

Package ‘JaBbA’

January 24, 2017

Title What the Package Does (one line, title case)

Version 0.0.0.9000

Description What the package does (one paragraph)

Depends R (>= 3.1.1),
Rcplex (>= 0.3),
Matrix,
gTrack

Imports data.table (>= 1.9),
GenomicRanges (>= 1.18),
igraph (>= 0.7),
gUtils,
gTrack

License What license is it under?

LazyData true

RoxygenNote 5.0.1

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abs2rel

abs2rel

Description

rescales CN values from relative to "absolute" (i.e. per cancer cell copy) scale given purity and ploidy By default, output is normalized to 1 (i.e. assumes that the total relative copy number signal mass over the genome is 1)

Usage

```
abs2rel(gr, purity = NA, ploidy = NA, gamma = NA, beta = NA,
        field = "cn", field.ncn = "ncn", total = 1)
```

Arguments

gr	GRanges input with meta data field corresponding to mean relative copy "mean" in that interval
purity	purity of sample
ploidy	ploidy of sample
gamma	gamma fit of solution (over-rides purity and ploidy)
beta	beta fit of solution (over-rides purity and ploidy)
field	meta character specifying meta data field in "gr" variable from which to extract signal, default "mean"
field.ncn	character specifying meta data field in "gr" variable from which to extract germline integer copy number, default "ncn", if doesn't exist, germline copy number is assumed to be zero

Details

takes in gr with signal field "field"

Value

numeric vector of integer copy numbers

adj2inc	<i>adj2inc</i>
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Description

converts adjacency matrix (of directed graph) into incidence matrix - ie an nodes x edges matrix, for each edge i connecting node j to k, column i will have -1 at position j and +1 at position k

Usage

```
adj2inc(A)
```

all.paths	<i>all.paths</i>
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Description

Low level function to enumerate all elementary paths and cycles through graph

Usage

```
## S3 method for class 'paths'
all(A, all = F, ALL = F, sources = c(), sinks = c(),
    source.vertices = sources, sink.vertices = sinks, exclude = NULL,
    verbose = FALSE, ...)
```

Arguments

A	nxn adjacency matrix
all	logical flag, if all = T, will include all sources (parentless vertices) and sinks (childless vertices) in path computati
ALL	logical flag, if ALL = T, will also include vertices without outgoing and incoming edges in paths
sources	graph indices to treat as sources (by default is empty)
sinks	graph indices to treat as sinks (by default is empty)
verbose	logical flag

Details

takes directed graph represented by n x n binary adjacency matrix A and outputs all cycles and paths between source.vertices, sink.vertices

Value

list of integer vectors corresponding to indices in A (i.e. vertices) \$paths = paths indices \$cycles = cycle indices

annotate.walks	<i>annotate.walks</i>
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Description

Low level function to annotate walks (GRanges list) with cds / promoter annotations

Usage

```
annotate.walks(walks, cds, promoters = NULL, filter.splice = T,
  verbose = F, prom.window = 1000, max.chunk = 1e+09, mc.cores = 1,
  exhaustive = FALSE)
```

Arguments

walks	GRangesList of walks to query (eg from traversal of JaBbA object)
cds	GRangesList of CDS annotation, one per transcript (each a coding region of an exon)
promoters	GRanges of promoters (same length as transcript)
filter.splice	flag whether to filter out splice variants of a given gene
verbose	flag
prom.window	window to use around each transcript to identify putative promoter if promoter is NULL
mc.cores	number of cores to use

Details

given: walks: input grl of walks on the genome tx: gr annotating transcript boundaries or grl annotating exon level transcripts e.g. refgene or grl with grl-level meta data fields \$s1, \$s2, \$e1, \$e2 annotating start and end positions of transcript and cds respectively, \$gene_sym representing gene label, \$chr - chromosome, \$str strand and gr level features (exon_frame) annotating the frame (0,1,2) of the first exon position in a + transcript and last exon position in a - transcript. Assumes that exons are ordered in each grl item in the order of transcription (i.e. right most exon for negative strand transcripts) (e.g. output of read_refGene(grl = T))

Value

a grl of putative fusions with annotations in values field: \$label \$type e.g. promoter-fusion, 5-UTR fusion, In-frame fusion, 3' truncated fusion, 5' truncated fusion, in-frame poly-fusion, 3' truncated poly-fusion, 5' truncated poly-fusion \$genes genes involved in fusion \$transcripts transcripts involved in fusion

annotates every possible altered transcript in region including - transcripts with truncated

cbs_stub	<i>run cbs</i>
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Description

run cbs

Usage

```
cbs_stub(cov.file, out.file, field = "ratio", alpha = 1e-05)
```

chromhmpx_stub	<i>run gene-chromhmm px analysis</i>
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Description

run gene-chromhmm px analysis

Usage

```
chromhmpx_stub(rg.span.file, chromhmm.file, this.ra.file, out.file,  
  verbose = T, max.dist = 1e+06)
```

chromoplexy	<i>chromoplexy</i>
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Description

Determines chromoplexy paths from standard JaBbA output

Usage

```
chromoplexy(kag = NULL, jab = NULL, sol = NULL, all = F, ref.only = F,
  filt.jab = T, reciprocal = TRUE, hijacked = TRUE, paths = F,
  dist = 1000, cn.dist = dist, verbose = F, interval = 400,
  chunksize = 5000)
```

Arguments

jab	JaBbA object
all	logical flag whether to enumerate all possible cycles, otherwise will return (an arbitrary) minimal decomposition into the shortest "chains" of balanced rearrangements
paths	logical flag if paths = T, will also try to compute paths (in addition to cycles), default = FALSE
dist	maximum distance at which to cluster junctions, i.e. in which to consider a deletion or amplification bridge
cn.dist	minimum distance at which to enforce copy concordance for deletion and amplification bridges

Details

Outputs all chromoplexy paths and cycles (i.e. paths and cycles in breakpoint graph) allowing quasi-reciprocal rearrangements with amplification / deletion bridge distance threshold "dist"

Value

paths and cycles of as list of vectors of aberrant edge index sequences, aberrant edges refer to edges described in kag\$ab.edges matrix

collapse.paths	<i>collapse.paths</i>
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Description

collapse simple paths in a graph G (adjacency matrix or igraph object) returns m x m new adjacency matrix and map of old vertex id's to new ones \$adj = m x m matrix #map = length n with indices 1 .. m

Usage

```
collapse.paths(G, segstats, verbose = T)
```

convex.basis	<i>karyoMIP utility functions</i>
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Usage

```
convex.basis(A, interval = 80, chunksize = 100, exclude.basis = NULL,
  exclude.range = NULL, maxchunks = Inf, verbose = F)
```

Details

convex.basis

Outputs a matrix K of the convex basis of matrix A

i.e. each column $x = K[,i]$ is a minimal solution (with respect to sparsity) to $Ax = 0, x \geq 0$

`exclude.basis = 0, 1` matrix of dimension $k \times \text{ncol}(A)$ specifying k sparsity patterns that we would like to exclude from the convex.basis. This can speed up computation since any non-negative combinations of vectors that satisfy an exclusion property will also be excludable, and thus we can remove such vectors as soon as we detect them..

`exclude.range = 0, 1` matrix of dimension $k \times \text{nrow}(A)$ specifying k sparsity patterns that we would like exclude, but these are specified in terms of the range of `abs(A)` .. i.e. we want to exclude all basis vectors v such that `nz(exclude.ranges[i,] C nz(abs(A)*v))` for some pattern i . Again any non-neg linear comb of any intermediate-basis vector that satisfies this property will satisfy it, as a result we can exclude these vectors when we see them.

cumseg	<i>segment</i>
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Description

Wrapper around `cumSeg` to segment numeric data in an input `GRanges` with signal meta data field (e.g. `$signal`) Returns a `GRanges` of piecewise constant regions with their associated value

Usage

```
cumseg(gr, field = "signal", log = T, type = "bic", alg = "stepwise",
  S = 1, verbose = F, mc.cores = 1, ...)
```

fusions	<i>fusions</i>
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Description

annotates all gene fusions given an n x n adjacency matrix A of n genomic segments seg and grl of transcripts (eg output of read_RefGene) seg must be (1) a tiling of the genome and (2) have copies of both + and - intervals for each genomic range (eg output of karyograph)

Usage

```
fusions(junctions = NULL, jab = NULL, cds = NULL, promoters = NULL,
        query = NULL, prom.window = 1000, verbose = T, max.chunk = 1e+10,
        cb.interval = 10000, cb.chunksize = 10000, cb.maxchunks = 1e+10,
        exhaustive = FALSE, debug = NULL, mc.cores = 1)
```

Arguments

junctions	GRangesList of junctions (each a length 2 GRanges)
jab	JaBbA object (overrides junctions input)
cds	CDS annotations (GrangesList of transcript composed of coordinates coding regions of exons)
promoters	GRanges of promoters (same length as transcript)
query	optional query limiting walks to specific regions of interest
prom.window	window to use around each transcript to identify putative promoter if promoter is NULL

Details

alternate input is a pile of junctions of ranges with strand orientation pointing AWAY from breakend
cds = gencode cds GRanges gff3 / gtf input
"gene_name" GRangesList meta data field is used in annotation and in not creating "splice" fusions that arise from different transcripts of the same gene.

Value

GRangesList of walks corresponding to transcript boundaires

genepx_stub	<i>run gene-gene px analysis</i>
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Description

run gene-gene px analysis

Usage

```
genepx_stub(rg.span.file, this.ra.file, out.file, verbose = T,
            max.dist = 1e+06)
```

gr.tile.map	<i>gr.tile.map</i>
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Description

Given two tilings of the genome (eg at different resolution) query and subject outputs a length(query) list whose items are integer vectors of indices in subject overlapping that overlap that query (strand non-specific)

Usage

```
gr.tile.map(query, subject, mc.cores = 1, verbose = FALSE)
```

Arguments

query	Query
subject	Subject
mc.cores	number of cores
verbose	Default FALSE

Note

Assumes that input query and subject have no gaps (including at end) or overlaps, i.e. ignores end() coordinates and only uses "starts"

jabba.alleles	<i>jabba.alleles</i>
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Description

Populates allelic values for JaBbA object. This does not explicitly impose junction balance constraints on alleles, but rather just computes the maximum likelihood estimate given allelic counts and the inferred total copy number on a given segment according to JaBbA

Usage

```
jabba.alleles(jab, het.sites, alt.count.field = "alt.count.t",
  ref.count.field = "ref.count.t", baf.field = "baf.t", split.ab = F,
  uncoupled = FALSE, conservative = FALSE, verbose = F)
```

Arguments

jab	JaBbA object
het.sites	GRanges with meta data fields (see below) for alt and rref count
alt.count.field	character specifying alt.count meta data field in input het.sites (default alt.count.t)
ref.count.field	character specifying ref.count meta data field in input het.sites (default ref.count.t)

<code>split.ab</code>	logical flag whether to split aberrant segments (segments with ab edge entering or leaving prior to computing allelic states (default FALSE))
<code>uncoupled</code>	logical flag whether to not collapse segments after inferring MLE estimate (default FALSE), if FALSE will try to merge adjacent segments and populate allele-specific junctions with copy numbers on the basis of the MLE fit on individual allelic segments
<code>conservative</code>	if TRUE then will leave certain allelic segments "unphased" if one cannot sync the high / low interval state with the incoming and / or outgoing junction state
<code>verbose</code>	logical flag

Value

list with following fields: `$segstats` = GRanges of input segments with `$cn.high` and `$cn.low` segments populated `$asegstats` = GRanges of allelic segments (length is $2 * \text{length}(\text{segstats})$) with high and low segments each having `$cn`, this is a "melted" `segstats` GRanges `$atd` = gTrack of allelic segments and supporting input `het.sites` `$adj` = allelic adjacency matrix of allele specific junctions `$ab.ix` = indices of aberrant edges in `$adj` `$ref.ix` = indices of reference edges in `$adj`

JaBbA.digest

JaBbA.digest

Description

JaBbA.digest

Usage

```
JaBbA.digest(jab, kag = NULL, verbose = T, keep.all = T)
```

Arguments

<code>jab</code>	JaBbA object "undigested"
<code>kag</code>	karyograph (original karyograph input to JaBbA), if NULL then will "redigest" JaBbA object
<code>verbose</code>	logical flag
<code>keep.all</code>	<code>keep.all</code> (default TRUE) whether to keep 0 copy junctions or collapse segments across these as well

Details

processes JaBbA object (1) collapsing segments with same copy number that lack loose ends (2) (optional) +/- adds segments corresponding to loose ends (3) outputting edges data frame with colors, and other formatting information (4) outputting junctions GRangesList with copy number, color, lty and other plotting components

TODO: replace with proper object instantiator

jabba.dist	<i>jabba.dist</i>
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Description

Computes distance between pairs of intervals on JaBbA graph

Usage

```
jabba.dist(jab, gr1, gr2, matrix = T, directed = FALSE, max.dist = Inf,  
  include.internal = TRUE, verbose = FALSE, EPS = 1e-09)
```

Arguments

jab	JaBbA object
gr1	interval set 1 GRanges
gr2	interval set 2 GRanges
matrix	flag whether to output a matrix
max.dist	numeric (default = Inf), if non-infinity then output will be a sparse matrix with all entries that are greater than max.dist set to zero

Details

Given "jabba" object and input granges gr1 and gr2 of (signed) intervals

Value

a length(gr1) x length(gr2) matrix whose entries ij store the distance between the 3' end of gr1[i] and 5' end of gr2[j]

jabba.hood	<i>jabba.hood</i>
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Description

Given JaBbA object and seed window "win", outputs a reduced set of windows within neighborhood of n coordinate (or nodes) within the seed region(s) on the graph (only includes edges with weight !=0)

Usage

```
jabba.hood(jab, win, d = 0, k = NULL, pad = 0, ignore.strand = TRUE,  
  bagel = FALSE, verbose = FALSE)
```

Arguments

jab	JaBbA object
win	GRanges of window of interest
d	= distance in coordinates on graph
k	Neighborhood on graph around window of interest to query
pad	pad level at which to collapse nearly reference adjacent intervals

Value

a reduced set of windows within neighborhood k of seed on the graph (only includes edges with weight !=0)

jabba.melt	<i>jabba.melt</i>
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Description

jabba.melt

Usage

```
jabba.melt(jab, anti = FALSE, verbose = FALSE, mc.cores = 1,
           max.del = 10)
```

Arguments

jab	JaBbA object "undigested"
verbose	logical flag
kag	karyograph (original karyograph input to JaBbA), if NULL then will "redigest" JaBbA object
keep.all	keep.all (default TRUE) whether to keep 0 copy junctions or collapse segments across these as well

Details

melt JaBbA graph into "events" that decompose the total ploidy into amplicons (or deleticons, if anti = TRUE) Each amplicons / deleticon is flanked by either (1) junctions (2) loose ends or (3) chromosome ends / telomeres

jabba.walk	<i>jabba.walk</i>
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Description

Computes walks around all aberrant edges in JABbA object

Usage

```
jabba.walk(sol, kag = NULL, digested = T, outdir = "temp.walk",
  junction.ix = NULL, loci = NULL, clustersize = 100, trim = FALSE,
  trim.w = 1e+06, prune = FALSE, prune.d1 = 1e+05, prune.d2 = 1e+05,
  maxiterations = Inf, mc.cores = 1,
  genes = read.delim("~/DB/COSMIC/cancer_gene_census.tsv", strings =
  F)$Symbol, verbose = T, max.threads = 4, customparams = T, mem = 6,
  all.paths = FALSE, nomip = F, tilim = 100, nsolutions = 100,
  cb.interval = 10000, cb.chunksize = 10000, cb.maxchunks = 1e+10)
```

Arguments

<code>sol</code>	JaBbA object
<code>outdir</code>	output directory
<code>junction.ix</code>	junction indices around which to build walks (default is all junctions)
<code>loci</code>	loci around which to build walks (over-rides <code>junction.ix</code>), alternatively can be a list of "all.paths" objects (i.e. each a list utput of initial <code>all.paths = TRUE</code> run +/- field <code>\$prior</code> for walk to re-eval a given <code>all.paths</code> combo)
<code>clustersize</code>	size of the cluster to output around the locus or junction of interest
<code>trim</code>	logical flag whether trim in neighborhood of junction (only applicable if <code>loci = NULL</code> , default = <code>TRUE</code>)
<code>trim.w</code>	integer width to which to trim to
<code>prune</code>	flag whether to prune trivial walks for whom a path can be drawn from first to last interval in a graph linking intervals with pairwise distance < d1 on the walk or distance < d2 on the reference
<code>prune.d1</code>	local distance threshold for walk pruning
<code>prune.d2</code>	referenc distance threshold for walk pruning
<code>mc.cores</code>	number of cores to use, default 1
<code>genes</code>	character vector of gene symbols with which to annotate walk (eg cancer genes)
<code>verbose</code>	logical flag

Details

Takes in JaBbA solution and computes local reconstructions around all aberrant edges (default). Reconstructions (i.e. Huts) consists of collections of walks, each walk associated with a copy number, and a given region (collection of genomic windows). The interval sum of walks in a given region, weighted by copy numbers will recapitulate the marginal copy profile (as estimated by JaBbA). The reconstruction is chosen to maximize parsimony.

Optional flags allow making huts around specific junctions or specified loci (GRangesList)

Walks are reconstructed locally within "clustersize" nodes of each aberrant edge, where clustersize is measured by the number of total edges. Larger cluster sizes may fail to be computationally tractable, i.e. with a highly rearranged genome in an area of dense interconnectivity.

Value

list of walk set around each locus or junction that is inputted to analysis, each list item is a list with the following fields \$win = input locus of interest, \$grl = GRangesList of walks, \$grs is a collapsed footprint of all walks in the walk list for this locu \$td gTrack of of the output, additional outputs for debugging: \$sol, \$K, \$Bc, \$eix, \$vix, \$h

jabba2vcf	<i>jabba2vcf</i>
-----------	------------------

Description

Converts jabba output to vcf file according to 4.2 "BND" syntax

Usage

```
jabba2vcf(jab, fn = NULL, sampleid = "sample", hg = read_hg(fft = T),
  cnv = FALSE)
```

Arguments

jab	JaBbA object
fn	output file name
sampleid	sample id
hg	human genome as BSgenome or ffTrack
cnv	flag whether to dump in CNV format

Value

returns character string or writes to file if specified

jbaMIP	<i>jbaMIP</i>
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Description

jbaMIP

Usage

```
jbaMIP(adj, segstats, beta = NA, gamma = NA, field.ncn = "ncn",
  tilim = 20, mipemphasis = 0, epgap = 0.01, ploidy.min = 0.1,
  ploidy.max = 20, beta.guess = beta, beta.min = beta.guess,
  beta.max = beta.guess, gamma.guess = NA, gamma.min = gamma.guess,
  gamma.max = gamma.guess, cn.sd = 1, cn.prior = cn.sd, partition = T,
  purity.prior.mean = NA, purity.prior.sd = 0.3,
  purity.prior.strength = 1, purity.prior = c(purity.prior.mean,
  purity.prior.sd), cn.fix = rep(NA, length(segstats)), cn.lb = cn.fix,
  cn.ub = cn.fix, loose.ends = c(), adj.lb = 0 * adj, adj.nudge = 0 *
  adj, ecn.out.ub = rep(NA, length(segstats)), ecn.in.ub = rep(NA,
  length(segstats)), gurobi = F, nsolutions = 1, verbose = F, debug = F,
  mc.cores = 1, ignore.edge = FALSE, ignore.cons = TRUE,
  edge.slack = TRUE, slack.prior = 1, ...)
```

Arguments

adj	n x n adjacency matrix interpreted as binary (this is the \$adj output of karyograph)
segstats	n x 1 GRanges object with "mean" and "sd" value fields
beta	numeric guess for beta (i.e. from ppgrid)
gamma	numeric guess for gamma (i.e. from ppgrid)
field.ncn	this field takes into account normal copy number in relative to absolute conversion
adj.lb	nxn matrix of lower bounds on particular copy numbers - this is used to force certain junctions into the graph
adj.nudge	nxn adjacency matrix of "nudge" rewards on individual junctions, NOTE: maximum value in this matrix
slack.prior	1/slack.prior = penalty for each additional copy number of each slack edge, the higher slack.prior the more slack we allow in the reconstruction, should be intuitively calibrated to the expected "incompleteness" of the reconstruction, 1/slack.prior should be calibrated with respect to $1/(k*sd)^2$ for each segment, so that we are comfortable with junction balance constraints introducing k copy number deviation from a segments MLE copy number assignment (the assignment in the absence of junction balance constraints)

Details

primary "heavy" lifting task of JaBbA. Sets up optimization problem given an input graph and segstats input and sends to CPLEX via RCplex

combines edge-conservation constraints from karyograph (n x n adjacency matrix connecting n genomic intervals) with segment abundance data (segstats - length n GRanges object with mean and sd fields corresponding to posterior means and sd's on the the relative "concentration" of each interval) to infer

(1) interval and edge absolute copy numbers on the karyograph (2) purity and ploidy (3) slack edges (if any needed)

basically solves ABSOLUTE problem (fitting integer grid to continuous segment intensities) while enforcing edge-conservation constraints.

Most important optional parameters include (1) cn.sd (expected deviation of absolute copy number from ploidy) (2) ploidy.min and ploidy.max → useful for probing alternate solutions, but can be generously set (3) adj.lb → enforces minimal edge absolute copy number, eg to force aberrant adjacency use (4) edge.slack - logical variable to determine whether or not to allow penalized relaxation of edge conservation constraints (5) nsolutions - number of alternate solutions

Value

output is a Rplex solution or list of Rplex solution with additional fields, each Rplex solution is a list and the additional fields added by jbaMIP are \$adj input n x n adjacency matrix populated with integer copy numbers \$segstats input segstats vector populated with meta data fields \$cn, \$ecn.in, \$ecn.out, \$edges.out, \$slack.in, \$slack.out \$purity purity value associated with relative-absolute affine copy number conversion for this solution \$ploidy purity value associated with relative-absolute affine copy number conversion for this solution \$

Additional fields for qc / technical debugging: \$nll.cn negative log likelihood corresponding to the CN fit in this solution \$nll.opt negative log likelihood corresponding to the MLE CN without junction constraints \$residual = value of residual between copy solution and MLE fit without junction constraints \$beta beta value associated with relative-absolute affine copy number conversion for this solution \$gamma gamma value associated with relative-absolute affine copy number conversion for this solution \$gap.cn total gap between MLE fit without junction constraints and JaBbA fit \$ploidy.constraints input ploidy constraints \$beta.constraints input beta constraints \$cn.prior input cn.prior \$slack.prior input slack.prior

jbaMIP.allelic

jbaMIP.allelic

Description

Takes adj and segstats from output from jbaMIP and a granges of het.sites with \$ref.count and \$alt.count

Usage

```
jbaMIP.allelic(adj, segstats, purity, gamma, partition = T,
  slack.prior = 0.001)
```

Arguments

adj	adjacency matrix populated with total copy counts on junctions
segstats	granges tiling genome populated with total copy counts on interval, mu_high, sd_high, mu_low, sd_low variables on alleles
purity	purity from solution
gamma	gamma param from jbaMIP
slack.prior	1/slack.prior = penalty for each slack i.e. loose end copy in solution

Details

assumes segstats has fields \$cn populated and adj has copy states

Value

list with fields \$segstats = input segstats annotated with fitted cn.high, cn.low columns \$asegstats = output "allelic" segstats, with \$cn, \$parent.node, \$slack.in, \$slack.out, \$phased fields filled in \$adj = output length(segstats) x length(segstats) x 2 "allelic" adjacency matrix with inferred allelic copy numbers on edges \$aadj = flattened output 2*length(segstats) x 2* length(segstats) "allelic" adjacency matrix with inferred allelic copy numbers on edges

jbaMIP.process	<i>jbaMIP.process</i>
----------------	-----------------------

Description

process jbaMIP solution "sol" given original graph "g" (karyograph() list output) into JaBbA object

Usage

```
jbaMIP.process(sol, allelic = F)
```

Arguments

sol	JaBbA object
allelic	logical flag specifying whether object is allelic

Details

output is

Value

list with items: \$B incidence matrix of augmented graph (including slack vertices) (vertices x edges) rownames of \$B are vertex names of \$G and colnames of B are named with character version of their \$G indices (i.e. column order of B respects the original edge order in the solution)

\$e edge constraints for downstream karyoMIP, i.e the copy numbers at the edges \$e.ij numedges x 2 vertex pair matrix denoting what are the vertex pairs corresponding to the cols of \$B and entries of \$e, \$eclass, \$etype etc \$eclass id for each unique edge / anti-edge equivalence class \$etype specifies whether edge is slack or nonslack

jbaMIP.summarize	<i>jbaMIP.summarize</i>
------------------	-------------------------

Description

summarizes miqp result (i.e. multiple solutions) outputting data frame with summary info

Usage

```
jbaMIP.summarize(sol)
```

junction.paths	<i>junction.paths</i>
----------------	-----------------------

Description

Applies "pigeonhole principle" to enumerate all junction paths in a karyograph that can be proven to have copy number greater than 0

Usage

junction.paths(cn, adj)

Arguments

cn length n vector of integer copy numbers
adj nxn matrix of junction copy numbers

Details

Takes as input adjacency matrix specifying junction copy numbers and numeric vector specifying node copy numbers.

Value

list with fields \$paths list of n paths, each path i consisting of an n_i x 2 matrix specifying sequences of n_i junctions (each an ij node pair) \$mcn minimum copy number associated with path i

karyograph	<i>karyograph</i>
------------	-------------------

Description

karyograph

Usage

karyograph(junctions, tile = NULL, label.edges = FALSE)

Arguments

junctions GRangesList of junctions, where each item is a length GRanges of signed locations
tile GRanges optional existing tiling of the genome (eg a copy number segmentation) from which additional segments will be created

Details

builds graph from rearrangement breakpoints +/- copy number endpoints used for downstream jbaMIP and karyoMIP functions

Input `bpp` is a `GRangesList` of signed locus pairs describing aberrant adjacencies. The convention is as follows: Each locus in the input breakpoint pair points to the direction that is being joined by the adjacencies i.e. (-) bp points to "left" or preceding segment (+) bp points to the "right" or the following segment

eg imagine `albp1lb clbp2ld` "+" bp point to the right (eg b or d), "-" bp point to the left (a or c)

Input "tile" is a set of intervals whose endpoints are also used to partition the genome prior to the building of the karyograph.

Output karyograph connects signed genomic intervals (in a signed tiling of the reference genome) with "aberrant" and "reference" edges. Reference edges connect intervals that are adjacent in the reference genome, and aberrant edges are inferred (upstream of this) through cancer genome paired end analysis. Note that every node, edge, and path in this karyograph has a "reciprocal path"

Value

a list with the following fields `$tile` = `GRanges` of length $2 \times n$ tiling of the genome corresponding to union of rearrangement breakpoints and copy number endpoints `$G` = `igraph` object representing karyograph, here are the edge and vertex features vertex features: `$chrom`, `$start`, `$end`, `$width`, `$strand`, `$size`, `$shape`, `$border.width`, `$label`, `$chrom.ord`, `$y`, `$col`, `$weight` edge features: `$bp.id`, `$weight`, `$from`, `$to`, `$col`, `$type`, `$line.style`, `$arrow.shape`, `$width`, important: `$type` specifies which edges are "aberrant" and "reference", `$bp.id` specifies which input rearrangement (item in junctions) a given aberrant edge came from (and is NA for reference edges) `$adj` = $2n \times 2n$ adjacency matrix whose nonzero entries `ij` show the edge.id in `$G` `$ab.adj` = $2n \times 2n$ binary matrix specifying aberrant edges `$ab.edges` = `length(junctions) \times` 'from', 'to' \times '+', '-' mapping junction id's (indices into input junctions lists) to source and sink vertices, in both orientations

karyograph_stub	<i>karyograph if (is.character(tryCatch(png(paste(out.file, '.ppgrid.png', sep = ''), height = 500, width = 500), error = function(e) 'bla'))))</i>
-----------------	---

Description

karyograph if (is.character(tryCatch(png(paste(out.file, '.ppgrid.png', sep = ''), height = 500, width = 500), error = function(e) 'bla'))))

Usage

```
karyograph_stub(seg.file, cov.file, nseg.file = NULL, het.file = NULL,
  dranger.file = NULL, out.file, ra.file = NULL, force.seqlengths = NULL,
  purity = NA, ploidy = NA, field = "ratio", mc.cores = 1,
  max.chunk = 1e+08, subsample = NULL)
```

karyoMIP	<i>karyoMIP</i>
----------	-----------------

Description

MIP to locally compute walks in an existing JaBbA reconstruction, note: usually many optimal solutions to a given run. Used by jabba.walk.

Usage

```
karyoMIP(K, e, eclass = 1:length(e), kclass = NULL, prior = rep(0,
  ncol(K)), cpenalty = 1, tilim = 100, epgap = 1, nsolutions = 50,
  objsense = "max", ...)
```

Arguments

K	E x k binary matrix of k "extreme" contigs across E edges
e	edge copy numbers across E edges
eclass	edge equivalence classes, used to constrain strand flipped contigs to appear in solutions together, each class can have at most 2 members
prior	prior log likelihood of a given contig being in the karyotype
cpenalty	karyotype complexity penalty - log likelihood penalty given to having a novel contig in the karyotype, should be calibrated to prior, i.e. higher than the contig-contig variance in the prior, otherwise complex karyotypes may be favored
tilim	time limit to optimizatoin
nsolutions	how many equivalent solutions to report

Details

TODO: Make user friendly, still pretty raw

takes |E| x k matrix of k extreme paths (i.e. contigs) across e edges of the karyograph and length |E| vector of edge copy numbers (eclass), length |E| vector of edge equivalence classes (both outputs of jbaMIP.process) and computes most likely karyotypes that fit the edge copy number profile subject to some prior likelihood over the k extreme paths

Value

Rplex solution list object with additional field \$kcn for path copy number, \$kclass for k class id, \$mval for mval

karyoMIP.to.path	<i>karyoMIP.to.path</i>
------------------	-------------------------

Description

for a karyoMIP solution and associated K matrix of $n \times e$ elementary paths (input to karyoMIP), and $v \times e$ edge signed incidence matrix

Usage

```
karyoMIP.to.path(sol, K, e, gr = NULL, mc.cores = 1, verbose = T)
```

Arguments

sol	solution to karyoMIP
K	matrix of elementary paths (input to karyoMIP)
e	$nrow(K) \times 2$ edge matrix representing vertex pairs (i.e. edges to which K is referring to)
gr	optional GRanges whose names are indexed by rownames of B
mc.cores	integer number of cores
verbose	flag

Value

A list with following items: \$path length k list of paths, cycles (each item i is vector denoting sequence of vertices in G) \$is.cycle length k logical vector whose component i denotes whether path i is cyclic \$cn length k integer vector whose component i denotes copy number of contig i \$path.grl if path.grl == T

karyoseg	<i>karyoseg</i>
----------	-----------------

Description

performs preliminary segmentation on preliminary karyograph using CBS on coverage in tiles of coverage samples on the genome only segments inter-junction reference intervals above min.width (which is usually defined as 20 times the coverage width) returns the new karyograph with the addition partitions (and reference interval pairs)

Usage

```
karyoseg(kag, cov)
```

karyoSim

karyoSim

Description

Simulate (random) evolution of rearrangements according to input junctions, which are provided as a GRangesList, and grouped into "events" by events list (list of numeric vectors or of lists of numeric vectors indexing "junctions")

Usage

```
karyoSim(junctions, events = NULL, p.chrom = 0, p.wgd = 0,
  p.chromgain = 0.5, cn = NULL, req.gr = NULL, req.tel = TRUE,
  neo.tel = NULL, haploid = T, local.junction = T, local.qrs = T,
  force.event = T, lambda.chrom = 0, lambda.chromgain = lambda.chrom,
  lambda.chromloss = lambda.chrom, full = F, random.event = T,
  precedence = NULL, dist = 1000, verbose = T, ...)
```

Details

Goal of the simulation is to instantiate a collection of junctions (+/- approach some copy number profile) through a sequence of rearrangements and whole-chromosome copy changes

Junctions are sequences of signed reference intervals that are contiguous on the on the tumor genome (usually pairs)

Each event consists of either (1) a "quasi reciprocal sequence" (QRS) of junctions, implemented during a single "cell cycle", and are specified by vectors of junction indices, (2) a set of sets of junctions, specified as a list of list of junction indices, again without repetition, corresponding to complex events spanning multiple QRS's or "cell cycles" e.g. a BFB, which involve a replication step in between each QRS. The subsequent QRS's (attempted to be) instantiated in cis to the last item in the previous QRS (3) a GRanges object specifying a reference locus and meta data field \$type = "loss" or "gain" specifying one or more pieces of reference genome that should be lost or gained at a given step.

- Events are interpreted as strings of one or more "quasi reciprocal sequence" (QRS) of junctions which may be closed / cyclic (if they begin and end with the same junction index) or open. in which case they will result in at least some interval loss. We restrict QRS's to contain at most one repeated junction, and this has to be the first and the last item in the sequence. "Quasi" reciprocal means that we allow some sequence to be lost or gained in between breaks. - Every QRS is instantiated in the current genome, by mapping junctions, which are specified in haploid reference coordinates to intervals on the current genome. By default, instantiation is chosen so that the source interval of every junction in the QRS is on the same chromosome as the target interval on the previous junction in the QRS. If this is the case, then we say that the current junction "follows" the previous one in this WQRS instantiation. Quasi-reciprocity is then applied by possibly adding intervals at the site of a break (i.e. if the target interval of the previous junction is upstream of the source interval of the next junction). In situations where an instance of a subsequent junction cannot be found to follow the current junction, then the chain is either (1) interpreted as "broken", i.e. equiv to an unbalanced rearrangement (if strict.chain = F or (2) the event is discarded (if strict.chain = T) - Junctions / events can have many possible instantiations at a given round of evolution. This is because a given haploid interval on the reference can be associated with many loci on the tumor chromosome (in the simplest case, two homologues of the same chromosome) By default, the following preferences are exercised for choosing junction instantiations: (1) if a chromosome strand can be found that contains

all the intervals in the junction (2) a chromosome whose both strands contain all the intervals in the junction (3) a set of chromosome that instantiates the event as a chain of junctions these preferences can be over-ridden by specifying `instant.local` and `instant.chain` flags - After every cycle we do a "clean up" which involves (1) rejoining any pairs of broken ends that were partners at the previous iteration (2) removing any fragments that lack a telomere (if `req.tel = T`) or lack other `req.gr` (3) replacing reverse complements of chromosomes from previous iteration that were rearranged in the previous iteration with the reverse complements of their alteration products in the current iteration. - Every junction is implemented <exactly once> during the evolutionary history, i.e. lightning does not strike twice, infinite sites model

`p.chrom` = prob of chrom event at each simulation step `p.chromloss` = probability of chromosomal loss | chrom event (default 0.5) `p.chromgain` = probability of chromosomal gain | chrom event (default 0.5) `lambda.chrom` = poisson lambda of number of different chromosomes gained or lost at a chromosomal event `lambda.chromgain` = poisson lambda of number of chromosomes gained at each "gain" event (default `lambda.chrom`) `lambda.chromloss` = poisson lambda of number of chromosomes lost at each "loss" event (default `lambda.chrom`)

`p.wgd` = prob of whole genome doubling at each simulation step

Optionally can provide a copy profile `cn.profile` (GRanges tiling genome with `$cn` meta data field) and heuristic will be applied to attempt to "evolve" the simulation towards the observed copy profile (to be implemented)

Output is provided as - (if `full = F`) list with fields `$chroms` = Named GRangesList of final tumor chromosomes in terms of reference intervals `$gChain` = gChain mapping reference to tumor genome `$cn` = gRanges in reference genome of copy counts of reference intervals `$events` = data frame of event indices with field `$id` (for event id), `$desc` (see below for description) - (if `full = T`) List of lists, each item `k` corresponding to each stage `k` of evolution and containing the following fields: `$chroms` = Named GRangesList of tumor chromosomes at that stage in terms of reference intervals `$gChain` = gChain mapping reference to current genome `k` `$gChain.last` = gChain mapping last evolution step to current (from reference in first item of history) `$cn` = gRanges in reference genome of copy counts of reference intervals `$event` = list with `$id`, `$desc` that gives the id and description of event for chromosomal loss / gain `$id` = 'chromgain', or 'chromloss', `$desc` = indices chromosomes for a event, `$id` event id, `desc` = junctions involved

`done.events` = `rowSums(event2junction)==0`

karyotrack

karyo track

Usage

```
karyotrack(kag, paths = NULL, col = "red", pad = 0)
```

Arguments

<code>kag</code>	output of karyograph
<code>paths</code>	GRanges or GRangesList

Details

Takes karyograph and outputs gTrack +/- highlighting of one or more paths defined as GRanges or GRangesList (for multiple paths) Edges will only be highlighted when the exact interval pair corresponding to the edge is included in the graph

Value

gTrack of karyograph with particular nodes / edges colored with specified colors

<code>loose.ends</code>	<i>loose.ends</i>
-------------------------	-------------------

Description

takes jbaMIP output and outputs a vector of ranges on the right or left end of the intervals that have type 1-4 labels where

Usage

```
loose.ends(sol, kag)
```

Arguments

<code>sol</code>	JaBbA object
<code>kag</code>	karyograph object

Details

type1 = cn drop, no junction on this side, slack type2 = cn drop, no junction on this side slack type3 = no cn diff, used junction on other side, slack type4 = no cn diff, unused junction on other side, no slack

Value

vector of ranges on the right or left end of the intervals that have type 1-4 labels where

<code>mmatch</code>	<i>mmatch</i>
---------------------	---------------

Description

Low level utility function to match rows of matrix A to matrix B

Usage

```
mmatch(A, B, dir = 1)
```

multicoco

multicoco

Description

multi-scale coverage correction

Usage

```
multicoco(cov, numlevs = 1, base = 100, fields = c("gc", "map"),
  iterative = TRUE, presegment = TRUE, min.segwidth = 5e+06,
  mono = FALSE, verbose = T, imageroot = NULL, FUN = NULL, ...,
  mc.cores = 1)
```

Details

Given gc and mappability coverage correction at k "nested" scales finds the coverage assignment at the finest scale that yields the best correction at every scale

FUN = is a function that takes in a data frame / granges with \$reads and other covariate functions \$gc, \$map and outputs a vector of corrected read counts

cov is a constant with GRanges of coverage samples with (by default) fields \$reads, \$map, \$gc

base = is the multiple with which to perform "numlevs" additional scales of correction

munlist

munlist

Description

unlists a list of vectors, matrices, data frames into a n x k matrix whose first column specifies the list item index of the entry and second column specifies the sublist item index of the entry and the remaining columns specifies the value(s) of the vector or matrices.

Usage

```
munlist(x, force.rbind = F, force.cbind = F, force.list = F)
```

Details

force.cbind = T will force concatenation via 'cbind' force.rbind = T will force concatenation via 'rbind'

pp.nll

pp.nll

Description

computes neg log likelihood (\$nll) for purity, ploidy combo and mle abs copy numbers (\$v), returns as list

Usage

```
pp.nll(segstats, purity = NA, ploidy = NA, gamma = NA, beta = NA,
       field = "mean", field.ncn = "ncn", v = NULL)
```

Arguments

segstats	granges of segstats with field \$mean and \$sd corresponding to mean and standard deviation on estimates of fragment density
purity	numeric purity value
ploidy	numeric ploidy value
gamma	numeric gamma value (over-rides purity, ploidy)
beta	numeric beta value (over-rides purity, ploidy)
field.ncn	character specifying meta data field specifying germline copy number, default value is "ncn", if absent then will assume 2
solution	over-ride if not interested in MLE / weighted least squares fit, this should be an integer vector of length(segstats)

Details

v can be over-ridden to compute NLL for other (eg non MLE) values

Value

negative log likelihood of MLE i.e. least squares model (or supplied solution) fit

pp2gb

pp2gb

Description

converts purity / ploidy to gamma / beta (or reverse)

Usage

```
pp2gb(purity = NA, ploidy = NA, mu = NA, w = NA, gamma = NA,
      beta = NA)
```

Arguments

purity	value between 0 and 1
ploidy	value nonnegative
mu	vector of n segment averages
w	vector of n segment widths
gamma	non-negative value
beta	non-negative value

Details

takes in gr with signal field "field"

Value

list with purity / ploidy / gamma / beta entries

ppgrid	<i>ppgrid</i>
--------	---------------

Description

least squares grid search for purity and ploidy modes

Usage

```
ppgrid(segstats, allelic = FALSE, purity.min = 0.01, purity.max = 1,
       ploidy.step = 0.01, purity.step = 0.01, ploidy.min = 1.2,
       ploidy.max = 6, plot = F, verbose = F, mc.cores = 10)
```

Arguments

segstats	GRanges object of intervals with meta data fields "mean" and "sd" (i.e. output of segstats function)
allelic	logical flag, if TRUE will also look for mean_high, sd_high, mean_low, sd_low variables and choose among top solutions from top copy number according to the best allelic fit
mc.cores	integer number of cores to use (default 1)

Value

data.frame with top purity and ploidy solutions and associated gamma and beta values, for use in downstream jbaMIP

proximity	<i>proximity</i>
-----------	------------------

Description

Takes a set of n "query" elements (GRanges object, e.g. genes) and determines their proximity to m "subject" elements (GRanges object, e.g. regulatory elements) subject to set of rearrangement adjacencies (GRangesList with width 1 range pairs)

Usage

```
proximity(query, subject, ra = GRangesList(), jab = NULL, verbose = F,
          mc.cores = 1, max.dist = 1e+06)
```

Arguments

query	GRanges of "intervals of interest" eg regulatory elements
subject	GRanges of "intervals of interest" eg genes
ra	GRangesList of junctions (each a length 2 GRanges, similar to input to karyograph)
jab	existing JaBbA object (overrides ra input)
verbose	logical flag
mc.cores	how many cores (default 1)
max.dist	maximum genomic distance to store and compute (1MB by default) should the maximum distance at which biological interactions may occur

Details

This analysis makes the (pretty liberal) assumption that all pairs of adjacencies that can be linked on a karyograph path are in cis (i.e. share a chromosome) in the tumor genome.

Value

list of n x m sparse distance matrices: \$ra = subject-query distance in the rearranged genome for all loci < max.dist in tumor genome \$wt = subject-query distance in the reference genome for all loci < max.dist in tumor genome \$rel = subject-query distance in ra relative to wild type for above loci NOTE: values x_ij in these matrices should be interpreted with a 1e-9 offset to yield the actual value y_ij i.e. y_ij = x_ij-1e-9, x_ij>0, y_ij = NA otherwise (allows for sparse encoding of giant matrices)

ramip_stub	<i>run ramip</i>
------------	------------------

Description

run ramip

Usage

```
ramip_stub(kag.file, out.file, mc.cores = 1, mem = 16, tilim = 1200,
  slack.prior = 0.001, gamma = NA, beta = NA, customparams = T,
  purity.min = NA, purity.max = NA, ploidy.min = NA, ploidy.max = NA,
  edge.nudge = 0, ab.force = NULL)
```

ra_breaks	<i>ra_breaks</i>
-----------	------------------

Description

takes in either file or data frame from dranger or snowman or path to BND / SV type vcf file and returns junctions in VCF format.

Usage

```
ra_breaks(rafile, keep.features = T, seqlengths = hg_seqlengths(),
  chr.convert = T, snowman = FALSE, swap.header = NULL,
  breakpointer = FALSE, seqlevels = NULL, force.bnd = FALSE, skip = NA,
  get.loose = FALSE)
```

Details

The default output is GRangesList each with a length two GRanges whose strands point AWAY from the break. If get.loose = TRUE (only relevant for VCF)

Rcplex2	<i>over-ride Rcplex</i>
---------	-------------------------

Description

over-ride Rcplex

Usage

```
Rcplex2(cvec, Amat, bvec, Qmat = NULL, lb = 0, ub = Inf,
  control = list(), objsense = c("min", "max"), sense = "L",
  vtype = NULL, n = 1)
```

reads.to.walks	<i>reads.to.walks</i>
----------------	-----------------------

Description

Utility function to realign reads to walks.

Usage

```
reads.to.walks(bam, walks, outdir = "./test", hg = read_hg(fft = T),
               mc.cores = 1, insert.size = 1000, verbose = T)
```

Arguments

bam	path to bam file
walks	GRangesList of walks
outdir	outdir to compute into
hg	human genome sequence BSgenome or ffTrack
mc.cores	number of cores
insert.size	>= max insert size of library so that all relevant read pairs are extracted from the original bam

Details

Takes bam and collection of walks (GRanges list of signed intervals on hg19 or other BSgenome) pulls reads in regions of walks, then uses bwa mem to realign reads to walks, returns paths to new bams These bams are in "walk coordinates"

Assumes bwa mem installed an runnable from command line.

Value

paths to new bams These bams are in "walk coordinates"

rel2abs	<i>rel2abs</i>
---------	----------------

Description

rescales CN values from relative to "absolute" (i.e. per cancer cell copy) scale given purity and ploidy

Usage

```
rel2abs(gr, purity = NA, ploidy = NA, gamma = NA, beta = NA,
        field = "ratio", field.ncn = "ncn")
```

Arguments

gr	GRanges input with meta data field corresponding to mean relative copy "mean" in that interval
purity	purity of sample
ploidy	ploidy of sample
gamma	gamma fit of solution (over-rides purity and ploidy)
beta	beta fit of solution (over-rides purity and ploidy)
field	meta data field in "gr" variable from which to extract signal, default "mean"
field.ncn	meta data field in "gr" variable from which to extract germline integer copy number, default "ncn", if doesn't exist, germline copy number is assumed to be zero

Details

takes in gr with signal field "field"

Value

numeric vector of integer copy numbers

runJaBbA	<i>runJAbbA</i>
----------	-----------------

Usage

```
JaBbA(ra, abu, seg, cfield = NULL, tfield = NULL, outdir = "./",
      nseg = NULL, hets = NULL, name = "tumor", cores = 4,
      field = "ratio", subsample = NULL, tilim = 1200, edgenudge = 0.1,
      slack.penalty = 1000, overwrite = F)
```

Arguments

ra	— path to junction VCF file, dRanger txt file or rds of GRangesList of junctions (with strands oriented pointing AWAY from junction)
abu	— path to cov file, rds of GRanges or .wig / .bed file of (normalized, GC corrected) fragment density
seg	— optional path to existing segmentation, if missing then will segment abu using DNACopy with standard settings
cfield	— character, junction confidence meta data field in ra
tfield	— character, tier confidence meta data field in ra
outdir	— out directory to dump into, default ./
nseg	— optional path to normal seg file with \$cn meta data field
hets	— optional path to hets.file which is tab delimited text file with fields seqnames, start, end, alt.count.t, ref.count.t, alt.count.n, ref.count.n
name	— prefix for sample name to be output to seg file
cores	— number of cores to use (default 1)

field	— field of abu GRanges to use as fragment density signal
subsample	— numeric between 0 and 1 specifying how much to sub-sample high confidence coverage data
tilim	— timeout for jbaMIP computation (default 1200 seconds)
edgenudge	— numeric hyper-parameter of how much to nudge or reward aberrant junction incorporation, default 0.1 (should be several orders of magnitude lower than average 1/sd on individual segments), a nonzero value encourages incorporation of perfectly balanced rearrangements which would be equivalently optimal with 0 copies or more copies.
slack.penalty	— penalty to put on every loose.end copy, should be calibrated with respect to $1/(k*sd)^2$ for each segment, i.e. that we are comfortable with junction balance constraints introducing k copy number deviation from a segments MLE copy number assignment (the assignment in the absence of junction balance constraints)
overwrite	— flag whether to overwrite existing output directory contents or just continue with existing files.
nseg	— path to data.frame or GRanges rds of normal seg file with coordinates and \$cn data field specifying germline integer copy number

Details

Module to run jbaMIP + preprocessing from text file or rds input and dump files out to text. Generates the following files in the output directory: karyograph.rds — file of unpopulated karyograph as an RDS file of a list object storing the output of karyograph TODO - incorporate DNACopy into this for one stop operation jabba.rds — file storing JaBbA object jabba.simple.rds — file storing JaBbA object simplified so that segments containing all unpopulated aberrant junctions are merged jabba.raw.rds — storing raw jbaMIP solution, this may be useful for debugging and QC jabba.png, jabba.simple.png — gTrack images of the above reconstructions jabba.seg.txt — tsv file with jabba.simple solution segments jabba.seg.rds — GRanges rds with jabba.simple solution segments jabba.adj.txt — tsv file with edges (i.e. node pairs) of adjacency matrix populated with inferred copy numbers and node ids indexing segments in jabba.seg.txt jabba.vcf, jabba.simple.vcf — BND-style vcf output of junctions in JaBbA output populated with rearrangement and interval copy numbers jabba.cnv.vcf, jabba.simple.cnv.vcf — ccopy number style VCF showing jabba copy number output

segstats

segstats is a step in the JaBbA pipeline

Description

segstats is a step in the JaBbA pipeline

Usage

```
segstats(target, signal = NULL, field = "signal", asignal = NULL,
  afields = c("ref.count", "alt.count"), prior_weight = 1,
  prior_mean = NA, prior_alpha = NA, prior_beta = NA, max.chunk = 1e+08,
  max.slice = 20000, na.thresh = 0.2, subsample = NULL, mc.cores = 1,
  nsamp_prior = 1000, ksamp_prior = 100)
```


Arguments

target	GRanges of segments on which segstats will be computed
signal	GRanges of coverage from which samples will be taken
field	field of "signal" GRanges from which coverage signal will be pulled
asignal	optional GRanges corresponding width 1 biallelic snp allele counts across the genome,
afields	length 2 character vector meta data fields of asignal GRanges that will be used to get allele counts (default is ref.count, alt.count)
subsample	number between 0 and 1 with which to subsample per segment for coverage (useful for superdense coverage eg 50 bases to avoid correlations between samples due to read overlap)
mc.cores	number of cores to run on (default 1)

Details

computes posterior mean's and sd's for a target tiling GRanges of segments (target) target must be a non-overlapping gapless strandless or two stranded tiling of the genome (eg output of gr.tile) if two stranded, then every stranded interval must have a mirror image interval included

given a GRanges of signals using value field "field" of signal GRanges assuming that the signals inside each interval in "target" are independent samples from a normal distribution of unknown mean and variance

will also compute means and std deviations on a gamma posterior of the the "high" and "low" alleles for a granges "asignal" representing allelic signal ref count and tot count across a set of locations (i.e. modeling the counts as a poisson random variable) Fields are specified by afields

outputs target GRanges with "\$mean" and "\$sd" fields populated

seg_stub	<i>segment this.cov = seg2gr(read.delim(cov.file, strings = F), seqlengths = hg_seqlengths())</i>
----------	---

Description

```
segment this.cov = seg2gr(read.delim(cov.file, strings = F), seqlengths = hg_seqlengths())
```

Usage

```
seg_stub(cov.file, out.file, field = "ratio", mc.cores = 4)
```

slack.breaks	<i>slack.breaks</i>
--------------	---------------------

Description

makes "breaks" GRL from slack edges in jbaMIP segstats output (GRanges object)

Usage

```
slack.breaks(segstats)
```

Details

makes fake slack chromosomes corresponding to each nonzero slack edge and connects points on the genome in the following orientation:

+ segment source slack → - granges at endpoint of segment + segment target slack → + granges at start point of segment - segment source slack → + granges at start point of segment - segment target slack → - granges at endpoint of segment

sparse_subset	<i>sparse_subset</i>
---------------	----------------------

Description

given $k_1 \times n$ matrix A and $k_2 \times n$ matrix B returns $k_1 \times k_2$ matrix C whose entries $ij = 1$ if the set of nonzero components of row i of A is a (+/- strict) subset of the nonzero components of row j of B

Usage

```
sparse_subset(A, B, strict = FALSE, chunksize = 100, quiet = FALSE)
```

superenhancer_stub	<i>run super-enhancer analysis</i>
--------------------	------------------------------------

Description

run super-enhancer analysis

Usage

```
superenhancer_stub(rg.span.file, se.bed.file, this.ra.file, out.file,
  verbose = T, max.dist = 1e+06)
```

*vaggregate**vaggregate*

Description

same as aggregate except returns named vector with names as first column of output and values as second

Note: there is no need to ever use aggregate or vaggregate, just switch to data.table

Usage

```
vaggregate(...)
```

Arguments

... arguments to aggregate

Value

named vector indexed by levels of "by"

Author(s)

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