FunciSNP: Functional Identification of SNPs with Phenotype by Coincidence with Chromatin Biofeatures Vignette

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Introduction

FunciSNP's aim is to assist in identifying putative functional SNP. Using information from the 1000 genomes database as well as known position of GWAS tagSNP currated for a particular trait or disease, FunciSNP integrates the two data along with sequence information provided by peaks identified from high-throughput sequencing. FunciSNP assumes user will provide processed peaks identified using any available peak algorithm, such as FindPeaks.

This vignette provides a HOW-TO guide to setup and run FunciSNP on your machine. FunciSNP was developed with the idea that a FunciSNP user will have uninterupted high-speed internet access as well as a desktop machine with more than 4 multiple cores. If user is using a windows machine, multiple cores options will not work and thus total time to complete initial FunciSNP analysis will take longer than expected.

Using a 64bit Linux machine running 11.04 Ubuntu with 24gig RAM and 8 cores and academic high-speed internet access, the amount of time to complete 99 tagSNP across 20 different biofeatures took less than 30 min to complete. We anticipate it would take about 2 hours to complete on a single core.

Load FunciSNP+other useful libraries

```
> #When package is offically posted in Bioconductor, uncomment next 2 lines.
> #source("http://bioconductor.org/biocLite.R")
> #biocLite("FunciSNP");
> ## Following two packages and options() are not required to run 'FunciSNP' but
> #will enhance the analysis experience.
> #library(setwidth); ## Automatically set the value of options("width") when the
> #terminal emulator is resized
> #library(colorout); ## colorize R output on terminal emulators
> options(width=80);
> ##FunciSNP library and other related libraries needed.
> library("org.Hs.eg.db");
> library("gplots");
> library("gtools");
> library("ggplot2");
> library("matlab");
> library(FunciSNP);
> package.version("FunciSNP");
[1] "0.1.7"
```

Identify FuncySNP using published GWAS SNPs and publicly available biological features (ENCODE ChIPseq peaks)

FuncySNP()

This is the main funtion of FunciSNP. It will identify correlated SNPs which are in linkage disequilibrium (LD) to a known disease associated tagSNP. It will also determine if the correlated SNP in LD to the tagSNP overlaps a genomic biological feature. Correlated SNPs are directly imported from the current public release of the 1000 genomes database. 1000 genomes ftp servers available for the 1000 genomes public data: 1) National Center for Biotechnology Information (NCBI) ftp://ftp-trace.ncbi.nih.gov/1000genomes/; 2) European Bioinformatics Institute (EBI) ftp://ftp.1000genomes.ebi.ac.uk/vol1/.

Correlated SNPs in LD to a tagSNP and overlapping genomic biological features are known as putative functional SNPs (also defined as 'FuncySNP' elsewhere in the package.).

As an example, we collected SNPs identified by GWAS for Glioblastoma multiforme (GBM). In this example, GBM includes lower grade glioma, thus we label all objects with 'glioma'.

```
> ## Full path to the example GWAS SNP regions file for Glioblastoma
> # (collected from SNPedia on Jan 2012)
> glioma.snp <- file.path(system.file('extdata', package='FunciSNP'),
+ dir(system.file('extdata',package='FunciSNP'), pattern='.snp$'));
> #glioma.snp;
> ## Full path to the example biological features BED files
```

```
> # derived from the ENCODE project for Glioblastoma U-87 cell lines.
> glioma.bio <- system.file('extdata',package='FunciSNP');</pre>
> #glioma.bio;
> ## FunciSNP analysis, extracts correlated SNPs from the
> # 1000 genomes db ("ncbi" or "ebi") and finds overlaps between
> # correlated SNP and biological features and then
> # calculates LD (Rsquare, Dprime, distance, p-value).
> ## Depending on number of CPUs and internet connection, this step may take
> # some time. Please consider using a unix machine to access multiple cores.
> # glioma <- FuncySNP(snp.regions.file=glioma.snp,
> #
              bio.features.loc = glioma.bio,
              bio.features.TSS=FALSE);
> # glioma;
> # summary(glioma);
> ## The glioma example set above is called directly by
> data(glioma);
> glioma;
TagSNP List with 4 Tag SNPs and
 778 nearby, potentially correlated SNPs, that overlap at least one biofeature
$`R squared: 0.1`
             Total R.squared.cuff.0.1 Percent
tagSNPs
                4
                                       75.00
1kSNPs
               778
                                   64
                                         8.23
                                    3 100.00
bio.features
                 3
$ R squared: 0.5
             Total R.squared.cuff.0.5 Percent
tagSNPs
                4
                                   3 75.00
1kSNPs
               778
                                        5.66
                                   44
bio.features
                 2
                                       100.00
$ R squared: 0.9
             Total R.squared.cuff.0.9 Percent
                                        25.00
tagSNPs
                                   1
1kSNPs
               778
                                         1.67
                                   13
                                    2 100.00
bio.features
                 2
> summary(glioma);
TagSNP List with 4 Tag SNPs and
 778 nearby, potentially correlated SNPs, that overlap at least one biofeature
Number of potentially correlated SNPs
overlapping at least x biofeatures, per Tag SNP at an R squared of
$`R squared: 0.1 in 4 Tag SNPs with a total of `
                        bio.1 bio.2
rs4977756
                            3
                                  2
rs498872
                            9
rs6010620
                           52
                                  9
TOTAL # CORRELATED SNPS
                           64
                                 11
```

```
$`R squared: 0.5 in 4 Tag SNPs with a total of `
                         bio.1 bio.2
rs4977756
                                    0
rs498872
                             2
                                    0
rs6010620
                            40
                                    6
TOTAL # CORRELATED SNPS
                            44
                                    6
$`R squared: 0.9 in 2 Tag SNPs with a total of `
                         bio.1
rs6010620
                            13
TOTAL # CORRELATED SNPS
                            13
> class(glioma);
[1] "TSList"
attr(,"package")
[1] "FunciSNP"
```

Annotating newly identified FuncySNPs

All known genomic features (exon, intron, 5'UTR, 3'UTR, promoter, lincRNA or in gene desert (intergentic)) are used to annotate each newly identified FuncySNP. Information described in this data.frame() is used for all summary plots, table, and to output results in BED format. This step should be completed after running FuncySNP().

```
> glioma.anno <- FunciSNPAnnotateSummary(glioma);</pre>
> class(glioma.anno);
[1] "data.frame"
> gl.anno <- glioma.anno;</pre>
> ## remove rownames for this example section.
> rownames(gl.anno) <- c(1:length(rownames(gl.anno)))</pre>
> dim(gl.anno);
[1] 862 28
> head(gl.anno); ##
  chromosome bio.feature.start bio.feature.end
                                                        bio.feature corr.snp.id
1
           5
                        1200710
                                         1201809 knownGene.TSS.hg19 chr5:1200720
                                         1201809 knownGene.TSS.hg19 chr5:1200766
2
           5
                        1200710
3
           5
                        1200710
                                         1201809 knownGene.TSS.hg19 chr5:1200817
4
           5
                        1200710
                                         1201809 knownGene.TSS.hg19 chr5:1200946
5
           5
                        1200710
                                         1201809 knownGene.TSS.hg19 chr5:1200976
           5
                        1200710
                                         1201809 knownGene.TSS.hg19 chr5:1201033
                                                    D.prime
  corr.snp.position tag.snp.id tag.snp.position
                                                                R.squared p.value
                                                          NA
                                                                       NA
1
            1200720
                     rs2736100
                                          1286516
                                                                                 1
2
            1200766
                     rs2736100
                                          1286516
                                                          NA
                                                                        NA
                                                                                 1
3
            1200817
                     rs2736100
                                          1286516
                                                          NA
                                                                                 1
```

```
4
            1200946 rs2736100
                                          1286516
                                                                                  1
                                                          NΑ
5
                                          1286516 1.0000000 0.0022585199
            1200976 rs2736100
                                                                                  1
6
            1201033 rs2736100
                                          1286516 0.1795671 0.0004069606
                                                                                  1
  distance.from.tag population.count population nearest.lincRNA.ID
              -85796
                                   286
                                                       TCONS_00010241
                                               ASN
1
2
              -85750
                                   286
                                                       TCONS_00010241
                                               ASN
3
              -85699
                                   286
                                               ASN
                                                       TCONS_00010241
4
              -85570
                                   286
                                               ASN
                                                       TCONS_00010241
5
              -85540
                                   286
                                               ASN
                                                       TCONS_00010241
6
              -85483
                                   286
                                               ASN
                                                       TCONS_00010241
  nearest.lincRNA.distancetoFeature nearest.lincRNA.coverage
                               -39302
1
                                                       upstream
2
                               -39348
                                                       upstream
3
                               -39399
                                                       upstream
4
                               -39528
                                                       upstream
5
                               -39558
                                                       upstream
6
                               -39615
                                                       upstream
  nearest.TSS.GeneSymbol
                                  nearest.TSS.refseq nearest.TSS.ensembl
                  SLC6A19 NM_001003841; NP_001003841
                                                          ENSG00000174358
                  SLC6A19 NM_001003841; NP_001003841
2
                                                          ENSG00000174358
3
                  SLC6A19 NM_001003841; NP_001003841
                                                          ENSG00000174358
4
                  SLC6A19 NM_001003841; NP_001003841
                                                          ENSG00000174358
5
                  SLC6A19 NM_001003841; NP_001003841
                                                          ENSG00000174358
6
                  SLC6A19 NM_001003841; NP_001003841
                                                          ENSG00000174358
  nearest.TSS.coverage nearest.TSS.distancetoFeature Promoter utr5 Exon Intron
               upstream
                                                   -990
                                                             YES
                                                                    NO
                                                                         NΩ
                                                                                 NΩ
2
                                                   -944
                                                             YES
                                                                    NO
                                                                         NO
                                                                                 NO
               upstream
3
                                                                                 NO
                                                   -893
                                                             YES
                                                                    NO
                                                                         NO
               upstream
                                                   -764
                                                                         NO
                                                                                 NO
4
                                                             YES
                                                                    NO
               upstream
5
               upstream
                                                   -734
                                                             YES
                                                                    NO
                                                                         NO
                                                                                 NO
6
                                                   -677
                                                             YES
                                                                    NO
                                                                         NO
                                                                                 NO
               upstream
  utr3 Intergenic
1
    NO
               NO
2
    NO
                NO
3
    NO
                NO
4
    NO
                NO
5
    NO
                NO
    NO
                NO
> names(gl.anno);
 [1] "chromosome"
                                            "bio.feature.start"
 [3] "bio.feature.end"
                                            "bio.feature"
 [5] "corr.snp.id"
                                            "corr.snp.position"
 [7] "tag.snp.id"
                                            "tag.snp.position"
 [9] "D.prime"
                                            "R.squared"
[11] "p.value"
                                            "distance.from.tag"
[13] "population.count"
                                            "population"
[15] "nearest.lincRNA.ID"
                                            "nearest.lincRNA.distancetoFeature"
[17] "nearest.lincRNA.coverage"
                                            "nearest.TSS.GeneSymbol"
```

```
[19] "nearest.TSS.refseq"
                                      "nearest.TSS.ensembl"
                                      "nearest.TSS.distancetoFeature"
[21] "nearest.TSS.coverage"
                                      "utr5"
[23] "Promoter"
[25] "Exon"
                                      "Intron"
[27] "utr3"
                                      "Intergenic"
> summary(gl.anno[,c(1:18,20:28)]);
 chromosome
                  bio.feature.start
                                    bio.feature.end
Length:862
                  Min. : 1200710
                                    Min. : 1201809
Class : character
                 1st Qu.: 62295044
                                    1st Qu.: 62295926
Mode :character
                 Median : 62326155
                                    Median : 62337392
                  Mean : 65165595
                                    Mean : 65169512
                  3rd Qu.: 62374564
                                    3rd Qu.: 62376020
                  Max. :118531575
                                    Max. :118532674
           bio.feature
                                corr.snp.id corr.snp.position
knownGene.TSS.hg19:372 chr11:118442863: 2 Min. : 1200720
TFBS_Nrsf_U87 : 22 chr11:118443036: 2 1st Qu.: 62295889
TFBS_Pol2_U87
                 :468 chr11:118443046: 2 Median: 62327508
                        chr11:118478342: 2 Mean : 65167605
                        chr20:62289690 : 2 3rd Qu.: 62375255
                        chr20:62289873 : 2 Max. :118532636
                        (Other)
                                    :850
    tag.snp.id tag.snp.position
                                    D.prime
                                                      R.squared
rs2736100: 96 Min. : 1286516 Min. :7.835e-04 Min.
                                                           :9.520e-08
               1st Qu.: 62309839
rs4977756: 25
                                  1st Qu.:9.338e-01 1st Qu.:7.765e-04
rs498872 :166 Median : 62309839
                                  Median :1.000e+00 Median :4.501e-03
rs6010620:575
               Mean : 65163135
                                  Mean :8.995e-01 Mean
                                                          :1.258e-01
               3rd Qu.: 62309839
                                  3rd Qu.:1.000e+00 3rd Qu.:2.804e-02
                                  Max. :1.000e+00 Max.
               Max. :118477367
                                                           :9.776e-01
                                  NA's
                                        :4.710e+02 NA's
                                                          :4.710e+02
   p.value
                    distance.from.tag population.count population
      :2.115e-163 Min. :-100000 Min. :286.0
Min.
                                                  ASN: 96
1st Qu.: 1.000e+00
                   1st Qu.: -19966
                                    1st Qu.:379.0
                                                    EUR: 766
Median : 1.000e+00
                    Median : 13942
                                    Median :379.0
Mean : 7.989e-01
                    Mean : 4470
                                    Mean :368.6
                    3rd Qu.: 25290
3rd Qu.: 1.000e+00
                                     3rd Qu.:379.0
                    Max. : 67371
Max. : 1.000e+00
                                    Max. :379.0
     nearest.lincRNA.ID nearest.lincRNA.distancetoFeature
TCONS_00010241: 96
                      Min. :-265183
                       1st Qu.: -92280
TCONS_00015797: 25
TCONS_00020001:166
                       Median : 59111
TCONS_00027984: 26
                       Mean :
                                 2073
                       3rd Qu.: 73343
TCONS_00028269:549
                       Max. : 246019
nearest.lincRNA.coverage
                          nearest.TSS.GeneSymbol
                                                    nearest.TSS.ensembl
```

TNFRSF6B

downstream:565

:305 ENSG00000243509:305

```
: 86
                          PHLDB1
                                                     ENSG00000019144: 86
inside
           : 9
                                                    ENSG00000197114: 68
upstream
          :288
                          ZGPAT
                                        : 68
                          RTEL1; TNFRSF6B: 37
                                                     ENSG00000229299: 59
                          SLC6A18
                                        : 34
                                                     ENSG00000026036: 37
                                        :202
                                                     ENSG00000244977: 36
                          (Other)
                                        :130
                                                     (Other)
                                                                    :271
                          NA's
nearest.TSS.coverage nearest.TSS.distancetoFeature Promoter
                                                                utr5
                             :-16454.0
                                                     NO:694
                                                               NO:825
downstream:103
                      Min.
          :311
                      1st Qu.: -3117.0
inside
                                                     YES:168
                                                               YES: 37
                                 -76.0
upstream :448
                      Median:
                      Mean
                                 890.4
                      3rd Qu.:
                               2305.8
                      Max.
                             : 28781.0
 Exon
           Intron
                      utr3
                               Intergenic
NO:776
          NO:413
                     NO:702
                               NO:810
YES: 86
          YES:449
                     YES:160
                               YES: 52
> rm(gl.anno);
```

Summary table used to describe newly identified FuncyS-NPs

Using a specified Rsquare value (0-1) to subset the data, a table is generated which summarizes the total number of FuncySNPs, associated tagSNPs, and number of overlapping biofeatures. This will provide user a first look at the total number of available FuncySNP at a particular Rsquare cutoff. If geneSum is set to TRUE, a list of gene names is reported instead.

> FunciSNPtable(glioma.anno, rsq=0.5);

```
Total R.squared.cuff.0.5 Percent tagSNPs 4 3 75.00 1kSNPs 778 44 5.66 bio.features 2 2 100.00
```

> FunciSNPtable(glioma.anno, rsq=0.5, geneSum=TRUE);

Summary of correlated SNPs overlapping biofeatures

This function helps in determining the number of correlated SNPs overlapping a number of different biofeatures. This is similar to running 'glioma' above. But

now you can specifically call the function and set a pre-determined rsq value to subset the data and thereby obtain a more objective result.

> FunciSNPsummaryOverlaps(glioma.anno)

	bio.1	bio.2
rs2736100	41	0
rs4977756	12	0
rs498872	59	3
rs6010620	236	40
TOTAL # CORRELATED SNPS	348	43

> FunciSNPsummaryOverlaps(glioma.anno, rsq=0.5)

	bio.1	bio.2
rs4977756	2	0
rs498872	2	0
rs6010620	40	6
TOTAL # CORRELATED SNPS	44	6

Summary of correlated SNPs for a number of different tagSNPs

After running FunciSNPsummaryOverlaps(), the next question one would like to know is which correlated SNPs overlapping a number of different biofeatures for a number of associated tagSNP. Thus, in the example above, we have determined that we are interested in learning more about the correlated SNPs associated with 'rs6010620' and which overlap at least 2 different biofeatures.

```
> rs6010620 <- FunciSNPidsFromSummary(glioma.anno, tagsnpid="rs6010620",</p>
          num.features=2, rsq=0.5)
```

```
> summary(rs6010620);
```

```
chromosome
                  bio.feature.start bio.feature.end
Length:12
                  Min. :62326155 Min.
                                            :62330994
Class : character
                  1st Qu.:62329895
                                     1st Qu.:62337392
Mode :character
                  Median :62354158
                                     Median :62355398
                         :62351007
                  Mean
                                     Mean
                                            :62353861
                  3rd Qu.:62370211
                                     3rd Qu.:62371310
                  Max.
                         :62371621
                                     Max.
                                            :62372970
```

```
bio.feature
                          corr.snp.id corr.snp.position
                                                            tag.snp.id
knownGene.TSS.hg19:6
                                     Min.
                                                       rs2736100: 0
                       rs1056441:2
                                            :62330439
TFBS_Nrsf_U87
                 :0
                       rs1291209:2
                                      1st Qu.:62330484
                                                       rs4977756: 0
TFBS_Pol2_U87
                 :6
                       rs1295810:2
                                     Median :62354704
                                                        rs498872 : 0
                       rs1741708:2
                                     Mean :62352184
                                                        rs6010620:12
                       rs6062498:2
                                      3rd Qu.:62370732
                       rs6122159:2
                                     Max.
                                            :62372041
                       (Other) :0
tag.snp.position
                     D.prime
                                     R.squared
```

p.value Min. :62309839 Min. :0.5073 Min. :1.555e-127 Min. :0.8380

```
1st Qu.:62309839
                 1st Qu.:0.8979
                                   1st Qu.:0.5365 1st Qu.:1.555e-127
Median :62309839 Median :0.9204
                                   Median :0.7588 Median :1.868e-117
                                   Mean :0.6967
Mean :62309839 Mean :0.9039
                                                   Mean : 1.046e-81
3rd Qu.:62309839
                 3rd Qu.:0.9234
                                   3rd Qu.:0.8092 3rd Qu.: 2.955e-89
Max. :62309839 Max. :0.9234
                                   Max. :0.8092 Max. : 6.274e-81
distance.from.tag population.count population
                                                 nearest.lincRNA.ID
       :20600
                 Min.
                       :379
                                  ASN: 0
                                             TCONS_00010241: 0
1st Qu.:20645
                 1st Qu.:379
                                  EUR:12
                                             TCONS_00015797: 0
Median :44865
                 Median:379
                                             TCONS_00020001: 0
Mean :42345
                 Mean :379
                                             TCONS_00027984: 0
3rd Qu.:60893
                 3rd Qu.:379
                                             TCONS_00028269:12
Max. :62202
                 Max. :379
nearest.lincRNA.distancetoFeature nearest.lincRNA.coverage
Min. : 71755
                                 downstream:12
1st Qu.: 71800
                                 inside : 0
Median : 96020
                                 upstream : 0
Mean : 93500
3rd Qu.:112048
Max. :113357
nearest.TSS.GeneSymbol
SLC2A4RG:6
TNFRSF6B:4
ZGPAT :2
ARCN1
       :0
ARFRP1 :0
CDKN2B :0
(Other) :0
NM_020062; NP_064446
NM_003823; NP_003814
NM_001083113; NM_001195653; NM_001195654; NM_032527; NM_181485; NP_001076582; NP_001182582; NP_0
NM_001003841; NP_001003841
{\tt NM\_001037335; NM\_033405; NP\_001032412; NP\_208384}
NM_001080441;NP_001073910
(Other)
    nearest.TSS.ensembl nearest.TSS.coverage nearest.TSS.distancetoFeature
ENSG00000125520:6
                                            Min. : 265
                        downstream:4
ENSG00000243509:4
                        inside :8
                                            1st Qu.: 726
ENSG00000197114:2
                        upstream :0
                                            Median:1764
ENSG00000019144:0
                                            Mean :1566
                                             3rd Qu.:2418
ENSG00000026036:0
ENSG00000049656:0
                                             Max. :2463
(Other)
Promoter utr5
                          Intron
                                 utr3
                                          Intergenic
                  Exon
                 NO :12
                          NO :8
                                  NO :6
NO:12 NO:12
                                          NO :10
```

YES:6

YES: 2

YES:4

YES: 0

YES: 0 YES: 0

```
> dim(rs6010620);
[1] 12 28
> class(rs6010620);
[1] "data.frame"
> ## See FunciSNPbed to visualize this data in a genome browser.
```

Plot FunciSNP results

FunciSNPplot is a function developed to plot various types of plots to summarize and assist end-user in making informed discoveries of FunciSNP results. Plots can be stored in a folder for future reference. Most plots were created in publication standard.

The following example plots the distribution of the Rsquare values for each correlated SNP. We recommend attempting this plot before subsetting any data by a specified rsq value. The distribution helps to identify a specific Rsquare value that will provide the most informative discovery.

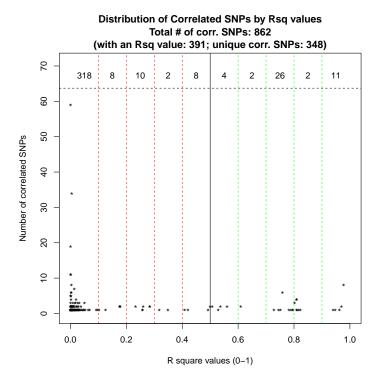


Figure 1: Distribution of Rsquare values of all Correlated SNPs. Each marked bin contains the total number of correlated SNPs. The sum of all the counts would total the number of correlated SNPs.

Using splitbysnp argument, the same type of plot as above is generated, however the total number of correlated SNPs are divided by the associated tagSNP.

```
> FunciSNPplot(glioma.anno, splitbysnp=TRUE)
> ggsave("glioma_dist_bysnp.pdf")
```

Distribution of correlated SNPs for each tagSNP

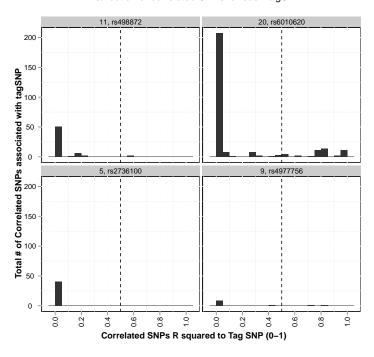


Figure 2: Distribution of Rsquare values of all Correlated SNPs divided by the tagSNP and it's location.

Using genomicSum argument will result the overall genomic distribution of the newly identified correlated SNPs. Using rsq value, the plot is divided into all correlated SNPs vs subset. This type of plot is informs the relative enrichment for genomic features.

```
> pdf("glioma_genomic_sum_rcut.pdf")
> FunciSNPplot(glioma.anno, rsq=0.5, genomicSum=TRUE, save=FALSE)
> dev.off()
pdf
2
```

tagSummary argument is unique in that it will automatically save the plots in a specific folder. This is done because this function will generate a summary plot for each biofeatures. The first plot is a scatter plot showing the relationship between Rsquare and Distance to tagSNP for each FuncySNP. The second plot is a histogram distribution of number of correlated SNPs at each Rsquare value. Each set of plot is further divided by tagSNP. Best if used with rsq value.

```
> ## Following will output a series of plots for each biofeature at rsq=0.5
> FunciSNPplot(glioma.anno, tagSummary=TRUE, rsq=0.5)
```

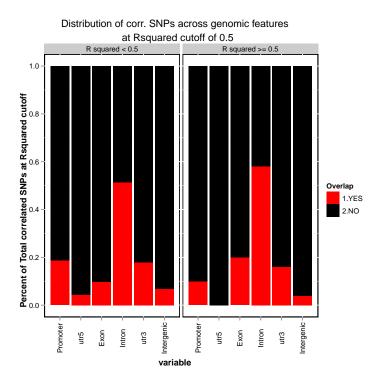


Figure 3: Stacked bar chart summarizing all correlated SNPs for each of the identified genomic features: exon, intron, 5UTR, 3UTR, promoter, lincRNA or in gene desert. Rsquare cutoff at 0.5. This plot is most informative if used with a rsq value.

Finished plotting 1/3Finished plotting 2/3Finished plotting 3/3

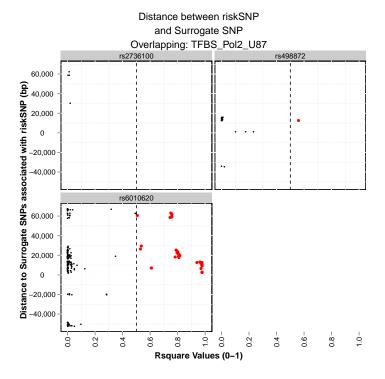


Figure 4: Scatter plot showing the relationship between Rsquare and Distance to tagSNP for each FuncySNP

Output results in BED format - visualize results

FunciSNPbed outputs a unique BED file which can be used to view in any genomic browser compatible with BED formats. To learn more about BED formats, see UCSC Genome Browser FAQ (http://genome.ucsc.edu/FAQ/FAQformat). Each tagSNP which is in LD to a corresponding FuncySNP overlapping at least one biofeature is colored black, while the FuncySNP is colored red. The initial position is provided by the first tagSNP and the first linked FuncySNP. We recommend using UCSC genome browser to view your BED files. This is useful so you can view all public and private tracks in relation to FunciSNP results.

```
> ## will output to current working directory.
> FunciSNPbed(glioma.anno, rsq=0.5);

Total corSNP (RED): 44
Total tagSNP (BLK): 3
> # FunciSNPbed(rs6010620, rsq=0.5);
```

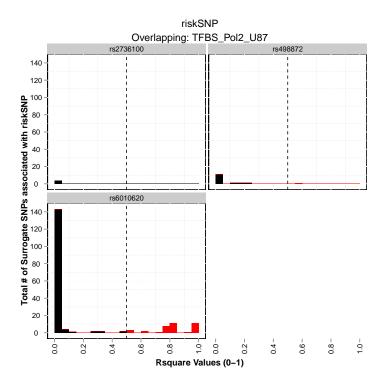


Figure 5: Histogram distribution of number of correlated SNPs at each Rsquare value

> sessionInfo()

R version 2.14.1 (2011-12-22)

Platform: x86_64-pc-linux-gnu (64-bit)

locale:

[1] LC_CTYPE=en_US.UTF-8 LC_NUMERIC=C

[3] LC_TIME=en_US.UTF-8 LC_COLLATE=en_US.UTF-8
[5] LC_MONETARY=en_US.UTF-8 LC_MESSAGES=en_US.UTF-8

[7] LC_PAPER=C LC_NAME=C

[9] LC_ADDRESS=C LC_TELEPHONE=C

[11] LC_MEASUREMENT=en_US.UTF-8 LC_IDENTIFICATION=C

attached base packages:

[1] tools splines grid stats graphics grDevices utils

[8] datasets methods base

other attached packages:

- [1] FunciSNP_0.1.7
- [2] VariantAnnotation_1.0.5
- [3] TxDb.Hsapiens.UCSC.hg19.knownGene_2.6.2
- [4] GenomicFeatures_1.6.7

```
[5] ChIPpeakAnno_2.2.0
 [6] limma_3.10.2
 [7] GO.db_2.6.1
 [8] BSgenome.Ecoli.NCBI.20080805_1.3.17
 [9] BSgenome_1.22.0
[10] multtest_2.10.0
[11] biomaRt_2.10.0
[12] GGtools_4.0.0
[13] ff_2.2-4
[14] bit_1.1-8
[15] annotate_1.32.1
[16] GGBase_3.14.0
[17] genefilter_1.36.0
[18] snpStats_1.4.1
[19] Matrix_1.0-3
[20] lattice_0.20-0
[21] survival_2.36-12
[22] rtracklayer_1.14.4
[23] RCurl_1.9-5
[24] Rsamtools_1.6.3
[25] Biostrings_2.22.0
[26] GenomicRanges_1.6.4
[27] IRanges_1.12.5
[28] matlab_0.8.9
[29] ggplot2_0.8.9
[30] proto_0.3-9.2
[31] reshape_0.8.4
[32] plyr_1.7.1
[33] gplots_2.10.1
[34] KernSmooth_2.23-7
[35] caTools_1.12
[36] bitops_1.0-4.1
[37] gdata_2.8.2
[38] gtools_2.6.2
[39] org.Hs.eg.db_2.6.4
[40] RSQLite_0.11.1
[41] DBI_0.2-5
[42] AnnotationDbi_1.16.11
[43] Biobase_2.14.0
loaded via a namespace (and not attached):
[1] digest_0.5.1
                    MASS_7.3-16
                                     parallel_2.14.1 XML_3.9-2
```

Questions or comments, please contact Simon G. Coetzee (scoetzee NEAR gmail POINT com) or Houtan Noushmehr (houtana NEAR gmail POINT com).

zlibbioc_1.0.0

[5] xtable_1.6-0