Pathogenic posIDs- adding grantham hi low scores to aa level df that has window burden info. from mod4

10/23/22 saved originaally as 'FIGURE4_CpDAA_missenseLevelScore..' jnb

3/13/23 updated name, added more score grantham columns, duplicated this jnb to make pathogenic CKY pos speicifc version

input data:

dir:

/Users/mariapalafox/Desktop/BRIDGE/DISORDER/dbNSFPmapping/MAXMEAN_SCORING

 Originally downloaded filtered CpDAAoCATEGORY dbNSFP files from hoffman "sh # INPUT FOR THIS MARKDOWN & FIGURE 3 scp mfpalafo@hoffman2.idre.ucla.edu:/u/home/m/mfpalafo/projectarboleda/patho_dbNSFP/MAXMEAN_SCORING/top10scores_MetaMerged_1323_MendelCpD
 .

Input files made in:

FIGURE_3c_missenseLevelScoresPt3_CpDAAoCATEGORY_MendelCpD.ipynb (pathogenic poslDs)

Input file names for this markdown:

top10scores_Patho_positions_granthamHighOrLowCorrectedVersion_CKYonly_850_r

- highest or lowest grantham missense scores for pathogenic posID filtered data from OMIMCpD protein subset
- could not make for OMIM protein subset becauses dont have all possible missense scores for every posID in OMIM, onnly downloaded OMIMCpD metamerge

output data:

dir:

/Users/mariapalafox/Desktop/BRIDGE/disorder/FIGURES/modules/STEP4/grantham/

```
In [13]: # packages
         import os
         import sys
         import numpy as np
         import pandas as pd
         import csv
         from ast import literal eval
         from statistics import mean
         %matplotlib inline
         sys.path.append("/Users/mariapalafox/Desktop/TOOLBOXPY")
         from all funx import *
         from IPython.display import display, HTML
         from IPython.display import Image
         display(HTML("<style>.container {width:90% !important;}</styl</pre>
         e>"))
         pd.set option('display.max columns', None)
         pd.set option('display.max colwidth', 1000) # you can't use n
         one here
         pd.set option('display.max seg items', 2000) # seg in column
         pd.set option('display.max rows', 10)
         %timeit
         # no scientific notation display, format decimal
         pd.options.display.float format = '{:.4f}'.format
         def mediumpic(name):
             display(Image(filename = name, width= 500, height=650))
         def smallpic(name):
             display(Image(filename = name, width= 300, height=600))
         def largepic(name):
             display(Image(filename = name, width= 800, height=1000))
```

filtered input data for CpDAA missense changes corresponding to the highest and lowest grantham score

- paper 1 si for K missense type odds ratio has different order than current project grantham reference excel MISSENSE/GRANTHAM_CpDAA_MissenseTypes.csv
- double check word doc and excel with other sources

highest grantham change:

C/W K/I or K/M Y/C

lowest grantham change:

C/S

K/R

Y/F

- Y missense types all = 2331, single codon
- C/S is most of any C missense type, with 14434 and other types at 7217
- K/R is similar to other missense type counts at 8908, but K/I and K/M are possible by only 1 of 2 codons, so these counts combined = 8908, why is K/N so high?

<u>Grantham conservative range described below from this link</u> (https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3431198/)

title: In silico analysis of missense substitutions using sequence-alignment based methods, 2008

OVERVIEW OF IN SILICO APPROACHES TO MISSENSE SUBSTITUTION ANALYSIS

Fundamentally, there are four classes of amino acid, sequence, or structural attribute that have been used to try to distinguish between neutral and pathogenic missense substitutions in silico: (1) pairwise comparison of the physico-chemical characteristics or evolutionary substitution frequencies between the wild-type and variant amino acid, (2) evolutionary conservation at the position at which a missense substitution is observed, (3) comparison between the variant amino acid and the evolutionarily tolerated amino acid range of variation at its position in the protein, and (4) protein structural considerations. In silico missense analysis algorithms may use data from just one of these classes, or combine data from two, three, or all four of the classes.

Pairwise amino acid comparisons may be based on data from amino acid substitution scoring matrices (e.g., PAM250, BLOSUM62). These matrices were derived from the frequencies with which the 20 amino acids are observed to substitute for each other in multiple sequence alignments of related proteins [Dayhoff et al., 1978; Henikoff and Henikoff, 1992]. BLOSUM62 scores range from 4–11 for identities, from 0–3 for commonly observed substitutions, and from (–1) to (–4) for substitutions rarely observed in related proteins. The average BLOSUM62 score is

lower for pathogenic substitutions than for neutral substitutions [Ferrer-Costa et al., 2002; Balasubramanian et al., 2005]. Alternatively, comparisons may be based on amino acid physical or chemical properties.

A score called the Grantham Difference describes the difference in side chain atomic composition, polarity, and volume between two amino acids [Grantham, 1974]. Substitutions with Grantham Differences of:

- 5-60 are generally considered "conservative",
- 60-100 "non-conservative", and
- > 100 "radical"

The average Grantham Difference for pathogenic substitutions is higher than for neutral substitutions [Miller et al., 2001; Abkevich et al., 2004; Balasubramanian et al., 2005]. However, these methods of pairwise amino acid comparisons alone have not led to popular missense substitution classification algorithms.

Grantham's distance [edit]

Grantham's distance depends on three properties: composition, polarity and molecular volume.^[4]

Distance difference D for each pair of amino acid i and j is calculated as:

$$D_{ij} = \left[lpha (c_i - c_j)^2 + eta (p_i - p_j)^2 + \gamma (v_i - v_j)^2
ight]^{rac{1}{2}}$$

where c = composition, p = polarity, and v = molecular volume; and are constants of squares of the inverses of the mean for each property, respectively equal to 1.833, 0.1018, 0.000399. According to Grantham's distance, most similar amino ϵ leucine and isoleucine and the most distant are cysteine and tryptophan.

Difference D for amino acids[4]

Arg	Leu	Pro	Thr	Ala	Val	Gly	lle	Phe	Tyr	Cys	His	Gln	Asn	Lys	Asp	Glu	Met	Trp	
110	145	74	58	99	124	56	142	155	144	112	89	68	46	121	65	80	135	177	Se
	102	103	71	112	96	125	97	97	77	180	29	43	86	26	96	54	91	101	Arg
		98	92	96	32	138	5	22	36	198	99	113	153	107	172	138	15	61	Le
			38	27	68	42	95	114	110	169	77	76	91	103	108	93	87	147	Pr
				58	69	59	89	103	92	149	47	42	65	78	85	65	81	128	Th
					64	60	94	113	112	195	86	91	111	106	126	107	84	148	Al
						109	29	50	55	192	84	96	133	97	152	121	21	88	Va
							135	153	147	159	98	87	80	127	94	98	127	184	GI
								21	33	198	94	109	149	102	168	134	10	61	lle
									22	205	100	116	158	102	177	140	28	40	Ph
										194	83	99	143	85	160	122	36	37	Ту
											174	154	139	202	154	170	196	215	Су
												24	68	32	81	40	87	115	Hi
													46	53	61	29	101	130	GI
														94	23	42	142	174	As
															101	56	95	110	Ly
																45	160	181	As
																	126	152	GI
																		67	Me

Wiki grantham difference table results

- cysteine change differences are all considered radical >100 substitutions
- lysine and tyrosine have more conservative possible substitutions

--- missense Grantham -----

7	*K/R	26
6	K/Q	53
5	K/E	56
4	K/T	78
3	K/N	94
2	*K/M	95
1	*K/I	102
6	*C/S	112
5	C/G	159
4	C/R	180
3	C/Y	194
2	C/F	205
1	*C/W	215
6	*Y/F	22
5	Y/H	83
4	Y/N	143
3	Y/S	144
2	Y/D	160
1	*Y/C	194

Grantham difference article blurb at top of markdown:

- 5-60 are generally considered "conservative",
- 60-100 "non-conservative", and
- > 100 "radical"

[1] PATHO positions in OMIM&CpD proteins

```
uniqueCount(patho, 'posID')
uniqueCount2(patho, 'UKBID')
# only grantham change missense counts for category overlap o
f CpDAA codons shown in counts
checkColumnValue(patho, 'CATEGORY')
#making nonredundant posID version of df
dfposID = patho.drop duplicates(['posID'])
print( ' ----- unique PATHO CKY in 415 proteins -----
uniqueCount(dfposID, 'posID')
print()
print('----PATHO type counts in 415 proteins----
----')
checkColumnValue(dfposID, 'aaref')
print()
print('-----')
checkColumnValue(dfposID, 'LP')
crosstabit(dfposID, 'aaref', 'LP')
print()
print('-----')
checkColumnValue(dfposID, 'RP')
crosstabit(dfposID, 'aaref', 'RP')
print()
print('why are low and high grantham counts not the same, eac
h posID should have 1 high and 1 low change..')
checkColumnValue(patho, 'missenseType')
# making seperate df's for highest grantham difference and lo
west grantham differences between aaref and aaalt in misse
nse exchange
print('----')
checkColumnValue(patho, 'Grantham.highest') # df.dtypes == b
001
gHigh = patho[patho['Grantham.highest']].copy()
print('high grantham df shape: ', gHigh.shape)
checkColumnValue(gHigh, 'missenseType')
uniqueCount(gHigh, 'posID')
uniqueCount2(gHigh, 'UKBID')
print()
print('----')
checkColumnValue(patho, 'Grantham.lowest') # df.dtypes == bo
gLow = patho[patho['Grantham.lowest']].copy()
```

```
print('low grantham df shape: ', gLow.shape)
checkColumnValue(gLow, 'missenseType')
uniqueCount(gLow, 'posID')
uniqueCount2(gLow, 'UKBID')
print('----')
kr = gLow[gLow['missenseType'] == 'K/R']
checkColumnValue(kr, 'codonpos')
yf = gLow[gLow['missenseType'] == 'Y/F']
checkColumnValue(yf, 'codonpos')
posID length: 1986
posID set length: 850
UKBID set length: 415
     CATEGORY Count
         NaN 1597
1 PATHOGENIC 314
    RAREST
2
                45
3
          VUS
                21
4 RARE 8
5 COMMON/BENIGN 1
----- unique PATHO CKY in 415 proteins ------
posID length: 850
posID set length: 850
-----PATHO type counts in 415 proteins-----
aaref Count
0 Y 320
1 C 286
2 K 244
-----total LP counts-----
        LP Count
        NaN 786
1 Unliganded 56
2 Liganded 8
LP Liganded Unliganded Total
aaref
                   34 41
         7
С
K
                   22 23
           1
                   56 64
Total
      8
```

-----total RP counts-----

```
RP Count
0
    NaN
         792
1
           42
    Low
2 Medium
           11
3
    High
         5
RP
     High Low Medium Total
aaref
С
       4 13
                       24
                 7
K
        1 14
                       19
                 4
        0 15
Y
                 0
                       15
Total
       5 42
                  11
                       58
why are low and high grantham counts not the same, each posI
D should have 1 high and 1 low change..
 missenseType Count
0
       C/S
              572
        Y/F
1
             320
2
        Y/C 320
       C/W 286
3
4
        K/R 244
5
       K/M 151
        K/I
              93
_____
  Grantham.highest Count
0
           False 1136
1
            True
                850
high grantham df shape: (850, 53)
 missenseType Count
0
        Y/C
              320
1
       C/W
             286
2
        K/M 151
3
        K/I 93
posID length: 850
posID set length: 850
UKBID set length: 415
  Grantham.lowest Count
0
    True 1136
1
         False 850
low grantham df shape: (1136, 53)
 missenseType Count
0
       C/S
              572
1
        Y/F
              320
```

```
2 K/R 244

posID length: 1136
posID set length: 850

UKBID set length: 415

codonpos Count
0 2 244

codonpos Count
0 2 320
```

[2] PATHO positions in OMIM&CpD proteins

```
In [16]: # duplicate cysteine posID issue
    cs = gLow[gLow['missenseType'] == 'C/S']
    checkColumnValue(cs, 'codonpos')
    checkColumnValue(cs, 'refcodon')

    cs.groupby('codonpos').agg(
        mean_CADD_phred =('CADD_phred', mean),
        mean_phyloP100way_vertebrate =('phyloP100way_vertebrate',
        mean)
    )
        # first position of codon less constrained than 2nd pos typ
        ically
```

Out[16]:

$mean_CADD_phred \quad mean_phyloP100way_vertebrate$

codonpos		
1	26.3442	7.4606
2	25.9940	7.7536

[3] PATHO positions in OMIM&CpD proteins

```
In [18]: # Position 1 slightly higher cadd and less connserved than po
        s 2
        # going to count the C/S change thats at codon position 1
        # arg for codonn pos 2, matches lysine and tyrosine substitut
        ion
        cs = cs[cs['codonpos'] == 1]
        #print(cs.shape)
        print('----high grantham-----
        ---')
        uniqueCount(gHigh, 'posID')
        checkColumnValue(gHigh, 'missenseType')
        print('high grantham K missense 151 + 93: ', 151 + 93, '\n')
        checkColumnValue(gHigh, 'codonpos')
        print('----')
        print('----')
        gLow = pd.concat([cs, kr, yf], axis=0, ignore index=True)
        uniqueCount(gLow, 'posID')
        checkColumnValue(gLow, 'missenseType')
        checkColumnValue(gLow, 'codonpos')
        print('-----
        checkColumnValue(gHigh, 'aaref')
        checkColumnValue(gLow, 'aaref')
        # check for exact count match
```

```
-----high grantham-----
posID length: 850
posID set length: 850
 missenseType Count
    Y/C
             320
1
       C/W
            286
       K/M 151
K/I 93
2
3
high grantham K missense 151 + 93: 244
  codonpos Count
0
           564
      2
1
       3
          286
_____
----- low grantham-----
posID length: 850
posID set length: 850
 missenseType Count
    Y/F
0
             320
       C/S 286
1
2
       K/R 244
  codonpos Count
     2
0
          564
1
       1
          286
 aaref Count
0 Y 320
1
   C 286
2 K
       244
```

aaref Count

С

2 K 244

Y 320

286

0

1

[4] PATHO positions in OMIM&CpD proteins

10 raw scores:

 8 were used to make mean and max raw/rankscore concordaance (MPC and phylop100way added later)

chose raw scores over rankscore for clustering input

```
In [19]: # renaming grantham highessts and lowesst misesnse change df
         columns
         gLow = gLow[['posID', 'missenseType', 'codonType',
                       'VEST4 score',
                          'REVEL score',
                          'MutPred score',
                          'MVP score',
                          'BayesDel addAF score',
                          'ClinPred score',
                          'LISTS2 score',
                          'CADD raw',
                          'MPC score',
                          'phyloP100way vertebrate']].copy()
         lowColumns = ['posID', 'lowestGrantham.missenseType', 'lowe
         stGrantham.codonType',
                          'lowestGrantham.VEST4 score',
                          'lowestGrantham.REVEL score',
                          'lowestGrantham.MutPred score',
                          'lowestGrantham.MVP score',
                          'lowestGrantham.BayesDel addAF score',
                          'lowestGrantham.ClinPred score',
                          'lowestGrantham.LISTS2 score',
                          'lowestGrantham.CADD raw',
                          'lowestGrantham.MPC score',
                          'lowestGrantham.phyloP100way vertebrate']
         # rename low granthaam
         gLow.columns = lowColumns
         gHigh = gHigh[['posID', 'missenseType', 'codonType',
                         'VEST4 score',
                          'REVEL score',
                          'MutPred score',
                          'MVP_score',
                          'BayesDel addAF score',
                          'ClinPred score',
                          'LISTS2 score',
                          'CADD raw',
                          'MPC score',
```

[5] PATHO positions in OMIM&CpD proteins

Importing window missense burden AA level df from module processing

```
In [20]: dir4 = "/Users/mariapalafox/Desktop/BRIDGE/disorder/FIGURES/m
         odules/STEP4/"
         os.chdir(dir4)
         # CHANGE
         burden = pd.read csv("CKY level rmd Background 1D 3Dcounts ad
         ded 1234 MendelCpD v2.csv", low memory=False)
         print('filtering for only positions in high and low grantham
         dfs')
         # CHANGE
         # burden = burden[burden['CpDAA'] == 1].copy() # not necessaa
         ry because small omimcpd dataframe and merging
         print(burden.shape, '\n')
         checkColumnValue(burden, 'aaref')
         uniqueCount2(burden, 'UKBID')
         uniqueCount(burden, 'posID')
         print('NOTE: all aa position rows are unique in burden df')
         print('-----
         ----')
         print('merging grantham hi low dfs with aa-level window burde
         n df')
         final = pd.merge(burden, qHigh, on=['posID'], how='inner')
         print(gHigh.shape)
         print(final.shape)
         final = pd.merge(final, gLow, on=['posID'], how='inner')
         print(gLow.shape)
         print(final.shape)
```

```
filtering for only positions in high and low grantham dfs
         (101845, 134)
           aaref Count
               K 60312
         1
               Y 25112
               C 16421
         UKBID set length: 1234
         posID length: 101845
         posID set length: 101845
         NOTE: all aa position rows are unique in burden df
         merging grantham hi low dfs with aa-level window burden df
         (850, 13)
         (793, 146)
         (850, 13)
         (793, 158)
In [21]: checkColumnValue(final, 'aaref')
         uniqueCount2(final, 'UKBID')
         uniqueCount(final, 'posID')
           aaref Count
               Y
                   304
               С
         1
                    260
                    229
         UKBID set length: 389
         posID length: 793
         posID set length: 793
```

[6] SAVING PATHO only AA level with grantham hi low predictions from 10 scores for

1. OMIM&CpD proteins n = 389 proteins, 793 CKY pathogenic positions

```
In [22]: dir4grantham = "/Users/mariapalafox/Desktop/BRIDGE/disorder/F
         IGURES/modules/STEP4/grantham/"
         os.chdir(dir4grantham)
         # accounting all cpdaa
         uniqueCount(final, 'posID')
         uniqueCount2(final, 'UKBID')
         checkColumnValue(final, 'aaref')
         crosstabit(final, 'aaref', 'RP')
         crosstabit(final, 'aaref', 'LP')
         final.to_csv("CKY_level_PATHO_HighLow_granthamScored_rmd_Back
         ground_1D_3Dcounts_included_389_OMIMCpD_793_posIDs_v3.csv", i
         ndex=False)
         posID length: 793
        posID set length: 793
        UKBID set length: 389
          aaref Count
         0
              Y
                   304
         1
              С
                   260
         2
              K
                   229
        RP High Low Medium Total
         aaref
                  4 13
                               7
                                     24
        С
        K
                  1 14
                               4
                                     19
        Y
                  0 15
                               0
                                     15
         Total 5 42
                              11
                                     58
        _{
m LP}
               Liganded Unliganded Total
         aaref
        С
                      7
                                 34
                                        41
         K
                      1
                                 22
                                        23
         Total
                      8
                                 56
                                        64
```

```
In [23]: final.head()
```

Out[23]:

	posID	genename	UKBID	CpD_UKBID	Mendelian_UKBID	pos	aaref	
0	A6NHR9_Y283	SMCHD1	A6NHR9	True	True	283	Υ	
1	O00329_C416	PIK3CD	O00329	True	True	416	С	
2	O00429_C367	DNM1L	O00429	True	True	367	С	
3	O00429_C431	DNM1L	O00429	True	True	431	С	
4	O00429_C446	DNM1L	O00429	True	True	446	С	

In [25]: final.columns

P rankscore.mean',

```
Out[25]: Index(['posID', 'genename', 'UKBID', 'CpD_UKBID', 'Mendelian
         UKBID', 'pos',
                 'aaref', 'refcodon', 'CpDAA', 'LP', 'RP', 'R', 'PATHO
         GENIC',
                'COMMONBENIGN', 'VUS', 'RARE', 'RAREST', 'BACKGROUN
         D',
                'Interpro domain', 'MPC score.mean', 'MPC score.max',
                'MPC rankscore.mean', 'MPC rankscore.max',
                 'phyloP100way vertebrate.mean', 'phyloP100way vertebr
         ate.max',
                 'phyloP100way vertebrate rankscore.mean',
                 'phyloP100way vertebrate rankscore.max', 'MAX.RANKSCO
         RE.SUM',
                 'MEAN.RANKSCORE.SUM', 'MAX.RAWSCORE.SUM', 'MEAN.RAWSC
         ORE.SUM',
                'VEST4 rankscore.max', 'REVEL rankscore.max', 'MutPre
         d rankscore.max',
                'MVP_rankscore.max', 'BayesDel_addAF_rankscore.max',
                'ClinPred rankscore.max', 'LISTS2 rankscore.max',
                'CADD_raw_rankscore.max', 'binary.VEST4.max', 'binar
         y.REVEL.max',
                 'binary.MutPred.max', 'binary.MVP.max', 'binary.Bayes
         Del.max',
                 'binary.ClinPred.max', 'binary.LIST.max', 'binary.CAD
         Dp.max',
                'VEST4 score.max', 'REVEL score.max', 'MutPred score.
         max',
                'MVP score.max', 'BayesDel addAF score.max', 'ClinPre
         d score.max',
                 'LISTS2 score.max', 'CADD phred.max', 'VEST4 rankscor
```

'REVEL rankscore.mean', 'MutPred rankscore.mean', 'MV

```
'BayesDel addAF rankscore.mean', 'ClinPred rankscore.
mean',
       'LISTS2 rankscore.mean', 'CADD raw rankscore.mean', '
binary.VEST4.mean',
       'binary.REVEL.mean', 'binary.MutPred.mean', 'binary.M
VP.mean',
       'binary.BayesDel.mean', 'binary.ClinPred.mean', 'bina
ry.LIST.mean',
       'binary.CADDp.mean', 'VEST4 score.mean', 'REVEL scor
e.mean',
       'MutPred score.mean', 'MVP score.mean', 'BayesDel add
AF score.mean',
       'ClinPred score.mean', 'LISTS2 score.mean', 'CADD phr
ed.mean',
       'UKBID VUSandPATHO', 'UKBID BACKGROUNDandPATHO', 'UKB
ID BENIGNandPATHO',
       'PATHOGENIC.withinRange0.count', 'PATHOGENIC.withinRa
nge3.count',
       'PATHOGENIC.withinRange6.count', 'PATHOGENIC.withinRa
nge15.count',
       'COMMONBENIGN.withinRange0.count', 'COMMONBENIGN.with
inRange3.count',
       'COMMONBENIGN.withinRange6.count', 'COMMONBENIGN.with
inRange15.count',
       'VUS.withinRange0.count', 'VUS.withinRange3.count',
       'VUS.withinRange6.count', 'VUS.withinRange15.count',
       'RARE.withinRange0.count', 'RARE.withinRange3.count',
       'RARE.withinRange6.count', 'RARE.withinRange15.coun
t',
       'RAREST.withinRange0.count', 'RAREST.withinRange3.cou
nt',
       'RAREST.withinRange6.count', 'RAREST.withinRange15.co
unt', 'Length',
       'posID.norm', 'posID.decile', 'PATHOGENIC.within0A.co
unt',
       'PATHOGENIC.within6A.count', 'PATHOGENIC.within8A.cou
nt',
       'PATHOGENIC.within10A.count', 'COMMONBENIGN.within0A.
count',
       'COMMONBENIGN.within6A.count', 'COMMONBENIGN.within8
A.count',
       'COMMONBENIGN.within10A.count', 'VUS.within0A.count',
       'VUS.within6A.count', 'VUS.within8A.count', 'VUS.with
in10A.count',
       'RARE.within0A.count', 'RARE.within6A.count', 'RARE.w
ithin8A.count',
       'RARE.within10A.count', 'RAREST.within0A.count',
       'RAREST.within6A.count', 'RAREST.within8A.count',
       'RAREST.within10A.count', 'BACKGROUND.withinRange0.co
unt',
       'BACKGROUND.withinRange3.count', 'BACKGROUND.withinRa
nge6.count',
```

```
'BACKGROUND.withinRange15.count', 'BACKGROUND.within0
A.count',
       'BACKGROUND.within6A.count', 'BACKGROUND.within8A.cou
nt',
       'BACKGROUND.within10A.count', 'resolved3D.posID',
       'highestGrantham.missenseType', 'highestGrantham.codo
nType',
       'highestGrantham.VEST4 score', 'highestGrantham.REVEL
_score',
       'highestGrantham.MutPred_score', 'highestGrantham.MVP
score',
       'highestGrantham.BayesDel addAF score',
       'highestGrantham.ClinPred score', 'highestGrantham.LI
STS2 score',
       'highestGrantham.CADD raw', 'highestGrantham.MPC scor
e',
       'highestGrantham.phyloP100way_vertebrate',
       'lowestGrantham.missenseType', 'lowestGrantham.codonT
ype',
       'lowestGrantham.VEST4 score', 'lowestGrantham.REVEL s
core',
       'lowestGrantham.MutPred score', 'lowestGrantham.MVP s
core',
       'lowestGrantham.BayesDel addAF score', 'lowestGrantha
m.ClinPred score',
       'lowestGrantham.LISTS2 score', 'lowestGrantham.CADD r
aw',
       'lowestGrantham.MPC score', 'lowestGrantham.phyloP100
way vertebrate'],
      dtype='object')
```

In []: # add describeprot data to this df for rmd clustering

"cluster1D window3 mean RAWSCORE droppedOverlapBtw Patho Beni qn CpDAA CKYfiltered Mendelian.csv" cluster1D window3 MeanRAWSCOREsum Pathogenic29324 Mendelian.c sv