johndoe[["CTV"]]
[1] "Structure: CTV (88.4cc), Dose: 1.76-1.87% (5500cGy prescribed), DVH: cumulative, Vo
> janedoe[[1]] <- johndoe[["CTV"]]</pre>
> janedoe[1:4]
[1] "List containing 4 DVH objects (CTV, LEFT_KIDNEY, STOMACH, DUODENUM)"
```

Note that pattern matching (using regular expressions) may also be employed to select a subset of DVH elements from a DVH list:

```
> janedoe["KIDNEY$"]
```

- [1] "List containing 2 DVH objects (LEFT_KIDNEY, RIGHT_KIDNEY)"
- > janedoe[c(2,"IGHT.*")]
- [1] "List containing 2 DVH objects (LEFT_KIDNEY, RIGHT_KIDNEY)"
- > janedoe["(?i)liver"] ## (case insensitive matching) ##
- [1] "List containing 1 DVH objects (LIVER)"

Additionally, the subset() function has been extended to support DVH lists. In particular, a user may specify which patient(s) to include (pattern matching is applied to patient names), which structures (pattern matching for structures as described above), and/or which DVHs according to specific dose constraints (e.g. select all structures satisfying a specific parameter):

```
> subset(janedoe, structure=c("liver", "kidney"), ignore.case=TRUE)
```

[1] "List containing 3 DVH objects (LIVER, LEFT_KIDNEY, RIGHT_KIDNEY)"

```
> subset(janedoe, constraint="volume > 200(cc)")
```

- [1] "List containing 6 DVH objects (LIVER, STOMACH, RIGHT_KIDNEY, PTV, SMALL_BOWEL, BODY
- > subset(janedoe, constraint="Dmean <= 500(cGy)")
- [1] "List containing 2 DVH objects (SMALL_BOWEL, BODY)"
- > subset(janedoe, constraint="D2cc < 20(Gy)")</pre>
- [1] "List containing O DVH objects ()"

cGy

0.00000000

Other list processing functions may be applied to DVH lists, enabling further data manipulation. The rev() function may be used to reverse the order of a DVH list, while the names() function may be used to extract (or set) the structure names for each DVH contained within the list. The length() function may be used to find the number of DVH objects contained within a DVH list, and the lapply() function can be used to perform a customizable set of operations on a DVH list and return a customizable set of values. Here are some examples employing each of these functions:

```
> names(janedoe)[1:4] <- c("A1", "B2", "C3", "D4")</pre>
> names(rev(janedoe[1:4]))
[1] "D4" "C3" "B2" "A1"
> length(johndoe)
[1] 10
> lapply(johndoe, function(DVH) { DVH[c("DMIN", "D50%", "DMAX", "V20%")] })
$LIVER
         сGу
                                    cGy
                                                   CC
               0.07174856 92.90000000 185.28500000
  0.00000000
$SMALL_BOWEL
        cGy
                      %
                                 cGy
                                              СС
 0.00000000 0.06320805 99.80000000 13.94640000
$DUODENUM
                           cGy
  0.00000 81.34012 102.40000 83.10130
$STOMACH
```

cGy

0.06726741 101.30000000 31.95860000

СС

\$CTV

cGy % cGy cc 96.8000 100.0081 102.6000 88.4095

\$PTV

cGy % cGy cc 84.10000 99.80143 102.60000 155.73500

\$BODY

cGy % cGy cc 0.000000e+00 6.202236e-02 1.026000e+02 1.893130e+03

\$LEFT_KIDNEY

cGy % cGy cc 0.00000000 0.07279133 44.40000000 18.60640000

\$RIGHT_KIDNEY

cGy % cGy cc 0.00000 24.25567 98.50000 85.93270

\$CORD

cGy % cGy cc 0.000000 4.078601 55.500000 17.496100

Patient name(s) and identifier(s) may be extracted from each DVH within the DVH list using the \$ modifier:

> janedoe[1:2]\$patients

\$LIVER

"Doe, Jane"

\$LEFT_KIDNEY

"Doe, Jane"

> janedoe[3:4]\$ID

\$STOMACH

"111111111"

\$DUODENUM

"111111111"

Note: for nested DVH list data generated using read.DVH(..., collapse=FALSE) on multiple input files, the as(.., Class="DVH.list") command can convert this list to a single DVH.list object. Subsequent structure processing may then proceed, as above.

3.4 DVH data

Each DVH structure contains a variety of data related to the structure itself as well as the distribution of radiation dose within the structure volume. Detailed slot list and parameters are described in the DVH-class documentation accompanying the RadOnc package. Specific parameters can be extracted using the [] modifier, which can take as its argument a character string representation of the desired dose/volume parameter. For instance, the volume of duodenum receiving 20Gy or the dose to the top 2.5% (2.3286cc) of the volume can be extracted from DVH data as follows:

These parameters are entirely flexible and may even specify a summarized range of dosimetric values, for instance "V10-20Gy" specifies the volume of a structure receiving between 10 and 20Gy. In addition, multiple parameters can be requested for a given DVH object at the same time. This functionality can also be applied to a DVH list using the \$ modifier.

```
> johndoe[["DUODENUM"]][c("V5%", "V20Gy", "D2.5%", "D2.3286cc", "Dmax")]
                                          cGv
      СС
               СС
 90.1912 76.5153 100.5605 100.5605 102.4000
> johndoe[1:4]$"V10-20Gy, V20Gy, Dmax"
$LIVER
                           cGy
       СС
                 СС
172.61312 21.35734
                     92.90000
$SMALL_BOWEL
                          cGy
       СС
                 СС
```

5.257440 9.370469 99.800000

\$DUODENUM

cc cc cGy 7.366491 76.515300 102.400000

\$STOMACH

cc cc cGy 5.85510 26.83795 101.30000

Note that specific parameter keywords include: "Dmax" (maximum dose), "Dmin" (minimum dose), "Dmean" (mean dose), "Dintegral" (estimated integral dose), "DRx" (prescription dose), and "volume" (total structure volume). If an improper parameter is specified however, NA results will be returned for the affected parameter(s):

Parameters may be further modified by "()" tags, which can alter the output format of data in the case of standard dosimetric parameters, for instance:

Integral dose presents a special case, where additional tags can specify the range over which the integral dose is calculated (default is over the entire dose range).

> johndoe[["LIVER"]][c("Dintegral(<20Gy)", "Dintegral(10-20Gy)")]

Gy*cc Gy*cc 3071.012 2493.167

3.5 Generalized equivalent uniform dose (gEUD) calculation

Generalized equivalent uniform doses (gEUD) can be calculated from one or more DVH or DVH.list objects using the gEUD() function. This calculation summarizes the entire DVH as a single dose which, when distributed evenly thorughout the structure volume, is biologically iso-effective with the heterogeneously distributed dose represented by the DVH (4). The calculation depends upon a user-specified tissue-specific parameter (a), which can account for behavior of both tumors and normal tissues. This parameter should be negative for target structures (e.g. tumor) and positive for organs at risk. For a = 1, the gEUD is equivalent to mean dose, while for a = Inf and a = -Inf, the gEUD is equivalent to maximum and minimum doses, respectively.

```
> gEUD(janedoe[1:3], 6:8)
      LIVER LEFT_KIDNEY
                            STOMACH
   2538.873
               1997.308
                           2842.618
> gEUD(janedoe[1:3], 1) == unlist(janedoe[1:3]$"Dmean")
      LIVER LEFT_KIDNEY
                            STOMACH
       TRUE
                   TRUE
                               TRUE
> gEUD(janedoe[1:3], Inf) == unlist(janedoe[1:3]$"Dmax")
      LIVER LEFT_KIDNEY
                            STOMACH
       TRUE
                   TRUE
                               TRUE
> gEUD(janedoe[1:3], -Inf) == unlist(janedoe[1:3]$"Dmin")
      LIVER LEFT_KIDNEY
                            STOMACH
       TRUE
                   TRUE
                               TRUE
```

3.6 Linear quadratic extrapolated (LQE) dose conversion

Linear quadratic extrapolation (LQE) has been used to model iso-effective doses (5) and is implemented herein for conversion of and comparison among one or more DVH or DVH.list objects to account for varying dose fractionation. Two identical DVH curves may in fact represent two widely divergent treatments if, for instance, one patient is treated to 6000cGy in four 1500cGy fractions whereas another is treated to 6000cGy in thirty 200cGy fractions. In the case below, the LQE() function is used to determine the isoeffective doses for 4500, 5500, and 6000cGy when delivered in 200cGy instead of 300cGy daily fractions (20x 300cGy is iso-effective with 36x 200cGy, assuming an alpha/Beta ratio of 3 [aB = 3]):

```
> LQE(c(4500, 5500, 6000), aB=3, fractions=c(300, 200))
[1] 5400 6600 7200
> LQE(c(4500, 5500, 6000), aB=3, N=c(20, 36))
[1] 5286.908 6558.679 7200.000
```

Note that the LQE() function may also be applied to DVH or DVH.list objects, wherein doses are converted according to specified fractionation and accounting for user-specified alpha/Beta ratio.

3.7 DVH plotting

Individual DVH plots can be generated by the plot() function, and may be altered to show dose and/or volume as relative or absolute values with DVH shown as cumulative or differential data.

> plot(janedoe[[3]], volume="relative", dose="absolute", type="cumulative")

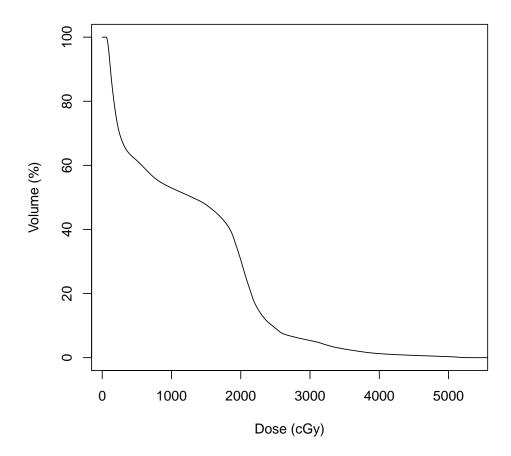


Figure 2: Standard dose-volume histogram for a single structure ("STOMACH") from patient Jane Doe. Data is shown as cumulative dose versus volume.

```
> plot(janedoe[1:3], plot.type="i", col=c("red", "green", "blue"),
+ legend="topright", legend.labels=names(janedoe[1:3]))
```

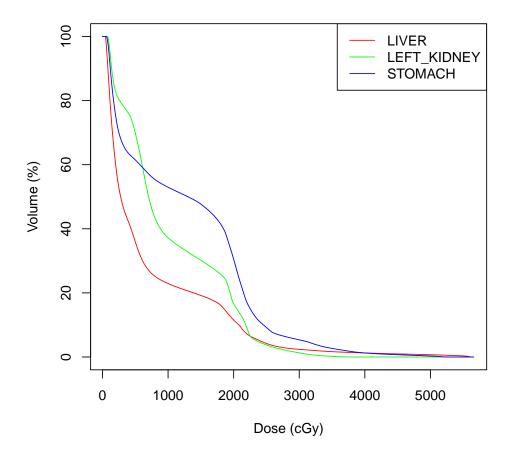


Figure 3: Standard dose-volume histogram for three structures from a single patient, Jane Doe. Data is shown as cumulative dose versus volume. Legend is displayed in the top right corner of the plot.

```
> plot(c(johndoe["STOMACH"], janedoe["STOMACH"]), #group 1
+ c(janedoe["LIVER"], johndoe["LIVER"]), #group 2
+ c(johndoe["DUODENUM"], janedoe["DUODENUM"]), #group 3
+ plot.type="g", dose="relative", col=c("blue", "red", "green"),
+ lwd=2, lty="dashed", fill.lty="solid", fill.transparency=0.3)
```

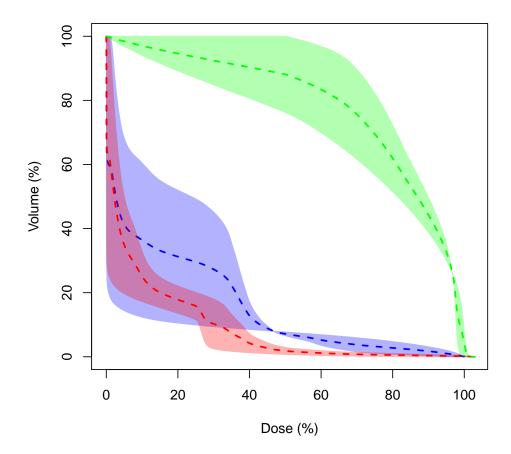


Figure 4: Mean dose-volume histograms are shown for three groups of DVHs, in this case corresponding to stomach, liver, and duodenum from two different patients (John Doe and Jane Doe). Data is shown as cumulative dose (relative) versus volume (relative). Shading represents the range of the data for each group (note that the width of the shading can be specified to represent other parameters instead of range – e.g. variance, standard deviation, interquartile range, median absolute deviation).

```
> group1 <- c("CTV", "PTV")
> group2 <- c("LIVER", "STOMACH", "SMALL_BOWEL")
> plot(c(johndoe[group1],janedoe[group1]),
+ c(janedoe[group2],johndoe[group2]),
+ plot.type="t", main="Target v. OAR t-Test", alpha=0.001,
+ col=c("red", "blue"), lty="dashed", fill.lty="solid")
```

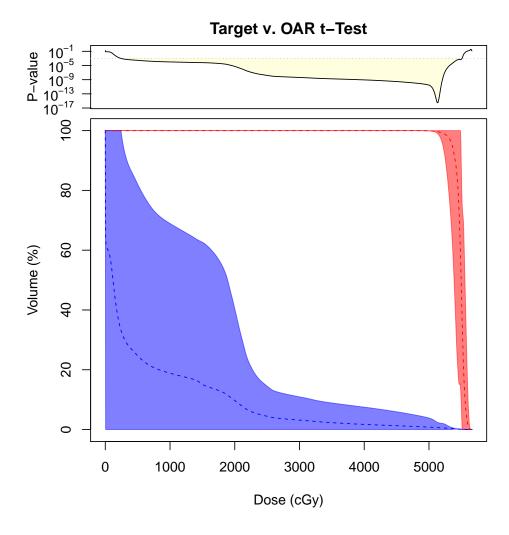


Figure 5: Mean dose-volume histograms are shown for two groups of DVHs, in this case corresponding to CTV/PTV and liver/stomach/small bowel from two different patients (John Doe and Jane Doe). Data is shown as cumulative dose (absolute) versus volume (relative). Shading represents the 99.9% confidence interval for each group (specified here by alpha=0.001). The corresponding p-values are shown in the upper panel, with corresponding significance threshold p<0.001.

> plot(janedoe[2:9], plot.type="b", volume="abs", dose="rel")

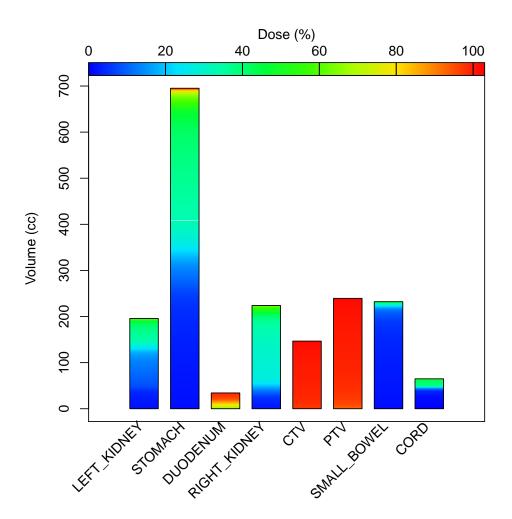


Figure 6: Bar representation of dose distributions for eight structures from a single patient (Jane Doe).

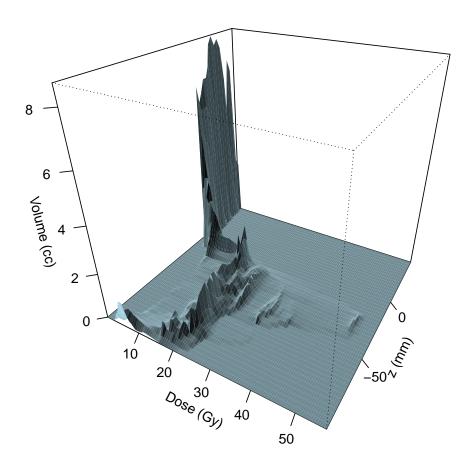


Figure 7: Axially-segmented DVH (zDVH) displayed as a differential DVH in a three dimensional projection. Plotting in this format simply requires input of a single zDVH object.

3.8 DVH statistics

Mean or median DVHs can be calculated using the mean() and median() functions, respectively. These functions take a DVH list as input and return a single object of class DVH representing the mean or median dose-volume histogram data calculated from the entire group.

```
> plot(janedoe)
> plot(median(janedoe), new=FALSE, col="red", lwd=2)
> plot(mean(janedoe), new=FALSE, col="blue", lwd=2, lty="dashed")
```

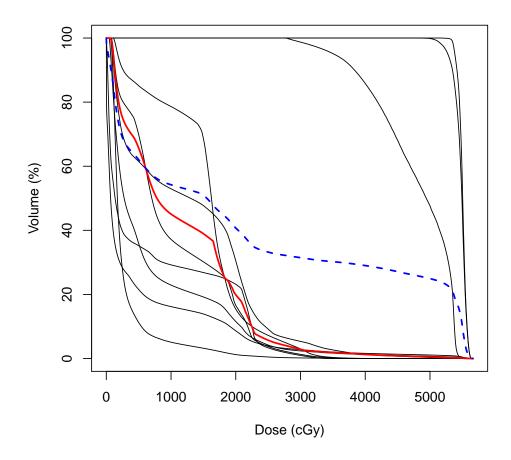


Figure 8: Mean and median DVHs are shown in blue dash and red, respectively.

Similarly, DVH list summation can be performed for two or more *non-overlapping* structures (e.g. bilateral lungs or kidneys) using the sum() function.

```
> L.kidney <- janedoe[["LEFT_KIDNEY"]]
> R.kidney <- janedoe[["RIGHT_KIDNEY"]]
> total.kidney <- sum(janedoe[c("LEFT_KIDNEY", "RIGHT_KIDNEY")])
> plot(total.kidney, type="diff", volume="abs")
> plot(L.kidney, new=FALSE, type="diff", volume="abs", col="red")
> plot(R.kidney, new=FALSE, type="diff", volume="abs", col="blue")
```

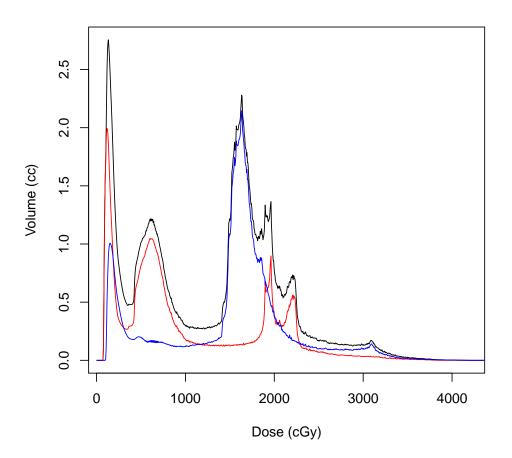


Figure 9: Plot of total kidney dose, calculated using sum() function. Left and right kidney doses are shown in red and blue, respectively.

In routine clinical practice and research, DVH comparisons are often performed at an individual parameter level (e.g. V20Gy from Group A compared to Group B). The *RadOnc* package enables automated comparison throughout the entire DVH. Functions such as t.test() and wilcox.test() are both enabled for DVH lists.

```
> AvB <- t.test(groupA, groupB)
> plot(AvB$dose, AvB$p, type="1", log="y", xlab="Dose (cGy)", ylab="p-value")
```

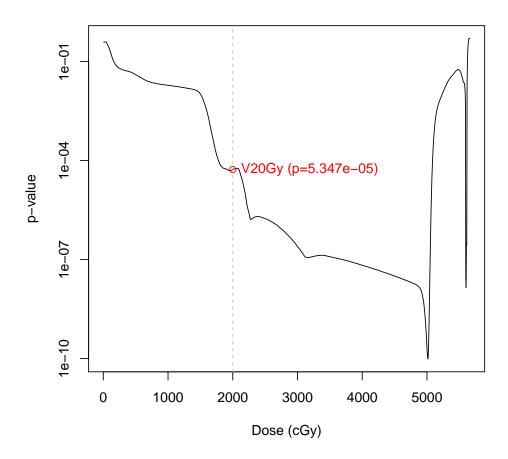


Figure 10: p-values from t.test() comparison as a function of dose. V20Gy is highlighted and its p-value corresponds closely to values generated from t-test of V20Gy directly.

4 Three-Dimensional Structure Analysis

4.1 DICOM-RT import

The read.DICOM.RT() function is designed to take input DICOM-RT file(s) and output a list of structure3D data objects containing all relevant data, in particular the axially-defined contours delineating each structure. Note that DICOM-RT file import was evaluated using Varian's Aria/Eclipse platform (v.10 and v.11). Other treatment planning systems may encode 3D structural information in a different format and this has not been evaluated in the current release of software.

For DICOM-RT data, the associated CT scan must be exported directly from the treatment planning system and should include all contoured structures of interest. In Eclipse, this can be accomplished via the "Export Wizard..." option in the "File" menu, accessed in either Countouring or Plan Evaluation modes. Note that the "Include structure set" option should be selected, and that Institution-specific filters will be required for proper data export. DICOM-RT data will consist of multiple files representing both the CT image as well as the relevant structure set(s).

DICOM-RT data may be imported using the read.DICOM.RT() function, with a mockup example shown here (note that "DICOM directory" should be replaced by the path to a specific directory containing the desired DICOM data):

> data <- read.DICOM.RT(path="<<DICOM directory>>", verbose=TRUE)

The DICOM-RT import process may take some time. We have included pre-loaded data for a single patient (included structures: spinal cord, mandible, teeth) which will be explored in this vignette.

4.2 3D structure manipulation

The read.DICOM.RT() function returns a structure.list object that can be manipulated in multiple ways. Subsets of structure lists can be obtained using the [] modifier, and any number of structure lists can be combined using the c() function. Additionally, single structure3D objects can be directly accessed using the [[]] modifier, and individual elements of a structure list may be directly replaced with other structure3D objects using the [[<- function.

```
> teeth[1:2]
[1] "List containing 2 structure3D objects (Tooth #1, Tooth #2)"
> c(cord, mandible)
[1] "List containing 2 structure3D objects (Spinal Cord, Mandible)"
> teeth[[1]]
[1] "Structure (Tooth #1) defined by 324 points in 23 axial slices"
> teeth[[1]] <- teeth[["Tooth #3"]]
> teeth
[1] "List containing 3 structure3D objects (Tooth #3, Tooth #2, Tooth #3)"
```

As with DVH.list objects, structure.list objects may be subsetted by pattern matching:

```
> teeth["Tooth.*"]
```

[1] "List containing 3 structure3D objects (Tooth #1, Tooth #2, Tooth #3)"

Other list processing functions may be applied to structure lists, enabling further data manipulation. The rev() function may be used to reverse the order of a structure list, while the names() function may be used to extract (or set) the structure names for each structure contained within the list. The length() function may be used to find the number of structures contained within a structure list, and the lapply() function can be used to perform a customizable set of operations on a structure list and return a customizable set of values. Here are some examples employing each of these functions:

```
> names(teeth) <- c("Larry", "Curly", "Moe")
> names(rev(teeth[1:3]))
[1] "Moe" "Curly" "Larry"
> length(teeth)
[1] 3
```

> lapply(teeth, function(tooth) { range(tooth) })

\$Larry

\$Curly

\$Moe

4.3 Plotting 3D structures

Three dimensional surfaces renderings can be generated by the plot() function. The RadOnc package does not currently contain the functionality to generate surface triangulations for a given structure, however future releases of the package will implement surface triangulation. Thus, data imported using read.DICOM.RT() will not currently be processed for surface triangulation and will generate an empty plot if plotting is attempted. External applications such as MeshLab (6) can be used to generate triangulations which, for advanced users, can be imported into a given structure3D object.

> plot(mandible)

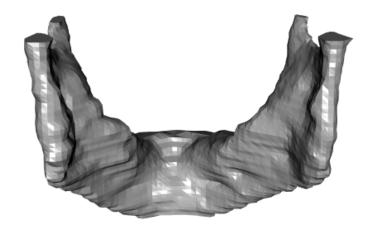


Figure 11: Three-dimensional surface reconstruction from triangulation of physician-contoured points for the mandible of a patient.

> plot(cord)



Figure 12: Three-dimensional surface reconstruction from triangulation of physician-contoured points for the spinal cord of a patient.

4.4 Structure comparison

Comparison of three-dimensional structures has numerous applications. In the case presented here, three physicians separately delineated a tooth on axial slices of a CT for a single patient. Variability among physician contours is demonstrated using the compareStructures() function:

> compareStructures(teeth, method="axial", plot=TRUE)

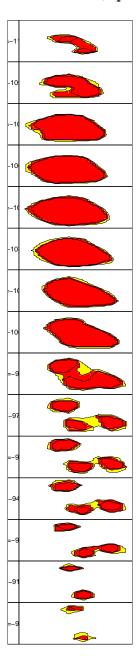


Figure 13: Axial comparison of overlap among three separate physician-defined contours for a single tooth. Red (and white) regions delineate consensus while decreasing degree of overlap is shown in decreasing shades of orange and yellow.

Structure comparison may also be performed by Hausdorff distance (7), which computes the distance between two points clouds, in this case structure surfaces. The absolute Hausdorff distance (hausdorff.method="absolute") yields the maximum distance required to connect any point from one point cloud to its closest neighbor in the other. This metric is highly subject to outliers, thus an aggregate metric is implemented by selecting the average distance (hausdorff.method="mean" or hausdorff.method="median") required to connect all points in one point cloud to their closest neighboring points in the other. Note that the Hausdorff distance between two completely superimposable point clouds is zero.

```
> compareStructures(teeth, method="hausdorff", hausdorff.method="mean")
```

```
Analyzing structure 1/2 (Tooth #1) ... FINISHED
Analyzing structure 2/2 (Tooth #3) ... FINISHED
Tooth #1 Tooth #3
Tooth #1 0.0000000 0.4847732
Tooth #3 0.4847732 0.0000000
```

Pairwise Dice similarity coefficients (DSC) may also be calculated (8):

> compareStructures(teeth, method="DSC")

```
Tooth #1 Tooth #2 Tooth #3
Tooth #1 1.0000000 0.8977074 0.8830091
Tooth #2 0.8977074 1.0000000 0.8786020
Tooth #3 0.8830091 0.8786020 1.0000000
```

References

- [1] R.E. Drzymala, R. Mohan, L. Brewster, J. Chu, M. Goitein, W. Harms, and M. Urie. Dose-volume histograms. *Int J Radiat Oncol Biol Phys*, 21(1):71–78, 1991.
- [2] R.F. Thompson. Radonc: an r package for analysis of dose-volume histogram and three-dimensional structural data. *JROI*, 6(1):71–83, 2014.
- [3] W. Straube, J. Matthews, W. Bosch, and J.A. Purdy. Dvh analysis: consequences for quality assurance of multi-institutional clinical trials. *Med Phys*, 32(6):2021, 2005.
- [4] C. Thieke, T. Bortfeld, A. Niemierko, and S. Nill. From physical dose constraints to equivalent uniform dose constraints in inverse radiotherapy planning. *Med Phys*, 30(9): 2332–2339, 2003.
- [5] G.W. Barendsen. Dose fractionation, dose-rate and iso-effect relationships for normal-tissue response. *Int J Radiat Oncol Biol Phys*, 8(11):1981–1997, 1982.
- [6] Open Source actively supported by the 3D-CoForm project. Meshlab. 2005. URL http://meshlab.sourceforge.net.
- [7] Felix Hausdorff. Grundzuge der Mengenlehre. Veit and Company, Leipzig, 1914.
- [8] L.R. Dice. Measures of the amount of ecologic association between species. *Ecology*, 26: 297–302, 1945.

A Acknowledgements

Many thanks to Brandon Whitcher for his phenomenal and generous support regarding the oro.dicom pacakge, and to Daniel Wollschlaeger for his insights and contributions regarding DVH file formatting, dose unit interconversion, and inclusion of CadPlan data. Thank you to Francis Gibbons, Usman Lula, Michael Young, and Lukasz Matulewicz for their contributions regarding inclusion of Elekta, RayStation, and TomoTherapy-formatted data. Thank you to Maura Kirk, Eric Ojerholm, and Abodunde Adegunloye for their help in identifying software bugs and suggesting additional package features.

B Previous Release Notes

- Initial release (v1.0.0) on 2013-07-07
- Subsequent release (v1.0.1) on 2013-10-23
- Subsequent release (v1.0.2) on 2013-11-25
- Subsequent release (v1.0.3) on 2014-01-17
- Subsequent release (v1.0.4) on 2014-01-31
- Subsequent release (v1.0.5) on 2014-02-19
- Subsequent release (v1.0.6) on 2014-02-20
- Subsequent release (v1.0.7) on 2014-02-27
- Subsequent release (v1.0.8) on 2014-05-31
- Subsequent release (v1.0.9) on 2014-11-22
- Subsequent release (v1.1.0) on 2015-04-22
- Last release (v1.1.1) on 2016-01-13

Please refer to NEWS.Rd file for more details.

> news(package="RadOnc")