

# Package ‘immApex’

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**Title** Tools for Adaptive Immune Receptor Sequence-Based Keras3 Modeling

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**Description** A set of tools to build tensorflow/keras3-based models in R from amino acid and nucleotide sequences focusing on adaptive immune receptors. The package includes pre-processing of sequences, unifying gene nomenclature usage, encoding sequences, and combining models. This package will serve as the basis of future immune receptor sequence functions/packages/models compatible with the scRepertoire ecosystem.

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immApex-package	<i>immApex: Tools for Adaptive Immune Receptor Sequence-Based Keras3 Modeling</i>
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## Description

A set of tools to build tensorflow/keras3-based models in R from amino acid and nucleotide sequences focusing on adaptive immune receptors. The package includes pre-processing of sequences, unifying gene nomenclature usage, encoding sequences, and combining models. This package will serve as the basis of future immune receptor sequence functions/packages/models compatible with the scRepertoire ecosystem.

## Author(s)

**Maintainer:** Nick Borcharding <ncborch@gmail.com>

## See Also

Useful links:

- <https://github.com/ncborcharding/immApex/>
- Report bugs at <https://github.com/ncborcharding/immApex/issues>

---

adjacencyMatrix	<i>Adjacency matrix from amino acid or nucleotide sequences</i>
-----------------	---

---

**Description**

Calculate frequency of adjacency between residues along a set of biological sequences.

**Usage**

```
adjacencyMatrix(  
  input.sequences = NULL,  
  normalize = TRUE,  
  sequence.dictionary = amino.acids  
)
```

**Arguments**

input.sequences	The amino acid or nucleotide sequences to use
normalize	Return the values as a function of total number of residues ( <b>TRUE</b> ) or frequencies ( <b>FALSE</b> )
sequence.dictionary	The letters to use in sequence generation (default are all amino acids)

**Value**

Adjacency matrix based on input.sequences.

**Examples**

```
new.sequences <- generateSequences(prefix.motif = "CAS",  
  suffix.motif = "YF",  
  number.of.sequences = 100,  
  min.length = 8,  
  max.length = 16)  
  
adj.matrix <- adjacencyMatrix(new.sequences,  
  normalize = TRUE)
```

---

formatGenes	<i>Ensure clean gene nomenclature using IMGT annotations</i>
-------------	--

---

**Description**

This function will format the genes into a clean nomenclature using the IMGT conventions.

**Usage**

```
formatGenes(
  input.data,
  region = "v",
  technology = NULL,
  species = "human",
  simplify.format = TRUE
)
```

**Arguments**

input.data	Data frame of sequencing data or scRepertoire outputs
region	Sequence gene loci to access - "v", "d", "j", or "c" or a combination using c("v", "d", "j")
technology	The sequencing technology employed - <b>'TenX'</b> , <b>'Adaptive'</b> , or <b>'AIRR'</b>
species	One or two word designation of species. Currently supporting: "human", "mouse", "rat", "rabbit", "rhesus monkey", "sheep", "pig", "platypus", "alpaca", "dog", "chicken", and "ferret"
simplify.format	If applicable, remove the allelic designation ( <b>TRUE</b> ) or retain all information ( <b>FALSE</b> )

**Value**

A data frame with the new columns of formatted genes added.

**Examples**

```
data(immapex_example.data)
formatGenes(immapex_example.data[["TenX"]],
            region = "v",
            technology = "TenX")
```

---

generateSequences      *Randomly Generate Amino Acid Sequences*

---

**Description**

Use this to make synthetic amino acid sequences for purposes of testing code, training models, or noise.

**Usage**

```
generateSequences(
  prefix.motif = NULL,
  suffix.motif = NULL,
  number.of.sequences = 100,
  min.length = 1,
  max.length = 10,
  sequence.dictionary = amino.acids
)
```

**Arguments**

prefix.motif	Add defined amino acid/nucleotide sequence to the start of the generated sequences.
suffix.motif	Add defined amino acid/nucleotide sequence to the end of the generated sequences
number.of.sequences	Number of sequences to generate
min.length	Minimum length of the final sequence. The min.length may be adjusted if incongruent with prefix.motif/suffix.motif lengths
max.length	Maximum length of the final sequence
sequence.dictionary	The letters to use in sequence generation (default are all amino acids)

**Value**

A vector of generated sequences

**Examples**

```
generateSequences(prefix.motif = "CAS",
                  suffix.motif = "YF",
                  number.of.sequences = 100,
                  min.length = 8,
                  max.length = 16)
```

---

 geometricEncoder

*Geometric Encoder from Amino Acid Strings*


---

**Description**

Use this to transform amino acid sequences into a geometric encoding of the sequence.

**Usage**

```
geometricEncoder(
  input.sequences,
  method.to.use = "BLOSUM62",
  theta = pi/3,
  verbose = TRUE
)
```

**Arguments**

input.sequences	The set of amino acid sequences
method.to.use	The method or approach for the conversion: <ul style="list-style-type: none"> <li>• BLOcks SUBstitution Matrices: BLOSUM45, BLOSUM50, BLOSUM62, BLOSUM80, BLOSUM100</li> </ul>

- Point Accepted Mutation Matrices: PAM30, PAM40, PAM70, PAM120, PAM250
- theta            The angle for geometric transformation
- verbose         Print messages corresponding to the processing step

### Value

Geometric encoded amino acid sequences in a matrix

### Examples

```
new.sequences <- generateSequences(prefix.motif = "CAS",
                                  suffix.motif = "YF",
                                  number.of.sequences = 100,
                                  min.length = 8,
                                  max.length = 16)

sequence.matrix <- geometricEncoder(new.sequences,
                                    method.to.use = "BLOSUM62",
                                    theta = pi/3)
```

---

getIMGT

*Get IMGT Sequences for Specific Loci*

---

### Description

Use this to access the ImMunoGeneTics (IMGT) sequences for a specific species and gene loci. More information on IMGT can be found at [imgt.org](http://imgt.org).

### Usage

```
getIMGT(
  species = "human",
  chain = "TRB",
  sequence.type = "aa",
  frame = "inframe",
  region = "v",
  max.retries = 3,
  verbose = TRUE
)
```

### Arguments

- species         One or two-word common designation of species.
- chain           Sequence chain to access, e.g., **TRB** or **IGH**.
- sequence.type   Type of sequence - **aa** (amino acid) or **nt** (nucleotide).
- frame           Designation for **all**, **inframe**, or **inframe+gap**.
- region          Gene loci to access.
- max.retries     Number of attempts to fetch data in case of failure.
- verbose         Print messages corresponding to the processing step.

**Value**

A list of allele sequences.

**Examples**

```
TRBV_aa <- getIMGT(species = "human",
  chain = "TRB",
  frame = "inframe",
  region = "v",
  sequence.type = "aa",
  max.retries = 3)
```

---

 getIR

*Extract Immune Receptor Sequences*


---

**Description**

Use this to extract immune receptor sequences from a Single-Cell Object or the output of [combineTCR](#) and [combineBCR](#).

**Usage**

```
getIR(input.data, chains, sequence.type = "aa")
```

**Arguments**

input.data	Single-cell object or the output of <a href="#">combineTCR</a> and <a href="#">combineBCR</a> from scRepertoire
chains	Immune Receptor chain to use - <b>TRA</b> , <b>TRB</b> , <b>IGH</b> , or <b>IGL</b>
sequence.type	Extract amino acid ( <b>aa</b> ) or nucleotide ( <b>nt</b> ) sequences

**Value**

A data frame of nucleotide or amino acid sequences

---

 immapex\_AA.data

*A list of amino acid properties*


---

**Description**

A list of amino acid properties that are used for [propertyEncoder](#) function.

This includes:

- atchleyFactors
- crucianiProperties
- FASGAI
- kideraFactors

- MSWHIM
- ProtFP
- stScales
- tScales
- VHSE
- zScales

**Usage**

```
data("immapex_AA.data")
```

**Value**

List of 10 amino acid properties for 20 amino acids

---

```
immapex_blosum.pam.matrices
```

*List of amino acid substitution matrices*

---

**Description**

A list of amino acid substitution matrices, using the Point Accepted Matrix (PAM) and BLOck Substitution Matrix (BLOSUM) approaches. A discussion and comparison of these matrices are available at [PMID: 21356840](#).

- BLOSUM45
- BLOSUM50
- BLOSUM62
- BLOSUM80
- BLOSUM100
- PAM30
- PAM40
- PAM70
- PAM120
- PAM250

**Usage**

```
data("immapex_blosum.pam.matrices")
```

**Value**

List of 10 substitution matrices



---

immapex\_example.data *Example contig data for Apex*

---

### Description

Contains a collection of bulk or paired TCR sequences in the respective formats in the form of a list from the following sources:

- TenX: 10k\_Human\_DTC\_Melanoma\_5p\_nextgem\_Multiplex from [10x Website](#).
- AIRR: Human\_colon\_16S8157851 from [PMID: 37055623](#).
- Adaptive: Adaptive\_2283\_D0 from [PMID: 36220826](#).

More information on the data formats are available: [AIRR](#), [Adaptive](#), and [TenX](#).

### Usage

```
data("immapex_example.data")
```

### Value

List of 3 example data sets for 10x, AIRR and Adaptive contigs.

---

immapex\_gene.list *A list of IMGT gene names by genes, loci, and species*

---

### Description

A list of regularized gene nomenclature to use for converting for data for uniformity. Data is organized by gene region, loci and species. Not all species are represented in the data and pseudogenes have not been removed.

### Usage

```
data("immapex_gene.list")
```

### Value

List of gene nomenclature by region, loci, and species.

inferCDR

*Infer portions of the CDR Loop from Vgene data***Description**

Use this to isolate sequences from the CDR loop using the V gene annotation. When there are multiple V gene matches for a single gene, the first allelic sequence is used.

**Usage**

```
inferCDR(
  input.data,
  reference = NULL,
  chain = "TRB",
  technology = NULL,
  sequence.type = "aa",
  sequences = c("CDR1", "CDR2")
)
```

**Arguments**

input.data	Data frame output of <a href="#">formatGenes</a>
reference	IMGT reference sequences from <a href="#">getIMGT</a>
chain	Sequence chain to access, like <b>TRB</b> or <b>IGH</b>
technology	The sequencing technology employed - <b>TenX</b> , <b>Adaptive</b> , <b>AIRR</b> , or <b>Omniscope</b>
sequence.type	Type of sequence - <b>aa</b> for amino acid or <b>nt</b> for nucleotide
sequences	The specific regions of the CDR loop to get from the data, such as <b>CDR1</b> .

**Value**

A data frame with the new columns of CDR sequences added.

**Examples**

```
# Getting the Sequence Reference
data(immapex_example.data)
TRBV_aa <- getIMGT(species = "human",
                  chain = "TRB",
                  frame = "inframe",
                  region = "v",
                  sequence.type = "aa")

# Ensuring sequences are formatted to IMGT
TenX_formatted <- formatGenes(immapex_example.data[["TenX"]],
                             region = "v",
                             technology = "TenX")

# Inferring CDR loop elements
TenX_formatted <- inferCDR(TenX_formatted,
                          chain = "TRB",
                          reference = TRBV_aa,
```

```
technology = "TenX",  
sequence.type = "aa",  
sequences = c("CDR1", "CDR2"))
```

---

mutateSequences	<i>Randomly Mutate Sequences of Amino Acids</i>
-----------------	---

---

## Description

Use this to mutate or mask sequences for purposes of testing code, training models, or noise.

## Usage

```
mutateSequences(  
  input.sequences,  
  n.sequences = 1,  
  mutation.rate = 0.01,  
  position.start = NULL,  
  position.end = NULL,  
  sequence.dictionary = amino.acids  
)
```

## Arguments

input.sequences	The amino acid or nucleotide sequences to use
n.sequences	The number of mutated sequences to return
mutation.rate	The rate of mutations to introduce into sequences
position.start	The starting position to mutate along the sequence <b>Default</b> = NULL will start the random mutations at position 1
position.end	The ending position to mutate along the sequence <b>Default</b> = NULL will end the random mutations at the last position
sequence.dictionary	The letters to use in sequence mutation (default are all amino acids)

## Value

A vector of mutated sequences

## Examples

```
sequences <- generateSequences(prefix.motif = "CAS",  
                              suffix.motif = "YF",  
                              number.of.sequences = 100,  
                              min.length = 8,  
                              max.length = 16)  
  
mutated_sequences <- mutateSequences(sequences,  
                                     n.sequence = 1,  
                                     position.start = 3,  
                                     position.end = 8)
```



---

positionalEncoder      *Adding Position-Specific Information to Sequences*

---

**Description**

Use this calculate positional encoding for recurrent neural networks using sin/cos and position information.

**Usage**

```
positionalEncoder(number.of.sequences, latent.dims = NULL)
```

**Arguments**

number.of.sequences      The number of sequences to generate position information  
latent.dims      The number of latent dimensions.

**Value**

A matrix of values

**Examples**

```
position.info <- positionalEncoder(number.of.sequences = 1000,  
                                  latent.dims = 64)
```

---

probabilityMatrix      *Position Probability Matrix for Amino Acid or Nucleotide Sequences*

---

**Description**

Use this to generate a position-probability or weight matrix for a set of given sequences.

**Usage**

```
probabilityMatrix(  
  input.sequences,  
  max.length = NULL,  
  convert.PWM = FALSE,  
  background.frequencies = NULL,  
  sequence.dictionary = amino.acids,  
  padding.symbol = ".",  
  verbose = TRUE  
)
```

**Arguments**

input.sequences	The amino acid or nucleotide sequences to use
max.length	Additional length to pad, NULL will pad sequences to the max length of input.sequences
convert.PWM	Convert the matrix into a positional weight matrix using log likelihood
background.frequencies	Provide amino acid or nucleotide frequencies for the positional weight matrix. If NULL, assumes uniform likelihood.
sequence.dictionary	The letters to use in sequence generation (default are all amino acids)
padding.symbol	Symbol to use for padding at the end of sequences
verbose	Print messages corresponding to the processing step

**Value**

A matrix with position specific probabilities or weights

**Examples**

```
new.sequences <- generateSequences(prefix.motif = "CAS",
                                  suffix.motif = "YF",
                                  number.of.sequences = 100,
                                  min.length = 8,
                                  max.length = 16)

PPM.matrix <- probabilityMatrix(new.sequences)
```

---

propertyEncoder

*Encoder from Amino Acid String by Properties*

---

**Description**

Use this to transform amino acid sequences a a matrix by amino acid properties derived from dimensional reduction strategies

**Usage**

```
propertyEncoder(
  input.sequences,
  max.length = NULL,
  method.to.use = NULL,
  convert.to.matrix = TRUE,
  summary.function = NULL,
  padding.symbol = ".",
  verbose = TRUE
)
```

**Arguments**

input.sequences	The amino acid sequences to use
max.length	Additional length to pad, NULL will pad sequences to the max length of input.sequences
method.to.use	The method or approach to use for the conversion: <ul style="list-style-type: none"> <li>• Individual sets: atchleyFactors, crucianiProperties, FASGAI, kideraFactors, MSWHIM, ProtFP, stScales, tScales, VHSE, zScales"</li> <li>• Multiple Sets: c("atchleyFactors", "VHSE")</li> </ul>
convert.to.matrix	Return a matrix (TRUE) or a 3D array (FALSE)
summary.function	Return a matrix that summarize the amino acid method/property Available summaries include: "median", "mean", "sum", variance ("vars"), or Median Absolute Deviation ("mads")
padding.symbol	Symbol to use for padding at the end of sequences
verbose	Print messages corresponding to the processing step

**Value**

Converted amino acid sequences by property in a matrix or 3D array

**Examples**

```
new.sequences <- generateSequences(prefix.motif = "CAS",
                                suffix.motif = "YF",
                                number.of.sequences = 100,
                                min.length = 8,
                                max.length = 16)

sequence.matrix <- propertyEncoder(new.sequences,
                                  method.to.use = "VHSE",
                                  convert.to.matrix = TRUE)
```

---

sequenceDecoder

*One Hot Decoder from One Hot Encoded Matrix or 3D Array*


---

**Description**

Use this to transform one hot encoded sequences back into amino acid or nucleotide sequences.

**Usage**

```
sequenceDecoder(
  sequence.matrix,
  encoder = "onehotEncoder",
  aa.method.to.use = NULL,
  call.threshold = 0.5,
  sequence.dictionary = amino.acids,
```

```
padding.symbol = ".",
remove.padding = TRUE
)
```

### Arguments

`sequence.matrix` The encoded sequences to decode in an array or matrix

`encoder` The method to prepare the sequencing information - "onehotEncoder" or "propertyEncoder"

`aa.method.to.use` The method or approach to use for the conversion:

- Individual sets: `atchleyFactors`, `crucianiProperties`, `FASGAI`, `kideraFactors`, `MSWHIM`, `ProtFP`, `stScales`, `tScales`, `VHSE`, `zScales`
- Multiple Sets: `c("atchleyFactors", "VHSE")`

`call.threshold` The relative strictness of sequence calling with higher values being more stringent

`sequence.dictionary` The letters to use in sequence generation (default are all amino acids)

`padding.symbol` Symbol to use for padding at the end of sequences

`remove.padding` Remove the additional symbol from the end of decoded sequences

### Value

Decoded amino acid or nucleotide sequences

### Examples

```
new.sequences <- generateSequences(prefix.motif = "CAS",
                                suffix.motif = "YF",
                                number.of.sequences = 100,
                                min.length = 8,
                                max.length = 16)

sequence.matrix <- onehotEncoder(new.sequences,
                                convert.to.matrix = TRUE)

decoded.sequences <- sequenceDecoder(sequence.matrix,
                                    padding.symbol = ".")
```

---

tokenizeSequences	<i>Generate Tokenized Sequences from Amino Acid String</i>
-------------------	--

---

### Description

Use this to transform amino acid sequences into tokens in preparing for deep learning models.



**Usage**

```
tokenizeSequences(  
  input.sequences,  
  add.startstop = TRUE,  
  start.token = "!",  
  stop.token = "^",  
  max.length = NULL,  
  convert.to.matrix = TRUE,  
  verbose = TRUE  
)
```

**Arguments**

input.sequences	The amino acid or nucleotide sequences to use
add.startstop	Add start and stop tokens to the sequence
start.token	The character to use for the start token
stop.token	The character to use for the stop token
max.length	Additional length to pad, NULL will pad sequences to the max length of input.sequences
convert.to.matrix	Return a matrix (TRUE) or a vector (FALSE)
verbose	Print messages corresponding to the processing step

**Value**

Tokenize sequences in a matrix or vector

**Examples**

```
new.sequences <- generateSequences(prefix.motif = "CAS",  
  suffix.motif = "YF",  
  number.of.sequences = 100,  
  min.length = 8,  
  max.length = 16)  
  
sequence.matrix <- tokenizeSequences(new.sequences,  
  add.startstop = TRUE,  
  start.token = "!",  
  stop.token = "^",  
  convert.to.matrix = TRUE)
```

---

variationalSequences *Generate Similar Sequences using Variational Autoencoder*

---

**Description**

Use this to simulate sequences using a variational autoencoder (VAE) and perturbation of the probability distributions.

**Usage**

```

variationalSequences(
  input.sequences,
  encoder.function = "onehotEncoder",
  aa.method.to.use = NULL,
  number.of.sequences = 100,
  encoder.hidden.dim = c(128, 64),
  decoder.hidden.dim = NULL,
  latent.dim = 16,
  batch.size = 16,
  epochs = 50,
  learning.rate = 0.001,
  epsilon.std = 1,
  call.threshold = 0.2,
  activation.function = "relu",
  optimizer = "adam",
  disable.eager.execution = FALSE,
  sequence.dictionary = amino.acids,
  verbose = TRUE
)

```

**Arguments**

input.sequences	The amino acid or nucleotide sequences to use
encoder.function	The method to prepare the sequencing information - "onehotEncoder" or "propertyEncoder"
aa.method.to.use	The method or approach to use for the conversion: <ul style="list-style-type: none"> <li>• Individual sets: atchleyFactors, crucianiProperties, FASGAI, kideraFactors, MSWHIM, ProtFP, stScales, tScales, VHSE, zScales"</li> <li>• Multiple Sets: c("atchleyFactors", "VHSE")</li> </ul>
number.of.sequences	Number of sequences to generate
encoder.hidden.dim	A vector of the neurons to use in the hidden layers for the encoder portion of the model
decoder.hidden.dim	A vector of the neurons to use in the hidden layers for the decoder portion of the model. If NULL assumes symmetric autoencoder
latent.dim	The size of the latent dimensions
batch.size	The batch size to use for VAE training
epochs	The number of epochs to use in VAE training
learning.rate	The learning rate to use in VAE training
epsilon.std	The epsilon to use in VAE training
call.threshold	The relative strictness of sequence calling with higher values being more stringent

activation.function      The activation for the dense connected layers  
optimizer                The optimizer to use in VAE training  
disable.eager.execution      Disable the eager execution parameter for tensorflow.  
sequence.dictionary        The letters to use in sequence mutation (default are all amino acids)  
verbose                    Print messages corresponding to the processing step

**Value**

A vector of mutated sequences

**Examples**

```
## Not run:  
sequences <- generateSequences(prefix.motif = "CAS",  
                               suffix.motif = "YF",  
                               number.of.sequences = 100,  
                               min.length = 8,  
                               max.length = 16)  
  
new.sequences <- variationalSequences(sequences,  
                                     encoder = "onehotEncoder",  
                                     encoder.hidden.dim = c(256, 128),  
                                     latent.dim = 16,  
                                     batch.size = 16)  
  
## End(Not run)
```

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