

Package ‘BIGDAWG’

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Type Package

Title Case-Control Analysis of Multi-Allelic Loci

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URL <http://tools.immunogenomics.org/>,
<https://github.com/IgDAWG/BIGDAWG>

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Description Data sets and functions for chi-squared Hardy-Weinberg and case-control association tests of highly polymorphic genetic data [e.g., human leukocyte antigen (HLA) data]. Performs association tests at multiple levels of polymorphism (haplotype, locus and HLA amino-acids) as described in Pappas DJ, Marin W, Hollenbach JA, Mack SJ (2016) <doi:10.1016/j.humimm.2015.12.006>. Combines rare variants to a common class to account for sparse cells in tables as described by Hollenbach JA, Mack SJ, Thomson G, Gourraud PA (2012) <doi:10.1007/978-1-61779-842-9_14>.

License GPL (>= 3)

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Description

This is the workhorse function for the amino acid analysis.

Usage

A(Locus, loci.ColNames, genos, grp, Strict.Bin, ExonAlign, Cores)

Arguments

Locus	Locus being analyzed.
loci.ColNames	The column names of the loci being analyzed.
genos	Genotype table.
grp	Case/Control or Phenotype groupings.
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test.
ExonAlign	Exon protein alignment filtered for locus.
Cores	Number of cores to use for analysis.

Note

This function is for internal BIGDAWG use only.

A.wrapper

Amino Acid Wrapper

Description

Wrapper function for amino acid analysis.

Usage

```
A.wrapper(
  loci,
  loci.ColNames,
  genos,
  grp,
  Exon,
  EPL,
  Cores,
  Strict.Bin,
  Output,
  Verbose
)
```

Arguments

loci	Loci being analyzed.
loci.ColNames	The column names of the loci being analyzed.
genos	Genotype table.
grp	Case/Control or Phenotype groupings.
Exon	Exon(s)for targeted analysis.
EPL	Protein Alignment List.

Cores	Number of cores to use for analysis.
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test
Output	Data return carryover from main BIGDAWG function
Verbose	Summary display carryover from main BIGDAWG function

Note

This function is for internal BIGDAWG use only.

AA.df.check	<i>Contingency Table Check</i>
-------------	--------------------------------

Description

Checks amino acid contingency table data frame to ensure required variation exists.

Usage

AA.df.check(x, Strict.Bin)

Arguments

x	contingency table.
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test.

Note

This function is for internal BIGDAWG use only.

AA.df.cs	<i>Contingency Table Amino Acid ChiSq Testing</i>
----------	---

Description

Runs ChiSq test on amino acid contingency table data frames.

Usage

AA.df.cs(x, Strict.Bin)

Arguments

x	contingency table.
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test.

Note

This function is for internal BIGDAWG use only.

AAtable.builder *Amino Acid Contingency Table Build*

Description

Build Contingency Tables for Amino Acid Analysis.

Usage

```
AAtable.builder(x, y)
```

Arguments

x	Filtered alignment list element.
y	Phenotype groupings.

Note

This function is for internal BIGDAWG use only.

AlignmentFilter *Alignment Filter*

Description

Filter Protein Exon Alignment File for Specific Alleles.

Usage

```
AlignmentFilter(Align, Alleles, Locus)
```

Arguments

Align	Protein Alignment Object.
Alleles	to be pulled.
Locus	Locus to be filtered against.

Note

This function is for internal BIGDAWG use only.

AlignObj.Create *Alignment Object Creator*

Description

Create Object for Exon Protein Alignments.

Usage

```
AlignObj.Create(Loci, Release, RefTab)
```

Arguments

Loci	Loci to be bundled.
Release	IMGT/HLA database release version.
RefTab	Data of reference exons used for protein alignment creation.

Note

This function is for internal BIGDAWG use only.

AlignObj.Update *Updated Alignment Object Creator*

Description

Synthesize Object for Exon Protein Alignments.

Usage

```
AlignObj.Update(Loci, Release, RefTab)
```

Arguments

Loci	Loci to be bundled.
Release	IMGT/HLA database release version.
RefTab	Data of reference exons used for protein alignment creation.

Note

This function is for internal BIGDAWG use only.

Append.System

Append Genetic System Locus Designation to Allele String

Description

Adds genetic system (HLA/KIR) to each allele name

Usage

```
Append.System(x, df.name)
```

Arguments

x	Vector Column genotypes to append
df.name	String SystemLocus name for each allele.

Note

This function is for internal use only.

BIGDAWG

BIGDAWG Main Wrapper Function

Description

This is the main wrapper function for each analysis.

Usage

```
BIGDAWG(
  Data,
  HLA = TRUE,
  Run.Tests,
  Loci.Set,
  Exon,
  All.Pairwise = FALSE,
  Trim = FALSE,
  Res = 2,
  EVS.rm = FALSE,
  Missing = 2,
  Strict.Bin = FALSE,
  Cores.Lim = 1L,
  Results.Dir,
  Return = FALSE,
  Output = TRUE,
```

```

Merge.Output = FALSE,
Verbose = TRUE
)

```

Arguments

Data	Name of the genotype data file.
HLA	Logical Indicating whether data is HLA class I/II genotyping data only.
Run.Tests	Specifics which tests to run.
Loci.Set	Input list defining which loci to use for analyses (combinations permitted).
Exon	Numeric Exon(s) for targeted amino acid analysis.
All.Pairwise	Logical indicating whether all pairwise loci should be analyzed in haplotype analysis.
Trim	Logical indicating if HLA alleles should be trimmed to a set resolution.
Res	Numeric setting what desired resolution to trim HLA alleles.
EVS.rm	Logical indicating if expression variant suffixes should be removed.
Missing	Numeric setting allowable missing data for running analysis (may use "ignore").
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test.
Cores.Lim	Integer setting the number of cores accessible to BIGDAWG (Windows limit is 1 core).
Results.Dir	Optional, string of full path directory name for BIGDAWG output.
Return	Logical Should analysis results be returned as list.
Output	Logical Should analysis results be written to output directory.
Merge.Output	Logical Should analysis results be merged into a single file for easy access.
Verbose	Logical Should a summary of each analysis be displayed in console.

Examples

```

## Not run:
### The following examples use the synthetic data set bundled with BIGDAWG

# Haplotype analysis with no missing genotypes for two loci sets
# Significant haplotype association with phenotype
# BIGDAWG(Data="HLA_data", Run.Tests="H", Missing=0, Loci.Set=list(c("DRB1", "DQB1")))

# Hardy-Weinberg and Locus analysis ignoring missing data
# Significant locus associations with phenotype at all but DQB1
# BIGDAWG(Data="HLA_data", Run.Tests="L", Missing="ignore")

# Hardy-Weinberg analysis trimming data to 2-Field resolution with no output to files (console only)
# Significant locus deviation at DQB1
BIGDAWG(Data="HLA_data", Run.Tests="HWE", Trim=TRUE, Res=2, Output=FALSE)

## End(Not run)

```

Build.Matrix*Build Output Matrix for GL2Tab Conversion*

Description

Initializes output matrix format for GL2Tab conversion

Usage

```
Build.Matrix(System, Loci)
```

Arguments

System	Character Genetic system HLA- or KIR
Loci	The loci for header names

Note

This function is for internal use only.

buildHAPnames*Haplotype Name Builder*

Description

Builds table of names for HAPsets

Usage

```
buildHAPnames(Combn, loci)
```

Arguments

Combn	Combination of loci to extraction from genos
loci	Character vector of unique loci being analyzed.

Note

This function is for internal BIGDAWG use only.

buildHAPsets	<i>Haplotype List Builder</i>
--------------	-------------------------------

Description

Builds table of haplotypes from combinations

Usage

```
buildHAPsets(Combn, genos, loci, loci.ColNames)
```

Arguments

Combn	Combination of loci to extraction from genos
genos	The genotype columns of the loci set being analyzed.
loci	Character vector of unique loci being analyzed.
loci.ColNames	Character vector of genos column names.

Note

This function is for internal BIGDAWG use only.

cci	<i>Case-Control Odds ratio calculation and graphing</i>
-----	---

Description

cci function port epicalc version 2.15.1.0 (Virasakdi Chongsuvivatwong, 2012)

Usage

```
cci(  
  caseexp,  
  controlex,  
  casenonex,  
  controlnonex,  
  cctable = NULL,  
  graph = TRUE,  
  design = "cohort",  
  main,  
  xlab,  
  ylab,  
  xaxis,  
  yaxis,  
  alpha = 0.05,
```

```

    fisher.or = FALSE,
    exact.ci.or = TRUE,
    decimal = 2
  )

```

Arguments

caseexp	Number of cases exposed
controlex	Number of controls exposed
casenonex	Number of cases not exposed
controlnonex	Number of controls not exposed
cctable	A 2-by-2 table. If specified, will supercede the outcome and exposure variables
graph	If TRUE (default), produces an odds ratio plot
design	Specification for graph; can be "case control", "case-control", "cohort" or "prospective"
main	main title of the graph
xlab	label on X axis
ylab	label on Y axis
xaxis	two categories of exposure in graph
yaxis	two categories of outcome in graph
alpha	level of significance
fisher.or	whether odds ratio should be computed by the exact method
exact.ci.or	whether confidence limite of the odds ratio should be computed by the exact method
decimal	number of decimal places displayed

Note

This function is for internal BIGDAWG use only.

cci.pval

Case Control Odds Ratio Calculation from Epicalc

Description

Calculates odds ratio and pvalues from 2x2 table

Usage

```
cci.pval(x)
```

Arguments

x List of 2x2 matrices for calculation, output of TableMaker.

Note

This function is for internal BIGDAWG use only.

cci.pval.list	<i>Case Control Odds Ratio Calculation from Epicalc list variation</i>
---------------	--

Description

Variation of the cci.pvalue function

Usage

```
cci.pval.list(x)
```

Arguments

x	List of 2x2 matrices to apply the cci.pvalue function. List output of TableMaker.
---	---

Note

This function is for internal BIGDAWG use only.

Check.Cores	<i>Check Cores Parameters</i>
-------------	-------------------------------

Description

Check cores limitation for OS compatibility

Usage

```
Check.Cores(Cores.Lim, Output)
```

Arguments

Cores.Lim	Integer How many cores can be used.
Output	Logical Should analysis results be written to output directory.

Check.Data

Check Data Structure

Description

Check data structure for successful conversion.

Usage

```
Check.Data(Data, System, Convert)
```

Arguments

Data	String Type of output.
System	Character Genetic system HLA or KIR
Convert	String Direction for conversion.

Note

This function is for internal use only.

Check.Params

Check Input Parameters

Description

Check input parameters for invalid entries.

Usage

```
Check.Params(  
  HLA,  
  Loci.Set,  
  Exon,  
  All.Pairwise,  
  Trim,  
  Res,  
  EVS.rm,  
  Missing,  
  Cores.Lim,  
  Return,  
  Output,  
  Merge.Output,  
  Verbose  
)
```

Arguments

HLA	Logical indicating whether data is HLA class I/II genotyping data only.
Loci.Set	Input list defining which loci to use for analyses (combinations permitted).
Exon	Numeric Exon(s) for targeted amino acid analysis.
All.Pairwise	Logical indicating whether all pairwise loci should be analyzed in haplotype analysis.
Trim	Logical indicating if HLA alleles should be trimmed to a set resolution.
Res	Numeric setting what desired resolution to trim HLA alleles.
EVS.rm	Logical indicating if expression variant suffixes should be removed.
Missing	Numeric setting allowable missing data for running analysis (may use "ignore").
Cores.Lim	Integer setting the number of cores accessible to BIGDAWG (Windows limit is 1 core).
Return	Logical Should analysis results be returned as list.
Output	Logical Should analysis results be written to output directory.
Merge.Output	Logical Should analysis results be merged into a single file for easy access.
Verbose	Logical Should a summary of each analysis be displayed in console.

Note

This function is for internal use only.

Check.Params.GLS	<i>Check Input Parameters for GLS conversion</i>
------------------	--

Description

Check input parameters for invalid entries.

Usage

```
Check.Params.GLS(
  Convert,
  File.Output,
  System,
  HZY.Red,
  DRB345.Check,
  Cores.Lim
)
```

Arguments

Convert	String Direction for conversion.
File.Output	String Type of output.
System	String Genetic system (HLA or KIR) of the data being converted
HZY.Red	Logical Reduction of homozygote genotypes to single allele.
DRB345.Check	Logical Check DR haplotypes for consistency and flag unusual haplotypes.
Cores.Lim	Integer How many cores can be used.

Note

This function is for internal use only.

CheckAlleles	<i>HLA Allele Legitimacy Check for Amino Acid Analysis</i>
--------------	--

Description

Checks available alleles against data to ensure complete overlap.

Usage

```
CheckAlleles(x, y)
```

Arguments

x	Exon protein list alignment object.
y	Genotypes from data file

Note

This function is for internal BIGDAWG use only.

CheckHLA	<i>HLA Formatting Check for Amino Acid Analysis</i>
----------	---

Description

Checks data to see if HLA data is properly formatted .

Usage

CheckHLA(x)

Arguments

x All columns of HLA genotyping data.

Note

This function is for internal BIGDAWG use only.

CheckLoci	<i>HLA Loci Legitimacy Check for Amino Acid Analysis</i>
-----------	--

Description

Checks available loci against data to ensure complete overlap.

Usage

CheckLoci(x, y)

Arguments

x Loci available in exon protein list alignment object.
y Unique column names

Note

This function is for internal BIGDAWG use only.

CheckRelease	<i>Function to Check Release Versions</i>
--------------	---

Description

This updates the protein alignment used in checking HLA loci and alleles as well as in the amino acid analysis.

Usage

```
CheckRelease(Package = T, Alignment = T, Output = F)
```

Arguments

Package	Logical to check for BIGDAWG package versions
Alignment	Logical to check the IMGT/HLA database version for the alignment bundled with BIGDAWG.
Output	Should any error be written to a file

Note

Requires active internet connection.

CheckString.Allele	<i>GL String Allele Check</i>
--------------------	-------------------------------

Description

GL String check for allele ambiguity formatting

Usage

```
CheckString.Allele(x)
```

Arguments

x	GL String to check against
---	----------------------------

Note

This function is for internal use only.

CheckString.Locus	<i>GL String Locus Check</i>
-------------------	------------------------------

Description

Check GL string for loci appearing in multiple gene fields.

Usage

```
CheckString.Locus(x, Loci)
```

Arguments

x	GL String to check against
Loci	Loci to check

Note

This function is for internal use only.

Condense.EPL	<i>Condensing Exon Specific Alignments to Single Dataframe</i>
--------------	--

Description

Combines multiple Exon Specific Alignments into a single Alignment object

Usage

```
Condense.EPL(EPL.Exon)
```

Arguments

EPL.Exon	Exon-Locus Specific Amino Acid Alignment.
----------	---

Note

This function is for internal BIGDAWG use only.

Create.Null.Table *Create Empty Table*

Description

Creates matrix of NA for no result tables.

Usage

```
Create.Null.Table(Locus, Names, nr)
```

Arguments

Locus	Locus being analyzed.
Names	Column names for final matrix.
nr	Number of rows.

Note

This function is for internal BIGDAWG use only.

DRB345.Check.Wrapper *DRB345 haplotype zygosity wrapper*

Description

Checks DR haplotypes for correct zygosity and flags unanticipated haplotypes

Usage

```
DRB345.Check.Wrapper(Genotype, Loci.DR)
```

Arguments

Genotype	Row of data set data frame following DRB345 parsing
Loci.DR	DRBx Loci of interest to test for consistency

Note

This function is for internal use only.

DRB345.Check.Zygoty *DRB345 haplotype zygoty checker single locus*

Description

Checks DR haplotypes for correct zygoty and flags unanticipated haplotypes for a single DRBx

Usage

DRB345.Check.Zygoty(Locus, Genotype)

Arguments

Locus	Locus of interest to test for consistency
Genotype	Row of data set data frame following DRB345 parsing

Note

This function is for internal use only.

DRB345.Exp	<i>DRB345 Expected</i>
------------	------------------------

Description

Checks DRB1 Genotype and Returns Expected DR345 Loci

Usage

DRB345.Exp(DRB1.Genotype)

Arguments

DRB1.Genotype	DRB1 Subject Genotypes
---------------	------------------------

Note

This function is for internal use only.

DRB345.parser

DRB345 Column Processing

Description

Separates DRB345 column pair into separate columns for each locus

Usage

DRB345.parser(Tab)

Arguments

Tab Data frame of sampleIDs, phenotypes, and genotypes

Note

This function is for internal BIGDAWG use only.

Err.Log

Error Code Display and Logging

Description

Displays error codes attributable to data formatting and Locus/Allele naming. Writes to log file.

Usage

Err.Log(Output, x, y = NULL, z = NULL)

Arguments

Output	Logical indicating if Error logging should be written to a file.
x	Log Code.
y	Misc information relevant to error.
z	Misc information relevant to error.

Note

This function is for internal BIGDAWG use only.

EVSremoval	<i>Expression Variant Suffix Removal</i>
------------	--

Description

Removes expression variant suffixes from HLA alleles in the exon protein alignment object.

Usage

EVSremoval(Locus, EPList)

Arguments

Locus	Locus to be filtered against.
EPList	Exon Protein Alignment Object

Note

This function is for internal BIGDAWG use only.

Exon.Filter	<i>Filter Exon Specific Alignment Sections</i>
-------------	--

Description

Filters the ExonPtnAlign object by locus and exon.

Usage

Exon.Filter(Locus, Exon, EPL.Locus, RefExons, E.Ptn.Starts)

Arguments

Locus	Locus being analyzed.
Exon	Exon being analyzed.
EPL.Locus	ExonPtnAlign object filtered by Locus
RefExons	Reference Exon Table
E.Ptn.Starts	Exon Protein Overlay Map

Note

This function is for internal BIGDAWG use only.

ExonPtnAlign.Create *Protein Exon Alignment Formatter*

Description

Dynamically creates an alignment of Allele exons for Analysis.

Usage

ExonPtnAlign.Create(Locus, RefTab)

Arguments

Locus	Locus alignment to be formatted.
RefTab	Reference exon protein information for alignment formatting.

Note

This function is for internal BIGDAWG use only.

ExonPtnList *Exon protein alignments.*

Description

Alignment object for use in the amino acid analysis.

Usage

ExonPtnList

Format

A list where each element is an alignment dataframe for a single locus.

Filler	<i>Replace or Fill 00:00 allele strings</i>
--------	---

Description

Replaces or Fills absent allele strings.

Usage

Filler(x, Locus = NULL, Type)

Arguments

x	Genotype
Locus	Locus column to adjust.
Type	String specifying whether to pad ('Fill') or leave blank ('Remove') absent calls

Note

This function is for internal use only.

Format.Allele	<i>Ambiguous Alleles Locus Name Formatting</i>
---------------	--

Description

Remove or Append Locus name from/to allele in an ambiguous allele string

Usage

Format.Allele(x, Type)

Arguments

x	Allele String
Type	String specifying whether to strip ('off') or append ('on') locus prefix

Note

This function is for internal use only.

`Format.Tab`*Tabular Data Locus Format Tool*

Description

Correctly orders the expanded GL string

Usage

```
Format.Tab(x, Order)
```

Arguments

<code>x</code>	Single row of converted GL string
<code>Order</code>	Single row data frame for mapping converted GL strings

Note

This function is for internal use only.

`getAllele.Count`*Recompute number of alleles*

Description

Using `Freq.Final`, recompute number of alleles

Usage

```
getAllele.Count(x)
```

Arguments

<code>x</code>	Locus specific contingency matrix <code>getCS.Mat</code> output.
----------------	--

Note

This function is for internal BIGDAWG use only.

`getCS.Mat`*Chi square matrices*

Description

Chi Square contingency matrix builder with rare cell binning

Usage

```
getCS.Mat(Locus, genos.sub, Allele.Freq, Allele.Combn)
```

Arguments

Locus	Locus of interest.
genos.sub	Genotypes for locus of interest.
Allele.Freq	Allele frequencies.
Allele.Combn	Allele combinations.

Note

This function is for internal BIGDAWG use only.

`getCS.stat`*Chi square test statistic*

Description

Calculate chi square test statistic

Usage

```
getCS.stat(Locus, Freq.Final)
```

Arguments

Locus	Locus of interest.
Freq.Final	Contingency Matrix getCS.Mat output.

Note

This function is for internal BIGDAWG use only.

GetField	<i>HLA trimming function</i>
----------	------------------------------

Description

Trim a properly formatted HLA allele to desired number of fields.

Usage

```
GetField(x, Res)
```

Arguments

x	HLA allele.
Res	Resolution desired.

Note

This function is for internal BIGDAWG use only.

getFileName	<i>File Name Extraction</i>
-------------	-----------------------------

Description

Function to extract file path.

Usage

```
getFileName(x)
```

Arguments

x	File name.
---	------------

Note

This function is for internal use only.

`GetFiles`*File Fetcher*

Description

Download Protein Alignment and Accessory Files

Usage

```
GetFiles(Loci)
```

Arguments

Loci HLA Loci to be fetched. Limited Loci available.

Note

This function is for internal BIGDAWG use only.

`getHap`*Haplotype Table Maker*

Description

Builds table of haplotypes

Usage

```
getHap(SID, HaploEM)
```

Arguments

SID Index number (i.e., row number) of sample ID from genotype matrix.

HaploEM Haplotype output object from haplo.stat::haplo.em function.

Note

This function is for internal BIGDAWG use only.

getObsFreq	<i>Observed Frequency</i>
------------	---------------------------

Description

Get observed frequency of genotypes

Usage

```
getObsFreq(x, genos.locus)
```

Arguments

x	Single genotype.
genos.locus	Locus genotypes.

Note

This function is for internal BIGDAWG use only.

GL2Tab.Loci	<i>Locus Ordering for GL2Tab</i>
-------------	----------------------------------

Description

Orders Locus Calls

Usage

```
GL2Tab.Loci(Locus, Genotype, System)
```

Arguments

Locus	Locus to condense
Genotype	Row of loci to condense
System	Character Genetic system HLA or KIR

Note

This function is for internal use only.

GL2Tab.Sub	<i>Genotype List String Expander</i>
------------	--------------------------------------

Description

Expands GL string into a table of adjacent loci

Usage

GL2Tab.Sub(x, System)

Arguments

x	Character GL string to expand
System	Character Genetic system HLA or KIR

Note

This function is for internal use only.

GL2Tab.wrapper	<i>Genotype List String to Tabular Data Conversion</i>
----------------	--

Description

Expands GL strings to columns of adjacent locus pairs.

Usage

GL2Tab.wrapper(df, System, Strip.Prefix, Abs.Fill, Cores)

Arguments

df	Data frame containing GL strings
System	Character Genetic system HLA or KIR
Strip.Prefix	Logical Should System/Locus prefixes be stripped from table data.
Abs.Fill	Logical Should absent loci special designations be used.
Cores	Integer How many cores can be used

Note

This function is for internal use only

GLSconvert	<i>Genotype List String Conversion</i>
------------	--

Description

Main Workhorse wrapper for cross converting columnar table to GL string representaion.

Usage

```
GLSconvert(
  Data,
  Convert,
  File.Output = "txt",
  System = "HLA",
  HZY.Red = FALSE,
  DRB345.Check = FALSE,
  Strip.Prefix = TRUE,
  Abs.Fill = FALSE,
  Cores.Lim = 1L
)
```

Arguments

Data	String File name or R Data Frame.
Convert	String Direction for conversion.
File.Output	String Type of File.Output.
System	String Genetic system (HLA or KIR) of the data being converted
HZY.Red	Logical Reduction of homozygote genotypes to single allele.
DRB345.Check	Logical Check DR haplotypes for consistency and flag unusual haplotypes.
Strip.Prefix	Logical Should System/Locus prefixes be stripped from table data.
Abs.Fill	Logical Should absent loci special designations be used.
Cores.Lim	Integer How many cores can be used.

H.MC	<i>Haplotype Analysis Function for Multicore</i>
------	--

Description

This is the workhorse function for the haplotype analysis.

Usage

```
H.MC(genos.sub, grp, Strict.Bin, Verbose)
```

Arguments

genos.sub	The genotype columns of the loci(locus) set being analyzed.
grp	Case/Control or Phenotype groupings.
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test
Verbose	Summary display carryover from main BIGDAWG function

Note

This function is for internal BIGDAWG use only.

H.MC.wrapper

Haplotype Wrapper for Multicore

Description

Wrapper for main H function

Usage

```
H.MC.wrapper(
  SID,
  Tabsub,
  loci,
  loci.ColNames,
  genos,
  grp,
  All.Pairwise,
  Strict.Bin,
  Output,
  Verbose,
  Cores
)
```

Arguments

SID	Character vector of subject IDs.
Tabsub	Data frame of genotype calls for set being analyzed.
loci	Character vector of unique loci being analyzed.
loci.ColNames	Character vector of genos column names.
genos	The genotype columns of the loci set being analyzed.
grp	Case/Control or Phenotype groupings.
All.Pairwise	Haplotype argument carryover from main BIGDAWG function
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test
Output	Data return carryover from main BIGDAWG function
Verbose	Summary display carryover from main BIGDAWG function
Cores	Cores carryover from main BIGDAWG function

Note

This function is for internal BIGDAWG use only.

HLA_data

Example HLA Dataset

Description

A synthetic dataset of HLA genotypes for using bigdawg.

Usage

HLA_data

Format

A data frame with 2000 rows and 14 variables

HWE

Hardy Weinberg Equilibrium Function

Description

This is the main function for the HWE analysis.

Usage

HWE(Tab)

Arguments

Tab data frame of genotype files post processing.

Note

This function is for internal BIGDAWG use only.

HWE.ChiSq	<i>Hardy Weinberg Equilibrium Function</i>
-----------	--

Description

This is the workhorse function for each group analysis.

Usage

```
HWE.ChiSq(genos.sub, loci, nloci)
```

Arguments

genos.sub	data frame of genotype files post processing.
loci	list of loci.
nloci	number of loci in list

Note

This function is for internal BIGDAWG use only.

HWE.wrapper	<i>Hardy-Weinberg Wrapper</i>
-------------	-------------------------------

Description

Wrapper for main HWE function

Usage

```
HWE.wrapper(Tab, Output, Verbose)
```

Arguments

Tab	Data frame of genotype files post processing.
Output	Data return carryover from main BIGDAWG function
Verbose	Summary display carryover from main BIGDAWG function

Note

This function is for internal BIGDAWG use only.

L *Locus Analysis Function*

Description

This is the workhorse function for the locus level analysis.

Usage

```
L(loci.ColNames, Locus, genos, grp, Strict.Bin)
```

Arguments

loci.ColNames	The column names of the loci being analyzed.
Locus	Locus being analyzed.
genos	Genotype table
grp	Case/Control or Phenotype groupings.
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test

Note

This function is for internal BIGDAWG use only.

L.wrapper *Locus Wrapper*

Description

Wrapper for main L function

Usage

```
L.wrapper(nloci, loci, loci.ColNames, genos, grp, Strict.Bin, Output, Verbose)
```

Arguments

nloci	Number of loci being analyzed.
loci	Loci being analyzed.
loci.ColNames	The column names of the loci being analyzed.
genos	Genotype table
grp	Case/Control or Phenotype groupings.
Strict.Bin	Logical specify if strict rare cell binning should be used in ChiSq test
Output	Data return carryover from main BIGDAWG function
Verbose	Summary display carryover from main BIGDAWG function

Note

This function is for internal BIGDAWG use only.

make2x2	<i>Creation of a 2x2 table using the indicated orientation.</i>
---------	---

Description

make2x2 function port epicalc version 2.15.1.0 (Virasakdi Chongsuvivatwong, 2012)

Usage

```
make2x2(caseexp, controlex, casenonex, controlnonex)
```

Arguments

caseexp	Number of cases exposed
controlex	Number of controls exposed
casenonex	Number of cases not exposed
controlnonex	Number of controls not exposed

Note

This function is for internal BIGDAWG use only.

makeComb	<i>Genotype Combination Maker</i>
----------	-----------------------------------

Description

Make data frame of possible genotype combinations

Usage

```
makeComb(x)
```

Arguments

x	Number of alleles.
---	--------------------

Note

This function is for internal BIGDAWG use only.

MergeData_Output *Data Object Merge and Output*

Description

Whole data set table construction of per haplotype for odds ratio, confidence intervals, and pvalues

Usage

```
MergeData_Output(BD.out, Run, OutDir)
```

Arguments

BD.out	Output of analysis as list.
Run	Tests that are to be run as defined by Run.Tests.
OutDir	Output directory defined by Results.Dir or default.

Note

This function is for internal BIGDAWG use only.

PgrpExtract *HLA P group Finder*

Description

Identify P group for a given allele if exists.

Usage

```
PgrpExtract(x, y)
```

Arguments

x	Allele of interest.
y	Formatted P groups.

Note

This function is for internal BIGDAWG use only.

PgrpFormat	<i>HLA P group File Formatter</i>
------------	-----------------------------------

Description

Format the hla_nom_p.txt read table object for a specific locus.

Usage

PgrpFormat(x, Locus)

Arguments

x	P group object from read.table command.
Locus	Locus to be filtered on.

Note

This function is for internal BIGDAWG use only.

PreCheck	<i>Data Summary Function</i>
----------	------------------------------

Description

Summary function for sample population within data file.

Usage

PreCheck(Tab, All.ColNames, rescall, HLA, Verbose, Output)

Arguments

Tab	Loci available in exon protein list alignment object.
All.ColNames	Column names from genotype data.
rescall	HLA resolution set for analysis.
HLA	HLA BIGDAWG argument passed to function
Verbose	Summary display carryover from BIGDAWG function.
Output	Data output carryover form BIGDAWG function

Note

This function is for internal BIGDAWG use only.

prepData *Prepare imported data*

Description

Prepare imported data for processing, checks, and analysis.

Usage

```
prepData(Tab)
```

Arguments

Tab Genotypes dataframe.

Note

This function is for internal BIGDAWG use only.

rmABstrings *Replace absent allele strings*

Description

Replaces allowable absent allele strings with ^ symbol.

Usage

```
rmABstrings(df)
```

Arguments

df Genotypes dataframe.

Note

This function is for internal BIGDAWG use only.

`RunChiSq`*Strict Chi-squared Contingency Table Test*

Description

Calculates chi-squared contingency table tests and bins all rare cells.

Usage`RunChiSq(x)`**Arguments**

`x` Contingency table.

Note

This function is for internal BIGDAWG use only.

`RunChiSq_c`*Contextual Binning Chi-squared Contingency Table Test*

Description

Calculates chi-squared contingency table tests and bins rare cells at 20

Usage`RunChiSq_c(x)`**Arguments**

`x` Contingency table.

Note

This function is for internal BIGDAWG use only.

Stripper	<i>Removes System and Locus from Alleles</i>
----------	--

Description

Removes the System and Locus designations for alleles calls in GL2Tab

Usage

```
Stripper(x)
```

Arguments

x	Allele
---	--------

Note

This function is for internal use only.

summaryGeno.2	<i>Haplotype missing Allele summary function</i>
---------------	--

Description

Summary function for identifying missing alleles in a matrix of genotypes.

Usage

```
summaryGeno.2(geno, miss.val = 0)
```

Arguments

geno	Matrix of genotypes.
miss.val	Vector of codes for allele missing values.

Note

This function is for internal BIGDAWG use only and is ported from haplo.stats.

Tab2GL.Loci	<i>Locus Condenser for Tab2GL</i>
-------------	-----------------------------------

Description

Condenses alleles calls of a single locus string using "+"

Usage

Tab2GL.Loci(Locus, Genotype, System, HZY.Red)

Arguments

Locus	Locus to condense
Genotype	Row of loci to condense
System	Character Genetic system HLA or KIR
HZY.Red	Logical Should homozygote genotypes be a single allele for non-DRB345.

Note

This function is for internal use only.

Tab2GL.Sub	<i>Genotype List String Condenser</i>
------------	---------------------------------------

Description

Condenses column of loci into a GL string using "^"

Usage

Tab2GL.Sub(x, System, HZY.Red)

Arguments

x	Row of loci to condense
System	Character Genetic system HLA or KIR
HZY.Red	Logical Should homozygote genotypes be a single allele for non-DRB345.

Note

This function is for internal use only.

Tab2GL.wrapper	<i>Genotype List String to Tabular Data Conversion</i>
----------------	--

Description

Expands GL strings to columns of adjacent locus pairs.

Usage

```
Tab2GL.wrapper(df, System, HZY.Red, Abs.Fill, Cores)
```

Arguments

df	Data frame containing GL strings
System	Character Genetic system HLA or KIR
HZY.Red	Logical Should homozygote genotypes be a single allele for non-DRB345.
Abs.Fill	Logical Should absent loci special designations be used
Cores	Integer How many cores can be used.

Note

This function is for internal use only.

TableMaker	<i>Table Maker</i>
------------	--------------------

Description

Table construction of per haplotype for odds ratio, confidence intervals, and pvalues

Usage

```
TableMaker(x)
```

Arguments

x	Contingency table with binned rare cells.
---	---

Note

This function is for internal BIGDAWG use only.

UpdateRelease	<i>Update function for protein alignment upon new IMGT HLA data release</i>
---------------	---

Description

This updates the protein alignment used in checking HLA loci and alleles as well as in the amino acid analysis.

Usage

```
UpdateRelease(Force = F, Restore = F, Output = F)
```

Arguments

Force	Logical specifying if update should be forced.
Restore	Logical specifying if the original alignment file be restored.
Output	Logical indicating if error reporting should be written to file.

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