# Package 'tame'

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**Title** Timing, Anatomical, Therapeutic and Chemical Based Medication Clustering

Version 0.2.0

**Description** Agglomerative hierarchical clustering with a bespoke distance measure based on medication similarities in the Anatomical Therapeutic Chemical Classification System, medication timing and medication amount or dosage. Tools for summarizing, illustrating and manipulating the cluster objects are also available.

```
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```

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cluster\_frequency

The Frequency of Assignment to Each Cluster

## Description

The function cluster\_frequency() calculates the number and frequency of individuals assigned to each cluster.

```
cluster_frequency(
  object,
  only = NULL,
  clusters = NULL,
  additional_data = NULL,
  ...
)
```

cluster\_frequency 3

## Arguments

object An object for which a summary is desired.

only <a href="data-masking">data-masking</a> Expressions that return a logical value, and are defined in

terms of the variables in object and/or additional\_data.

The default NULL selects all clusterings in object.

clusters <tidy-select> An unquoted expression naming the cluster or clusters in object

one wants to see summaries of. Names can be used as if they were positions in the data frame, so expressions like I:IV can be used to select a range of clusters.

The default NULL selects all clusters in the chosen clusterings of object.

additional\_data

A data frame with additional data that may be (left-)joined onto the parameters in object. This is often used in conjuction with only to select specific cluster-

ings based on additional\_data.

... Additional arguments passed to the specific summary sub-function.

#### **Details**

cluster\_frequency() calculates the number of individuals assigned to each cluster and the associated frequency of assignment.

#### Value

cluster\_frequency() returns a data frame with class cluster\_frequency.

- Clustering the name of the clustering.
- Cluster the cluster name.
- Count the number of individuals assigned to the cluster.
- Percent the percent of individuals assigned to the cluster.

### **Examples**

```
clust <- medic(complications, id = id, atc = atc, k = 3:5)
# make frequency tables
cluster_frequency(clust, k == 5)
cluster_frequency(clust, k < 5, I:III)</pre>
```

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comedication\_count

Frequency tables for medication amount

### **Description**

The function comedication\_count() calculates the number of unique medications for each individual and presents the count frequencies by cluster.

### Usage

```
comedication_count(
  object,
  only = NULL,
  clusters = NULL,
  count_grouper = function(x) {
     cut(x, breaks = c(0, 1, 2, Inf), labels = c("1",
     "2", "3+"))
},
  additional_data = NULL,
  ...
)
```

## **Arguments**

object An object for which a summary is desired.

only <data-masking> Expressions that return a logical value, and are defined in

terms of the variables in object and/or additional\_data.

The default NULL selects all clusterings in object.

clusters <tidy-select> An unquoted expression naming the cluster or clusters in object

one wants to see summaries of. Names can be used as if they were positions in the data frame, so expressions like I:IV can be used to select a range of clusters.

The default NULL selects all clusters in the chosen clusterings of object.

count\_grouper A function for grouping counts. As a standard it groups counts as 1 medication,

2 medications, and 3+ medications.

additional\_data

A data frame with additional data that may be (left-)joined onto the parameters in object. This is often used in conjuction with only to select specific cluster-

ings based on additional\_data.

... Additional arguments passed to the specific summary sub-function.

### **Details**

comedication\_count() calculates the number of ATC codes an individual has, and then outputs the number of individuals within a cluster that has that many ATC codes. Moreover, various relevant percentages or calculated. See Value below for more details on these percentages.

complications 5

#### Value

comedication\_count() returns a data frame of class comedication\_count

- Clustering the name of the clustering.
- Cluster the name of the cluster.
- Medication Count a number of medications. The numbers or groups are given by the count\_grouper() function.
- Number of People the number of individuals in cluster who has Medication Count number of comedications in study.
- Number of medications the number of medications of individuals who has Medication Count number of comedications in the cluster.
- Percentage of All People the percentage of individuals is study who has Medication Count number of comedications in the cluster.
- Percentage of People in Cluster the percentage of individuals in the cluster who has Medication Count number of comedications.
- Percentage of All Medications the percentage of medication in study from individuals who has Medication Count number of comedications in cluster.
- Percentage of Medication in Cluster the percentage of medication in cluster from individuals who has Medication Count number of comedications.
- Percentage of People with the Same Medication Count percentage of individuals among those with Medication Count number of comedications in this cluster.
- Percentage of Medication with the Same Medication Count percentage of medication among medication of individuals with Medication Count number of comedications in this cluster.

## **Examples**

```
clust <- medic(complications, id = id, atc = atc, k = 3:5)
comedication_count(clust, k == 5, clusters = I:III)</pre>
```

complications

A Simulated Data Set About Pregnancy Complications

### Description

We use this data set in all the examples in the package.

#### Usage

complications

#### **Format**

An object of class data. frame with 149 rows and 8 columns.

6 eczema

 $default\_atc\_groups$ 

Default ATC groups for summaries

## Description

This function finds the default ATC groups for the summaries. It is used in the summary.medic function.

## Usage

```
default_atc_groups(object, min_n = 2)
```

## Arguments

object

A medic object.

min\_n

The minimum number of ATC groups to be found.

#### Value

A data frame with two columns: regex and atc\_groups.

eczema

A Simulated Data Set About Eczema

## **Description**

A Simulated Data Set About Eczema

## Usage

eczema

## **Format**

An object of class data. frame with 50644 rows and 7 columns.

employ 7

employ

Employ a Clustering to New Data

#### **Description**

Employ a clustering to new data

#### Usage

```
employ(
  object,
  new_data,
  only = NULL,
  additional_data = NULL,
  assignment_method = "nearest_cluster",
  parallel = FALSE,
  ...
)
```

#### **Arguments**

object A medic clustering object for which employment is desired.

new\_data A data frame in which to look for variables with

only <data-masking> Expressions that return a logical value, and are defined in

terms of the variables in object and/or additional\_data and specifies which

clusterings should be employed to the new data.

additional\_data

A data frame with additional data that may be (left-)joined onto the parameters in object. This is often used in conjuction with only to select specific cluster-

ings based on additional\_data.

assignment\_method

A character naming the employment method. The default assignment method "nearest\_cluster" matches people in new\_data to their nearest cluster in the chosen clusterings from object. As finding exact matches (the next assignment method) is contained within this strategy the "exact\_only" matches are also reported in additional columns in the output.

The assignment method "exact\_only" only matches a person from new\_data to a cluster if they are a perfect match to anyone in object. Thus, people from new\_data are not guaranteed assignment to a cluster.

parallel

A logical or an integer. If FALSE, the default, no parallelization is done.

If TRUE or an integer larger than 2L parallelization is implemented via parLapply from the **parallel** package. When parallel is TRUE the number of clusters is set to detectCores - 1, and when parallel is an integer then the number of clusters is set to parallel. For more details on the parallelization method see parallel::parLapply.

paraliel::parLappiy

. . Additional arguments affecting the employment procedure.

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#### Value

employ returns a medic object.

### **Examples**

```
part1 <- complications[1:100,]
part2 <- complications[101:149,]

clust <- medic(part1, id = id, atc = atc, k = 3)

# Nearest cluster matching
employ(clust, part2)

# Only exact matching
employ(clust, part2, assignment_method = "exact_only")</pre>
```

enrich

Enrich Clustering Parameter

### **Description**

Enrich the parameter information in a clustering with user-defined data.

#### Usage

```
enrich(object, additional_data = NULL, by = NULL)
```

#### **Arguments**

object A medic object for enrichment. additional\_data

A data frame with additional data that may be (left-)joined onto the parameters in object.

by

A character vector of variables to join by. This variables is passed to the by term in a dplyr::left\_join() and inherits its behavior:

If NULL, the default, the join will perform a natural join, using all variables in common across the parameters and additional\_data.

To join by different variables on parameters and additional\_data, use a named vector. For example, by =  $c("k" = "cluster_size")$  will match parameters\$k to additional\_data\$cluster\_size.

To join by multiple variables, use a vector with length > 1. For example, by =  $c("k", "summation_method")$  will match parametersk to additional\_datak and parameters $summation_method$  to

additional\_data\$summation\_method. Use a named vector to match different variables in parameters and additional\_data.

For example, by =  $c("k" = "cluster\_size", "summation\_method" = "sm")$  will match parameters\$k to additional\_data\$cluster\_size and parameters\$summation\_method to additional\_data\$sm.

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## **Details**

The enrich() function is a joining function used for enriching the clustering characteristics with user-defined data. This function is used in all of the investigative functions with a additional\_data statement such as summary(), cluster\_frequency() and medication\_frequency().

#### Value

An object of class medic.

## **Examples**

```
clust <- medic(
   complications,
   id = id,
   atc = atc,
   timing = first_trimester:third_trimester,
   k = 3:5
)

new_parameters <- data.frame(k = 3:5, size = c("small", "small", "large"))
enrich(clust, new_parameters)</pre>
```

is.medic

Test if an object is a medic-object

## Description

Test if an object is a medic-object

## Usage

```
is.medic(object)
```

## Arguments

object

Any object.

### Value

TRUE is the object inherits from the medic class and has the required elements.

## **Examples**

```
clust <- medic(complications, id = id, atc = atc, k = 3)
is.medic(clust)</pre>
```

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medic

Medication clustering (based on ATC and timing)

## **Description**

The medic method uses agglomerative hierarchical clustering with a bespoke distance measure based on medication ATC codes similarities, medication timing and medication amount or dosage.

#### Usage

```
medic(
  data,
 k = 5,
  id,
  atc,
  timing,
  base_clustering,
  linkage = "complete",
  summation_method = "sum_of_minima",
  alpha = 1,
  beta = 1,
  gamma = 1,
  p = 1,
  theta = (5:0)/5,
  parallel = FALSE,
  return_distance_matrix = FALSE,
  set_seed = FALSE,
)
## S3 method for class 'medic'
print(x, ...)
```

## **Arguments**

data	A data frame containing all the variables for the clustering.
k	a vector specifying the number of clusters to identify.
id	<pre><tidy-select> An unquoted expression naming the variable in data describ- ing person id.</tidy-select></pre>
atc	<tidy-select> An unquoted expression naming the variable in data containing ATC codes.</tidy-select>
timing	<tidy-select> An unquoted expression naming the variable or variables in data describing medication timing. Variable names can be used as if they were positions in the data frame, so expressions like x:y can be used to select a range of variables. Moreover, pattern matching selection helpers such as starts_with or num_range may also be used to select timing variables.</tidy-select>

medic 11

base\_clustering

<tidy-select> An unquoted expression naming the variable in data that gives

an initial clustering to start the medic from or NULL.

linkage The agglomeration method to be used in the clustering. This should be (an un-

ambiguous abbreviation of) one of "ward.D", "ward.D2", "single", "complete", "average" (= UPGMA), "mcquitty" (= WPGMA), "median" (= WPGMC) or "centroid" (= UPGMC). See stats::hclust for more information. For a discussion

of linkage criterion choice see details below.

summation\_method

The summation method used in the distance measure. This should be either

"double\_sum" or "sum\_of\_minima". See details below for more information.

alpha A number giving the tuning of the normalization. See *details* below for more

information.

A number giving the power of the individual medication combinations. See

details below for more information.

gamma A number giving the weight of the timing terms. See *details* below for more

information.

p The power of the Minkowski distance used in the timing-specific distance. See

details below for more information.

theta A vector of length 6 specifying the tuning of the ATC measure. See *details* 

below for more information.

parallel A logical or an integer. If FALSE, the default, no parallelization is done.

If TRUE or an integer larger than 2L parallelization is implemented via parLapply from the **parallel** package. When parallel is TRUE the number of clusters is set to detectCores - 1, and when parallel is an integer then the number of clusters is set to parallel. For more details on the parallelization method see

parallel::parLapply.

return\_distance\_matrix

A logical.

set\_seed A logical or an integer.

... Additional arguments not currently in use.

x A medic object for printing.

## Details

The medic method uses agglomerative hierarchical clustering with a bespoke distance measure based on medication ATC codes and timing similarities to assign medication pattern clusters to people.

Two versions of the distance measure are available:

The double sum:

$$d(p_i, p_j) = N_{\alpha}(M_i \times M_j) \sum_{m \in M_i} \sum_{n \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta}.$$

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and the sum of minima:

$$d(p_i, p_j) = \frac{1}{2} (N_{\alpha}(M_i) \sum_{m \in M_i} \min_{n \in M_j} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_j} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} \min_{m \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} ((1 + D_{\theta}(m, n))(1 + \gamma T_p(t_{im}, t_{jn})) - 1)^{\beta} + N_{\alpha}(M_j) \sum_{n \in M_i} ((1 + D_{\theta}(m, n))(1 +$$

#### **Normalization:**

$$N_{\alpha}(x) = |x|^{-\alpha}$$

If the normalization tuning, alpha, is 0, then no normalization is preformed and the distance measure becomes highly dependent on the number of distinct medications given. That is, people using more medication will have larger distances to others. If the normalization tuning, alpha, is 1 - the default - then the summation is normalized with the number of terms in the sum, in other words, the average is calculated.

#### **ATC distance:**

The central idea of this method, namely the ATC distance, is given as

$$D_{\theta}(x,y) = \sum_{i=1,\dots,5} 1\{xandy match on level i, but not level i+1\}\theta_i$$

The ATC distance is tuned using the vector theta.

Note that two ATC codes are said to match at level i when they are identical at level i. E.g. the two codes N06AB01 and N06AA01 match on level 1, 2, and 3 as they are both "N" at level 1, "N06" at level 2, and "N06A" at level 3, but at level 4 they differ ("N06AB" and "N06AA" are not the same).

#### **Timing distance:**

The timing distance is a simple Minkowski distance:

$$T(x,y) = (\sum_{t \in T} |x_t - y_t|^p)^{1/p}.$$

When p is 1, the default, the Manhattan distance is used.

## Value

An object of class *medic* which describes the clusters produced the hierarchical clustering process. The object is a list with components:

data the inputted data frame data with the cluster assignments appended at the end.

clustering a data frame with the person id as given by id, the .analysis\_order and the clusters found.

variables a list of the variables used in the clustering.

**parameters** a data frame with all the inputted clustering parameters and the corresponding method names. These method names correspond to the column names for each cluster in the clustering data frame described right above.

key a list of keys used internally in the function to keep track of simplified versions of the data.

**distance\_matrix** the distance matrices for each method if return\_distance\_matrix is TRUE otherwise NULL.

call the matched call.

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### Methods (by generic)

• print(medic): Print method for medic-objects

#### See Also

```
summary.medic for summaries and plots.

employ for employing an existing clustering to new data.

enrich for enriching the meta data in the medic object with additional data.
```

## **Examples**

```
# A simple clustering based only on ATC
clust <- medic(complications, id = id, atc = atc, k = 3)
# A simple clustering with both ATC and timing
clust <- medic(
   complications,
   id = id,
   atc = atc,
   timing = first_trimester:third_trimester,
   k = 3
)</pre>
```

medication\_frequency ATC Code Frequency Within Clusters

## **Description**

The function medications() calculates the frequency of the different unique ATC codes within each cluster.

```
medication_frequency(
  object,
  only = NULL,
  clusters = NULL,
  additional_data = NULL,
  ...
)
```

#### **Arguments**

object An object for which a summary is desired.

only <data-masking> Expressions that return a logical value, and are defined in

terms of the variables in object and/or additional\_data.

The default NULL selects all clusterings in object.

clusters <tidy-select> An unquoted expression naming the cluster or clusters in object

one wants to see summaries of. Names can be used as if they were positions in the data frame, so expressions like I:IV can be used to select a range of clusters.

The default NULL selects all clusters in the chosen clusterings of object.

additional\_data

A data frame with additional data that may be (left-)joined onto the parameters in object. This is often used in conjuction with only to select specific cluster-

ings based on additional\_data.

... Additional arguments passed to the specific summary sub-function.

#### **Details**

medication\_frequency() calculates the number of individuals with a specific ATC code within a cluster. Moreover, it calculates the percentage of people with this medication assigned to this cluster and the percent of people within the cluster with this medication.

#### Value

medication\_frequency() returns a data frame with class medication\_frequency.

- Clustering the name of the clustering.
- Cluster the cluster name.
- atc ATC codes.
- Count number of individuals with this ATC code in this cluster.
- Percent of All Medication the percentage of individuals in the study with this ATC code and cluster.
- Percent of Medication in Cluster the percent of individuals in the cluster with this ATC code.

## **Examples**

```
clust <- medic(complications, id = id, atc = atc, k = 3:5)
medication_frequency(clust, k == 5, clusters = I:III)</pre>
```

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```
parameters_constructor
```

Internal option constructor

## **Description**

Given the input of the medic this function checks the input and constructs a data frame with the analysis parameters specified by the user.

## Usage

```
parameters_constructor(
  data,
  id,
  k = 5,
  atc,
  timing,
  base_clustering,
  linkage = "complete",
  summation_method = "sum_of_minima",
  alpha = 1,
  beta = 1,
  gamma = 1,
  p = 1,
  theta = (5:0)/5,
  ...
)
```

## Arguments

data	A data frame containing all the variables for the clustering.
id	<tidy-select> An unquoted expression naming the variable in data describing person id.</tidy-select>
k	a vector specifying the number of clusters to identify.
atc	<tidy-select> An unquoted expression naming the variable in data containing ATC codes.</tidy-select>
timing	<tidy-select> An unquoted expression naming the variable or variables in data describing medication timing. Variable names can be used as if they were positions in the data frame, so expressions like x:y can be used to select a range of variables. Moreover, pattern matching selection helpers such as starts_with or num_range may also be used to select timing variables.</tidy-select>

base\_clustering

<tidy-select> An unquoted expression naming the variable in data that gives an initial clustering to start the medic from or NULL.

linkage	The agglomeration method to be used in the clustering. This should be (an unambiguous abbreviation of) one of "ward.D", "ward.D2", "single", "complete", "average" (= UPGMA), "mcquitty" (= WPGMA), "median" (= WPGMC) or "centroid" (= UPGMC). See stats::hclust for more information. For a discussion of linkage criterion choice see <i>details</i> below.	
summation_method		
	The summation method used in the distance measure. This should be either "double_sum" or "sum_of_minima". See <i>details</i> below for more information.	
alpha	A number giving the tuning of the normalization. See <i>details</i> below for more information.	
beta	A number giving the power of the individual medication combinations. See <i>details</i> below for more information.	
gamma	A number giving the weight of the timing terms. See <i>details</i> below for more information.	
p	The power of the Minkowski distance used in the timing-specific distance. See <i>details</i> below for more information.	
theta	A vector of length 6 specifying the tuning of the ATC measure. See <i>details</i> below for more information.	
	Additional arguments not currently in use.	

## Value

A data.frame with the parameters for clustering.

## **Examples**

```
parameters_constructor(
  data = complications,
  k = 3,
  id = id,
  atc = atc
)
```

```
plot_cluster_frequency
```

Plot Cluster Frequency

## Description

This function plots the cluster frequency.

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### **Usage**

```
plot_cluster_frequency(object, ...)

## S3 method for class 'medic'
plot_cluster_frequency(object, ...)

## S3 method for class 'summary.medic'
plot_cluster_frequency(object, ...)

## S3 method for class 'cluster_frequency'
plot_cluster_frequency(object, scale = "percent", with_population = FALSE, ...)
```

#### **Arguments**

object The object containing the cluster frequency data.
... Additional arguments passed to the plotting functions.
scale The scale of the y-axis. Must be either "percent" or "count".
with\_population

Logical value indicating whether to include the population cluster.

#### Value

A ggplot object.

### See Also

```
cluster_frequency
plot_medication_frequency
plot_comedication_count
plot_timing_trajectory
plot_timing_atc_group
plot_summary
```

## **Examples**

```
clust <- medic(complications, id = id, atc = atc, k = 3)
clust |> plot_cluster_frequency()
clust |> cluster_frequency() |> plot_cluster_frequency()
clust |> summary() |> plot_cluster_frequency()
```

```
plot_comedication_count
```

Plot Comedication Count

## Description

This function plots the comedication count.

## Usage

```
plot_comedication_count(object, ...)
## S3 method for class 'medic'
plot_comedication_count(object, ...)
## S3 method for class 'summary.medic'
plot_comedication_count(object, ...)
## S3 method for class 'comedication_count'
plot_comedication_count(
   object,
   scale = "percent",
   scope = "cluster",
   focus = "people",
   with_population = FALSE,
   ...
)
```

## **Arguments**

object	The object containing the comedication count data.	
	Additional arguments passed to the plotting functions.	
scale	The scale of the y-axis. Must be either "percent" or "count".	
scope	The scope of the plot. Must be one of "cluster", "global" or "medication count".	
focus	The focus of the plot. Must be either "people" or "medication".	
with_population		
	Logical value indicating whether to include the population cluster.	

## Value

A ggplot object.

#### See Also

```
comedication_count
plot_cluster_frequency
plot_medication_frequency
plot_timing_trajectory
plot_timing_atc_group
plot_summary
```

### **Examples**

```
clust <- medic(complications, id = id, atc = atc, k = 3)
clust |> plot_comedication_count()
clust |> comedication_count() |> plot_comedication_count()
clust |> summary() |> plot_comedication_count()
```

```
\verb"plot_medication_frequency"
```

Plot Medication Frequency

## Description

This function plots the medication frequency.

```
plot_medication_frequency(object, ...)
## S3 method for class 'medic'
plot_medication_frequency(object, ...)
## S3 method for class 'summary.medic'
plot_medication_frequency(object, ...)
## S3 method for class 'medication_frequency'
plot_medication_frequency(
   object,
   scale = "percent",
   scope = "cluster",
   with_population = FALSE,
   ...
)
```

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## **Arguments**

object The object containing the medication frequency data.

... Additional arguments passed to the plotting functions.

scale The scale of the y-axis. Must be either "percent" or "count".

scope The scope of the plot. Must be one of "cluster", "global" or "medication".

with\_population

Logical value indicating whether to include the population cluster.

#### Value

A ggplot object.

#### See Also

```
medication_frequency
plot_cluster_frequency
plot_comedication_count
plot_timing_trajectory
plot_timing_atc_group
plot_summary
```

### **Examples**

```
clust <- medic(complications, id = id, atc = atc, k = 3)

clust |> plot_medication_frequency()
clust |> medication_frequency() |> plot_medication_frequency()
clust |> summary() |> plot_medication_frequency()
```

plot\_summary

Plot Summary

### **Description**

This function plots the summary of the clustering results.

```
plot_summary(object, ...)
## S3 method for class 'medic'
plot_summary(object, only = NULL, clusters = NULL, additional_data = NULL, ...)
## S3 method for class 'summary.medic'
```

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```
plot_summary(
  object,
  n_breaks = 5,
  plot_individual = FALSE,
  labels = FALSE,
  alpha_individual = 0.1,
  label_y_value = 0.1,
  ...
)
```

#### **Arguments**

object The object containing the summary data.

... Additional arguments passed to the plotting functions.

only <data-masking> Expressions that return a logical value, and are defined in

terms of the variables in object and/or additional\_data.

The default NULL selects all clusterings in object.

clusters <tidy-select> An unquoted expression naming the cluster or clusters in object

one wants to see summaries of. Names can be used as if they were positions in the data frame, so expressions like I:IV can be used to select a range of clusters.

The default NULL selects all clusters in the chosen clusterings of object.

additional\_data

A data frame with additional data that may be (left-)joined onto the parameters in object. This is often used in conjuction with only to select specific cluster-

ings based on additional\_data.

n\_breaks The number of breaks for the time scale.

plot\_individual

Logical value indicating whether to plot individual trajectories.

labels Logical value indicating whether to include labels.

alpha\_individual

The alpha value for the individual trajectories.

label\_y\_value A number between 0 and 1 that defines the height of the label text hight.

#### Value

A ggplot object.

### See Also

```
summary
plot_cluster_frequency
plot_medication_frequency
plot_comedication_count
plot_timing_trajectory
plot_timing_atc_group
```

### **Examples**

```
clust <- medic(</pre>
  complications,
  id = id,
 atc = atc,
  k = 3,
  timing = first_trimester:third_trimester
clust |> plot_summary()
clust |> summary() |> plot_summary()
# If the clustering object contains more than one clustering, it is necessary
# to filter the clustering, as only one clustering can be plotted at a time.
clust <- medic(</pre>
  complications,
  id = id,
 atc = atc,
  k = 3:5,
  timing = first_trimester:third_trimester
clust |> plot_summary(only = k == 4)
clust |> summary(only = k == 4) |> plot_summary()
```

plot\_timing\_atc\_group Plot Timing ATC Group

## Description

This function plots the timing ATC group.

```
plot_timing_atc_group(object, ...)
## S3 method for class 'medic'
plot_timing_atc_group(object, ...)
## S3 method for class 'summary.medic'
plot_timing_atc_group(object, ...)
## S3 method for class 'timing_atc_group'
plot_timing_atc_group(
   object,
   focus = "average",
   with_population = FALSE,
   max_lines = 50,
   ...
)
```

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### **Arguments**

object The object containing the timing ATC group data.
... Additional arguments passed to the plotting functions.

focus The focus of the plot. Must be either "average", "individual" or "both".

with\_population

Logical value indicating whether to include the population cluster.

max\_lines The maximum number of lines to plot.

#### Value

A ggplot object.

#### See Also

```
timing_atc_group
plot_cluster_frequency
plot_medication_frequency
plot_comedication_count
plot_timing_trajectory
plot_summary
```

## **Examples**

```
clust <- medic(
  complications,
  id = id,
  atc = atc,
  k = 3:5,
  timing = first_trimester:third_trimester
)

clust |> plot_timing_atc_group()
  clust |> timing_atc_group() |> plot_timing_atc_group()
  clust |> summary() |> plot_timing_atc_group()
```

```
plot_timing_trajectory
```

Plot Timing Trajectory

## Description

This function plots the timing trajectory.

#### Usage

```
plot_timing_trajectory(object, ...)

## S3 method for class 'medic'
plot_timing_trajectory(object, ...)

## S3 method for class 'summary.medic'
plot_timing_trajectory(object, ...)

## S3 method for class 'timing_trajectory'
plot_timing_trajectory(
   object,
   focus = "average",
   with_population = FALSE,
   max_lines = 50,
   ...
)
```

## **Arguments**

object The object containing the timing trajectory data.

... Additional arguments passed to the plotting functions.

focus The focus of the plot. Must be either "average", "individual" or "both".

with\_population

Logical value indicating whether to include the population cluster.

max\_lines The maximum number of lines to plot.

#### Value

A ggplot object.

#### See Also

```
timing_trajectory
plot_cluster_frequency
plot_medication_frequency
plot_comedication_count
plot_timing_atc_group
plot_summary
```

## **Examples**

```
clust <- medic(
  complications,
  id = id,
  atc = atc,</pre>
```

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```
k = 3:5,
  timing = first_trimester:third_trimester
)

clust |> plot_timing_trajectory()
  clust |> timing_trajectory() |> plot_timing_trajectory()
  clust |> summary() |> plot_timing_trajectory()
```

print.summary.medic

Print Summary of Medication

#### **Description**

This function prints a summary of medication information.

#### Usage

```
## S3 method for class 'summary.medic'
print(x, ...)
```

### **Arguments**

x An object of class summary.medic.

... currently only included for compatibility with generic. Has no effect.

## **Details**

This function prints various information about medication, including cluster frequency, medication frequency, number of different medication taken in the study period, average exposure trajectories, and average exposure trajectories by ATC groups.

#### Value

The function is called for its side effects and does not return any value.

refactor

Refactor Cluster Levels

#### **Description**

Refactor the levels of the chosen clusters.

```
refactor(object, ..., inheret_parameters = TRUE)
```

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## Arguments

. . .

object A medic object.

<data-masking> Name-value pairs. ... is passed to dplyr::mutate, and
therefor inherits its behavior:

The name gives the name of the new clustering in the output. The value can be:

- A vector of length 1, which will be recycled to the correct length.
- A function of another clustering.

When a recording uses the name of an existing clustering, this new clustering will overwrite the existing one.

inheret\_parameters

A logical. If TRUE a new clustering overwriting an existing clustering inherits the parameters of the old.

#### Value

A medic object with relevant clusterings refactored.

### **Examples**

```
clust <- medic(complications, id = id, atc = atc, k = 3:4)

# Refactor one clustering
refactor(
   clust,
   `cluster_1_k=4` = dplyr::recode(`cluster_1_k=4`, IV = "III")
)

# Refactor all clusterings
refactor(
   clust,
   dplyr::across(
    dplyr::everything(),
    ~dplyr::recode(., IV = "III")
)
)</pre>
```

str.summary.medic

Summary of a medic-object using str function

### Description

Summary of a medic-object using str function

```
## S3 method for class 'summary.medic'
str(object, ...)
```

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#### **Arguments**

object A medic object.

... Additional arguments passed to str.default.

This function provides a summary of an object by using the str function. It is a modified version of the str.default function from the utils package, with

the maximum level set to 2.

summary.medic

Summary of medic object

## Description

Make cluster characterizing summaries.

### Usage

```
## $3 method for class 'medic'
summary(
  object,
  only = NULL,
  clusters = NULL,
  outputs = "all",
  additional_data = NULL,
  ...
)
```

#### **Arguments**

object An object for which a summary is desired.

only <data-masking> Expressions that return a logical value, and are defined in

terms of the variables in object and/or additional\_data.

The default NULL selects all clusterings in object.

clusters <tidy-select> An unquoted expression naming the cluster or clusters in object

one wants to see summaries of. Names can be used as if they were positions in the data frame, so expressions like I:IV can be used to select a range of clusters.

The default NULL selects all clusters in the chosen clusterings of object.

outputs A character vector naming the desired characteristics to output. The default

names all possible output types.

additional\_data

A data frame with additional data that may be (left-)joined onto the parameters in object. This is often used in conjuction with only to select specific cluster-

ings based on additional\_data.

.. Additional arguments passed to the specific summary sub-function.

#### Value

A list of clustering characteristics of class summary.medic is returned. It can contain any of the following characteristics:

#### **Cluster Frequencies:**

The number of individuals assigned to each cluster and the associated frequency of assignment.

### **Medication Frequencies:**

The number of individuals with a specific ATC code within a cluster. Moreover, it calculates the percentage of people with this medication assigned to this cluster and the percent of people within the cluster with this medication.

#### **Comedication Count:**

The number of ATC codes an individual has, and then outputs the number of individuals within a cluster that has that many ATC codes. Moreover, various relevant percentages or calculated. See Value below for more details on these percentages.

## **Timing Trajectories:**

The number of unique timing trajectories in each cluster, and the average timing trajectories in each cluster.

## Timing and ATC group interactions:

The number of people with unique timing trajectory and ATC group, as given by atc\_groups, in each cluster.

## **Examples**

```
clust <- medic(
  complications,
  id = id,
  atc = atc,
  k = 3:5,
   timing = first_trimester:third_trimester)
summary(clust)</pre>
```

summary\_crop

Crop Clustering Summary

#### **Description**

Functions for cropping summarized cluster data.

### Usage

```
summary_crop(object, ...)
## S3 method for class 'cluster_frequency'
summary_crop(object, top_n = 5L, min_count = 0, min_percent = 0, ...)
## S3 method for class 'medication_frequency'
summary_crop(
 object,
  top_n = 5L
 min_count = 0,
 min_percent = 0,
 scope = "cluster",
)
## S3 method for class 'comedication_count'
summary_crop(object, ...)
## S3 method for class 'timing_trajectory'
summary_crop(object, sample_n_individual = 100L, weighted_sample = TRUE, ...)
## S3 method for class 'timing_atc_group'
summary_crop(
 object,
 sample_n_individual = 100L,
 weighted_sample = TRUE,
 min_count = 0L,
  . . .
)
## S3 method for class 'summary.medic'
summary_crop(object, which = "all", ...)
```

### **Arguments**

object	The summary object to be cropped.
	Additional arguments to be passed to the specific method.
top_n	integer. In the case of cluster_frequency it is the number of clusters to keep. In the case of medication_frequency it is the number of medications to keep. If inf, all clusters or medications are kept.
min_count	integer. The minimum count of a cluster or medication to keep it in the summary. If 0, the default, the minimum count is zero, i.e. there is not a minimum count.
min_percent	numeric. The minimum percentage of a cluster or medication to keep it in the summary. If 0, the default, the minimum percentage is zero, i.e. there is not a minimum percentage.

scope

character. The scope of the summary crops top\_n, min\_count and min\_percent. The options are "cluster" and "global". The default is "cluster". If "cluster", the crop is based on the percentage of medication in the cluster. If "global", the crop is based on the percentage of all medication.

sample\_n\_individual

a logical or integer. If FALSE, no individual timing trajectories are sampled. If integer, sample\_n\_individual is the number of individual timing trajectories to sample. To sample all individual timing trajectories, set sample\_n\_individual to Inf.

weighted\_sample

a logical, but only used if sample\_n\_individual is an integer. If TRUE, the individual timing trajectories are sampled weighted by the number of medications in the individual timing trajectory. If FALSE, the individual timing trajectories are sampled uniformly.

which

A character vector specifying which summaries to crop. The options are "cluster\_frequency", "medication\_frequency", "comedication\_count", "timing\_trajectory", and "timing\_atc\_group". The default is "all".

#### Value

A summary object, which is a modified version of the input summary object.

#### cluster\_frequency summary crop

Extracts the top top\_n clusters by count. If top\_n is Inf, all clusters are kept. If min\_count is greater than 0, clusters with a count less than min\_count are removed. If min\_percent is greater than 0, clusters with a percentage less than min\_percent are removed. The remaining clusters are grouped into a "Remaining" cluster.

#### medication\_frequency summary crop

Extracts the top top\_n medications by count. If top\_n is Inf, all medications are kept. If min\_count is greater than 0, medications with a count less than min\_count are removed. If min\_percent is greater than 0, medications with a percentage less than min\_percent are removed. The remaining medications are grouped into a "Remaining" cluster.

The scope argument determines the scope of the crop. If scope is "cluster", the crop is based on the percentage of medication in the cluster. If scope is "global", the crop is based on the percentage of all medication.

### comedication\_count summary crop

TO DO

#### timing\_trajectory summary crop

Samples sample\_n\_individual individual timing trajectories. If sample\_n\_individual is Inf, all individual timing trajectories are kept. If weighted\_sample is TRUE, the individual timing trajectories are sampled weighted by the number of medications in the individual timing trajectory.

#### timing\_atc\_group summary crop

Samples sample\_n\_individual individual timing trajectories. If sample\_n\_individual is Inf, all individual timing trajectories are kept. If weighted\_sample is TRUE, the individual timing trajectories are sampled weighted by the number of medications in the individual timing trajectory.

## summary.medic summary crop

Crops multiple summaries. The which argument is a character vector specifying which summaries to crop. The options are "cluster\_frequency", "medication\_frequency", "comedication\_count", "timing\_trajectory", and "timing\_atc\_group". If which is "all", all summaries are cropped.

The ... argument is passed to the specific methods, e.g. top\_n and min\_count are passed to cluster\_frequency and medication\_frequency.

#### See Also

summary, cluster\_frequency, medication\_frequency, comedication\_count, timing\_trajectory,
timing\_atc\_group

## **Examples**

```
clust <- medic(</pre>
  complications,
  id = id.
 atc = atc,
  k = 3:5,
  timing = first_trimester:third_trimester
# Crop the cluster frequency summary
clust |>
  cluster_frequency() |>
  summary_crop(top_n = 3)
clust |>
  summary() |>
  summary_crop(which = "cluster_frequency", top_n = 3)
# Crop the medication frequency summary
clust |>
  medication_frequency() |>
  summary\_crop(top\_n = 3)
clust |>
  summary() |>
  summary_crop(which = "medication_frequency", top_n = 3)
# Crop the co-medication count summary
clust |>
```

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```
comedication_count() |>
  summary_crop(min_count = 10)
clust |>
  summary() |>
  summary_crop(which = "comedication_count", min_count = 10)
# crop the timing trajectory summary
clust |>
  timing_trajectory() |>
  summary_crop()
clust |>
  summary() |>
  summary_crop(which = "timing_trajectory")
# crop the timing ATC group summary
clust |>
  timing_atc_group() |>
  summary_crop()
clust |>
  summary() |>
  summary_crop(which = "timing_atc_group")
# crop multiple summaries
clust |>
  summary() |>
  summary_crop(
   which = c("cluster_frequency", "medication_frequency"),
   top_n = 3
  )
```

timing\_atc\_group

Timing and ATC pattern interactions

### **Description**

The function timing\_atc\_group() calculates the frequencies of distinct timing and ATC combinations within clusters.

```
timing_atc_group(
  object,
  only = NULL,
  clusters = NULL,
```

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```
atc_groups = default_atc_groups,
  additional_data = NULL,
)
```

#### **Arguments**

object An object for which a summary is desired.

only <data-masking> Expressions that return a logical value, and are defined in

terms of the variables in object and/or additional\_data.

The default NULL selects all clusterings in object.

clusters <tidy-select> An unquoted expression naming the cluster or clusters in object

> one wants to see summaries of. Names can be used as if they were positions in the data frame, so expressions like I:IV can be used to select a range of clusters.

The default NULL selects all clusters in the chosen clusterings of object.

A data frame specifying the ATC groups to summaries by or a function that atc\_groups

returns such a data.frame. The data.frame must have two columns:

• regex giving regular expressions specifying the wanted ATC groups.

atc\_groups the name of this ATC grouping.

As a standard the anatomical level (first level) of the ATC codes is used.

additional\_data

A data frame with additional data that may be (left-)joined onto the parameters in object. This is often used in conjuction with only to select specific clusterings based on additional\_data.

Additional arguments passed to the specific summary sub-function.

#### **Details**

timing\_atc\_group() calculates both the number of people with unique timing trajectory and ATC group, as given by atc\_groups, in each cluster.

### Value

timing\_atc\_group() returns a list of class timing\_atc\_group with two data frames:

#### average:

- Clustering the name of the clustering.
- Cluster the name of the cluster.
- ATC Groups the name of the ATC group. The groups are given by the atc\_groups input.
- timing variables the average timing value in the ATC group and cluster.
- Number of Medications the number of medications in the ATC group in the cluster.
- Percentage of Medications the percentage of medication in the cluster with this ATC
- Number of Distinct Timing Trajectories the number of unique timing trajectories in the ATC group in the cluster.

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#### individual:

- Clustering the name of the clustering.
- Cluster the name of the cluster.
- timing variables a unique timing pattern in the ATC group and cluster.
- Number of Medications with Timing Trajectory the number of medications with this unique timing trajectory and ATC group.

#### **Examples**

```
clust <- medic(
  complications,
  id = id,
  atc = atc,
  k = 3:5,
  timing = first_trimester:third_trimester
)
timing_atc_group(clust, k == 5, clusters = I:III)</pre>
```

timing\_trajectory

Timing pattern frequency within clusters

## Description

timing\_trajectory() calculates the average timing paths within clusters.

## Usage

```
timing_trajectory(
  object,
  only = NULL,
  clusters = NULL,
  additional_data = NULL,
  ...
)
```

#### **Arguments**

object An object for which a summary is desired.

only <a href="masking">data-masking</a> Expressions that return a logical value, and are defined in

terms of the variables in object and/or additional\_data.

The default NULL selects all clusterings in object.

clusters <tidy-select> An unquoted expression naming the cluster or clusters in object

one wants to see summaries of. Names can be used as if they were positions in the data frame, so expressions like I:IV can be used to select a range of clusters.

The default NULL selects all clusters in the chosen clusterings of object.

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```
additional_data
```

A data frame with additional data that may be (left-)joined onto the parameters in object. This is often used in conjuction with only to select specific clusterings based on additional\_data.

.. Additional arguments passed to the specific summary sub-function.

#### **Details**

timing\_trajectory() calculates both the number of unique timing trajectories in each cluster and the average timing trajectories in each cluster.

#### Value

timing\_trajectory() returns a list of class timing\_trajectory with two data frames:

#### average:

- Clustering the name of the clustering.
- Cluster the cluster name.
- timing variables the average timing value in the cluster.
- Count the number of people in the cluster.

#### individual:

- Clustering the name of the clustering.
- Cluster the cluster name.
- timing variables unique timing pattern in the cluster.
- Count number of people with this unique timing pattern.

## **Examples**

```
clust <- medic(
  complications,
  id = id,
  atc = atc,
  k = 3:5,
  timing = first_trimester:third_trimester
)
timing_trajectory(clust, k == 5, clusters = I:III)</pre>
```

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