Package ‘growthcurver’

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Type Package

Title Simple Metrics to Summarize Growth Curves

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Description Fits the logistic equation to microbial growth curve data (e.g., repeated absorbance measurements taken from a plate reader over time). From this fit, a variety of metrics are provided, including the maximum growth rate, the doubling time, the carrying capacity, the area under the logistic curve, and the time to the inflection point. Method described in Sprouffske and Wagner (2016) <doi:10.1186/s12859-016-1016-7>.

LazyData TRUE

Depends R (>= 4.0)

Imports minpack.lm (>= 1.2), stats (>= 4.0), graphics (>= 4.0), grDevices (>= 4.0)

URL https://github.com/sprouffske/growthcurver

BugReports https://github.com/sprouffske/growthcurver/issues

License GPL (>= 2)

Suggests testthat, knitr, dplyr, ggplot2, rmarkdown

VignetteBuilder knitr

RoxygenNote 7.1.1

NeedsCompilation no

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R topics documented:

`gcfit` .......................... 2
`gcvals` .......................... 3
`growthdata` .......................... 4
`NAiT` .......................... 7
`SummarizeGrowth` .......................... 7
`SummarizeGrowthByPlate` .......................... 9

Index 12

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gcfit .............................. Creates an object of class gcfit.

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Description

This is a constructor function for the "gcfit" class. This class is most often obtained as the return value when calling `SummarizeGrowth`.

Usage

`gcfit(gc_vals, log_mod, data_t, data_n)`

Arguments

`gc_vals`  An object of class gcvals that contains the summarized metrics from fitting the growth model to a set of experimental observations. This is where the fitness proxy parameters can be found. See `gcvals` for more information the information found in this object.

`log_mod`  An object of class nlsModel that contains the results of fitting the logistic growth model to the data

`data_t`  A numeric vector of times

`data_n`  A numeric vector of cell count or absorbance readings

Value

An object of class gcfit, which is a list of three objects, that combines the parameters (vals = gc_vals, model = log_mod, data = list(data_t, data_n))
gcvals

Creates an object of type gcvals.

Description

Constructor function for the "gcvals" class. This object is most often obtained when calling SummarizeGrowth (it is the first parameter in the gcvals object).

Usage

```r
gcvals(
  k,
  k_se,
  k_p,
  n0,
  n0_se,
  n0_p,
  r,
  r_se,
  r_p,
  sigma,
  df,
  t_mid,
  dt,
  auc_l,
  auc_e,
  note
)
```

Arguments

- **k**: The carrying capacity parameter
- **k_se**: The standard error of the carrying capacity parameter
- **k_p**: The p value of the carrying capacity parameter
- **n0**: The initial population size
- **n0_se**: The standard error of the initial population size
- **n0_p**: The p value of the initial population size
- **r**: The growth rate
- **r_se**: The standard error of the growth rate
- **r_p**: The p value of the growth rate
- **sigma**: Residual standard error from non-linear least squares fit of the model to the data
- **df**: Degrees of freedom
- **t_mid**: The time at the inflection point of the logistic curve (occurs at half of the carrying capacity)
dt

The maximum doubling time, obtained by evaluating the unrestrained growth of the population with growth rate r

auc_l

The area under the curve of the fitted logistic equation from time 0 to time t

auc_e

The area under the curve of the measurements.

note

Feedback on common problems with fitting the logistic curve to the data

Value

An object of class gcvals.

---

growthdata  Simulated growth curve data

Description

A dataset containing absorbance measurements over time of microbes growing in a plate reader for 1 day. The growth curves for a whole plate are included.

Usage

growthdata

Format

A data frame with 145 observations and 97 variables:

time  time, in hours
A1 absorbance readings of well A1
A2 absorbance readings of well A2
A3 absorbance readings of well A3
A4 absorbance readings of well A4
A5 absorbance readings of well A5
A6 absorbance readings of well A6
A7 absorbance readings of well A7
A8 absorbance readings of well A8
A9 absorbance readings of well A9
A10 absorbance readings of well A10
A11 absorbance readings of well A11
A12 absorbance readings of well A12
B1 absorbance readings of well B1
B2 absorbance readings of well B2
B3 absorbance readings of well B3
growthdata

B4 absobance readings of well B4
B5 absorbance readings of well B5
B6 absorbance readings of well B6
B7 absorbance readings of well B7
B8 absorbance readings of well B8
B9 absorbance readings of well B9
B10 absorbance readings of well B10
B11 absorbance readings of well B11
B12 absorbance readings of well B12
C1 absorbance readings of well C1
C2 absorbance readings of well C2
C3 absorbance readings of well C3
C4 absorbance readings of well C4
C5 absorbance readings of well C5
C6 absorbance readings of well C6
C7 absorbance readings of well C7
C8 absorbance readings of well C8
C9 absorbance readings of well C9
C10 absorbance readings of well C10
C11 absorbance readings of well C11
C12 absorbance readings of well C12
D1 absorbance readings of well D1
D2 absorbance readings of well D2
D3 absorbance readings of well D3
D4 absorbance readings of well D4
D5 absorbance readings of well D5
D6 absorbance readings of well D6
D7 absorbance readings of well D7
D8 absorbance readings of well D8
D9 absorbance readings of well D9
D10 absorbance readings of well D10
D11 absorbance readings of well D11
D12 absorbance readings of well D12
E1 absorbance readings of well E1
E2 absorbance readings of well E2
E3 absorbance readings of well E3
E4 absorbance readings of well E4
<table>
<thead>
<tr>
<th>Well</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>E5</td>
<td>absorbance readings of well E5</td>
</tr>
<tr>
<td>E6</td>
<td>absorbance readings of well E6</td>
</tr>
<tr>
<td>E7</td>
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<td>absorbance readings of well G12</td>
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<tr>
<td>H5</td>
<td>absorbance readings of well H5</td>
</tr>
</tbody>
</table>
NAT\(T\) absorbance readings of well H6
H7 absorbance readings of well H7
H8 absorbance readings of well H8
H9 absorbance readings of well H9
H10 absorbance readings of well H10
H11 absorbance readings of well H11
H12 absorbance readings of well H12

---

### Description

This function gives the number of cells or absorbance (\(N\)) at time \(t\) when the parameters to the logistic equation are \(K\), \(N_0\), and \(r\).

### Usage

\[NAT\(T(k, n_0, r, t)\]

### Arguments

- \(k\) The carrying capacity
- \(n_0\) The initial population size (absorbance or individuals)
- \(r\) The exponential "growth rate"
- \(t\) The time at which you want to know \(N\)

### Value

The number of cells, or \(N\), at time \(t\)

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### SummarizeGrowth

Summarize Growth Curves

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### Description

This function finds the parameters that describe the input data’s growth. It does so by fitting the logistic curve to your growth curve measurements.

### Usage

\[\text{SummarizeGrowth(data}_t, \text{data}_n, t_\text{trim} = 0, \text{bg}_\text{correct} = \text{"min"}, \text{blank} = \text{NA})\]
Arguments

data_t A vector of timepoints (data_n must also be provided and be the same length).
data_n A vector of cell counts or absorbance readings.
t_trim Measurements taken after this time should not be included in fitting the curve. If stationary phase is variable, this may give you a better fit. A value of 0 means no trimming. Defaults to no trimming (0).
bg_correct The background correction method to use. No background correction is performed for the default "none". Specifying "min" subtracts the smallest value in a column from all the rows in that column, and specifying "blank" subtracts the values from the blank vector from the data_n vector.
blank A vector of absorbance readings from a blank well (typically contains only media) used for background correction. The corresponding blank value is subtracted from the data_n vector for each timepoint. Defaults to NA.

Details

The logistic curve equation is

\[ N_t = \frac{N_0 K}{N_0 + (K - N_0)e^{-rt}} \]

where \( N_t \) is the number of cells (or the absorbance reading) at time \( t \), \( N_0 \) is the initial cell count (or absorbance reading), \( K \) is the carrying capacity, and \( r \) is the growth rate.

The fitness proxies returned are the parameters of the logistic equation and the area under the curve (a measure that integrates the effects of \( N_0 \), \( K \), and \( r \)). See gcfit for more documentation on these.

Value

An object of type gcfit containing the "fitness" proxies, as well as the input data and the fitted model.

See Also

See the accompanying Vignette for an example of how to use and interpret SummarizeGrowth. https://CRAN.R-project.org/package=growthcurver/vignettes/Growthcurver-vignette.html.

See also gcfit.

Examples

# We can check that the parameters that are found are the same
# as we use to generate fake experimental data. To do so, let's first
# generate the "experimental" data using the logistic equation,
# e.g., absorbance readings from a single well in a plate reader over time.

k_in <- 0.5  # the initial carrying capacity
m0_in <- 1e-5  # the initial absorbance reading
r_in <- 1.2  # the initial growth rate
N <- 50  # the number of "measurements" collected during the growth
# curve experiment
data_t <- 0:N * 24 / N  # the times the measurements were made (in hours)
data_n <- NATI(k = k_in, n0 = n0_in, r = r_in, t = data_t)  # the measurements

# Now summarize the "experimental" growth data that we just generated
gc <- SummarizeGrowth(data_t, data_n)

# Get the possible metrics for fitness proxies
gc$vals$r  # growth rate is a common choice for fitness
gc$vals$t_gen  # doubling time, or generation time, is also common
gc$vals$k
gc$vals$n0
gc$vals$auc_l
gc$vals$auc_e
gc$vals$t_mid

# Compare the data with the fit visually by plotting it
plot(gc)

---

**SummarizeGrowthByPlate**

**Summarize Growth Curves**

**Description**

This function finds the parameters that describe the input data’s growth for a plate of growth curves. It does so by fitting the logistic curve to your growth curve measurements.

**Usage**

```r
SummarizeGrowthByPlate(
  plate,
  t_trim = 0,
  bg_correct = "min",
  plot_fit = FALSE,
  plot_file = "growthcurver.pdf"
)
```

**Arguments**

- `plate` A data.table with at least two columns. One column contains timepoints that measurements were taken (e.g., hours) and must be named "time". An optional column can be included called "blank" that contains the blank readings for background correction (make sure to select the "blank" bg_correct option if you provide a blank column). Each remaining column contains the absorbance readings from a single well in a plate.
t_trim Measurements taken after this time should not be included in fitting the curve. If stationary phase is variable, this may give you a better fit. A value of 0 means no trimming. Defaults to no trimming (0).

bg_correct The background correction method to use. No background correction is performed for "none". Specifying "min" subtracts the smallest value in a column from all the rows in that column, and specifying "blank" subtracts the values from the blank vector from the data_n vector.

plot_fit TRUE if you want to generate a pdf file that plots all columns provided in the plate along with the growthcurver’s fit. The default value is FALSE, which generates no plots.

plot_file The name of the file to save the plots to if you set plot_fit to TRUE. The default file is called "growthcurver.pdf".

Details

The logistic curve equation is

\[ N_t = \frac{N_0 K}{N_0 + (K - N_0)e^{-rt}} \]

where \( N_t \) is the number of cells (or the absorbance reading) at time \( t \), \( N_0 \) is the initial cell count (or absorbance reading), \( K \) is the carrying capacity, and \( r \) is the growth rate.

The fitness proxies returned are the parameters of the logistic equation and the area under the curve (a measure that integrates the effects of \( N_0 \), \( K \), and \( r \)). See \texttt{gcfit} for more documentation on these.

This method expects that your data adhere to a particular format.

- The data are provided in a data.frame
- One column in the data.frame is named "time" and contains the time measurements (e.g., hours).
- Each remaining column contains the readings from a single well in a plate reader. The name of the column will be used to identify the sample in the output data.
- There are no missing values or non-numeric data in the data.frame.

Value

A data.table containing the summary metrics and residual error from the fit of the logistic curve to the data. The names of the input columns are used to identify each well (or sample).

See Also

See the accompanying Vignette for an example of how to use and interpret SummarizeGrowthByPlate. \url{https://CRAN.R-project.org/package=growthcurver/vignettes/Growthcurver-vignette.html}

Examples

```r
#Get the summary metrics for the entire plate of sample data provided #with the Growthcurver package
```
#First, load the example data provided with Growthcurver. Note that there is
#a column named "time" -- this is necessary for Growthcurver to know which
#column contains the time measurements. In this dataset, the repeated
#measurements from a single well in a plate are given in a column of data.

myPlate <- growthdata
names(myPlate)

#Next, do the analysis for all the columns.
summary_plate <- SummarizeGrowthByPlate(plate = myPlate)

#The output is a data frame that contains the information on the best
#fit for each column of data.
head(summary_plate)  # Use head to display just the first few rows
Index

* datasets
  growthdata, 4

gcfit, 2, 8, 10
gcvals, 2, 3, 3
growthdata, 4

NAT, 7

SummarizeGrowth, 2, 3, 7
SummarizeGrowthByPlate, 9