

main

December 10, 2020

1 Extract unique female specific SZ-associated genes

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[1]: import functools
import numpy as np
import pandas as pd
from scipy.stats import mannwhitneyu
from statsmodels.stats.multitest import fdrcorrection

[2]: @functools.lru_cache()
def get_res_df():
    return pd.read_csv('.././.././.././interaction_sex_sz/cmc_dlpfc/_m/genes/
↳residualized_expression.tsv', sep='\t').T

@functools.lru_cache()
def get_pheno_df():
    return pd.read_csv('/ceph/users/jbenja13/projects/sex_sz_ria/input/
↳commonMind/phenotypes/combine_files/_m/CMC_phenotypes_all.csv').
↳set_index("RNAseq:Sample_RNA_ID")

@functools.lru_cache()
def get_res_pheno_df():
    return pd.merge(get_pheno_df(), get_res_df(), left_index=True,
↳right_index=True)

[3]: def get_de(feature):
    f = pd.read_csv('.././.././female_analysis/_m/%s/diffExpr_szVctl_full.txt' %
↳feature, sep='\t')\
        .rename(columns={'gene_id': 'gencodeID'})
    f['ensemblID'] = f.gencodeID.str.replace("\\.*", "")
    f.set_index('ensemblID', inplace=True)
    m = pd.read_csv('.././.././male_analysis/_m/%s/diffExpr_szVctl_full.txt' %
↳feature, sep='\t')\
        .rename(columns={'gene_id': 'gencodeID'})
    m['ensemblID'] = m.gencodeID.str.replace("\\.*", "")
    m.set_index('ensemblID', inplace=True)
```

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    a = pd.read_csv('/ceph/projects/v3_phase3_paper/inputs/cmc/_m/
    ↪CMC_MSSM-Penn-Pitt_DLPFC_mRNA_IlluminaHiSeq2500_gene-adjustedSVA-differentialExpression-inc
    ↪tsv', sep='\t')\
        .rename(columns={"MAPPED_genes": 'gene_name'}).set_index('genes')
    return f, m, a

def get_unique(x, y, thres=0.05):
    return x.merge(pd.DataFrame(index = list(set(x[(x['adj.P.Val'] <= thres)].
    ↪index) -
                                                set(y[(y['adj.P.Val'] <= thres)].
    ↪index)))),
                    left_index=True, right_index=True)

def subset_sz_male():
    df = get_res_pheno_df()
    ctl = df[(df['Dx'] == 'Control') & (df['Sex'] == 'XY')].copy()
    sz = df[(df['Dx'] == 'SCZ') & (df['Sex'] == 'XY')].copy()
    return ctl, sz

def add_pvals_adjustPval(df):
    ctl, sz = subset_sz_male()
    m_pval = []
    for gene_id in df.Feature:
        stat, pval = mannwhitneyu(ctl[gene_id], sz[gene_id])
        m_pval.append(pval)
    fdr_m = fdr correction(m_pval)
    return pd.concat([df.set_index('Feature'),
                      pd.DataFrame({'Male_Pval': m_pval,
                                    'Male_FDR': fdr_m[1]},
                                    index=df.Feature)], axis=1)

```

1.1 Genes

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[4]: f, m, a = get_de('genes')
f['Feature'] = f.gencodeID
f['ensemblID'] = f.index
#genes = get_unique(get_unique(f, m), a)
genes = get_unique(f, m).rename(columns={'chromosome_name': 'Chrom',
    ↪'hgnc_symbol': 'Symbol'})
genes = genes[['Feature', 'gencodeID', 'Symbol', 'ensemblID',
               'Chrom', 'logFC', 't', 'adj.P.Val']].sort_values('adj.P.Val')
genes.Chrom = 'chr'+genes.Chrom
genes = add_pvals_adjustPval(genes)
genes = genes[~(genes['Male_Pval'] <= 0.05)] ## Stringents

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genes['Type'] = 'gene'
genes.shape
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[4]: (583, 10)

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[5]: genes[(genes['t']>0)].head(5)
```

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[5]:
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	gencodeID	Symbol	ensemblID	Chrom	\
Feature					
ENSG00000153132.12	ENSG00000153132.12	CLGN	ENSG00000153132	chr4	
ENSG00000179083.6	ENSG00000179083.6	FAM133A	ENSG00000179083	chrX	
ENSG00000165733.7	ENSG00000165733.7	BMS1	ENSG00000165733	chr10	
ENSG00000183023.18	ENSG00000183023.18	SLC8A1	ENSG00000183023	chr2	
ENSG00000236268.5	ENSG00000236268.5	LINC01361	ENSG00000236268	chr1	

	logFC	t	adj.P.Val	Male_Pval	Male_FDR	Type
Feature						
ENSG00000153132.12	0.389937	5.559139	0.000123	0.283020	0.313711	gene
ENSG00000179083.6	0.261268	5.004488	0.000535	0.272019	0.302973	gene
ENSG00000165733.7	0.150918	4.986552	0.000535	0.169787	0.205428	gene
ENSG00000183023.18	0.245819	4.925477	0.000632	0.082759	0.108963	gene
ENSG00000236268.5	0.404532	4.865744	0.000700	0.110632	0.140404	gene

1.2 DE summary

1.2.1 DE (feature)

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[6]: gg = len(set(genes['gencodeID']))

print("\nGene:\t\t%d" % (gg))
```

Gene: 583

```
[7]: genes.to_csv('female_specific_DE_genes.txt', sep='\t', index=True, header=True)
```

1.3 Number of DEGs on allosomes

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[8]: genes[(genes['Chrom'].isin(['chrX', 'chrY']))].groupby(['Type', 'Chrom']).size()
```

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[8]: Type  Chrom
gene  chrX    21
dtype: int64
```

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[ ]:
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