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SUPPLEMENTARY INFORMATION

Sample code. Python 2.7

Task[1] Compute amino acid entropy (**H**).
Task[2] Sort for amino acid positions where **H**>0 in human but not in mosquito
Task[3] Compute cumulative mutual information (**cMI_{xx}**) within the Exclusive human subset (**H_x**)

Task[1] Compute Shannon entropy (H) in bits. Input probability of occurrence of each amino acid.

a, c, d, e, f, g, h, i, k, l, m, n, p, q, r, s, t, v, w, y = amino acid probabilities

from cmath import *

def main(a, c, d, e, f, g, h, i, k, l, m, n, p, q, r, s, t, v, w, y):

 if a==0:
 Ha=0
 else: Ha=a*log(1/a,2)

 if c==0:
 Hc=0
 else: Hc=c*log(1/c,2)

 if d==0:
 Hd=0
 else: Hd=d*log(1/d,2)

 if e==0:
 He=0
 else: He=e*log(1/e,2)

 if f==0:
 Hf=0
 else: Hf=f*log(1/f,2)

 if g==0:
 Hg=0
 else: Hg=g*log(1/g,2)

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if h==0:  
    Hh=0  
else: Hh=h*log(1/h,2)
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if i==0:  
    Hi=0  
else: Hi=i*log(1/i,2)
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if k==0:  
    Hk=0  
else: Hk=k*log(1/k,2)
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if l==0:  
    Hl=0  
else: Hl=l*log(1/l,2)
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if m==0:  
    Hm=0  
else: Hm=m*log(1/m,2)
```

```
if n==0:  
    Hn=0  
else: Hn=n*log(1/n,2)
```

```
if p==0:  
    Hp=0  
else: Hp=p*log(1/p,2)
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```
if q==0:  
    Hq=0  
else: Hq=q*log(1/q,2)
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```
if r==0:  
    Hr=0  
else: Hr=r*log(1/r,2)
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if s==0:  
    Hs=0  
else: Hs=s*log(1/s,2)
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if t==0:  
    Ht=0  
else: Ht=t*log(1/t,2)
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if v==0:  
    Hv=0
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else: Hv=v*log(1/v,2)

if w==0:
    Hw=0
else: Hw=w*log(1/w,2)

if y==0:
    Hy=0
else: Hy=y*log(1/y,2)

H=Ha+Hc+Hd+He+Hf+Hg+Hh+Hi+Hk+Hl+Hm+Hn+Hp+Hq+Hr+Hs+Ht+Hv+Hw+Hy

H=abs(H)

return H

# Task[2] sort for positions where H>0 in human origin polyproteins and H=0 in Aedes
origin polyproteins
# Hx = exclusive human subset with H>0; n=3423 polyprotein amino acid positions

Hx=zeros(n)
for ndx2 in range(0, n, 1):
    x1 = H_human[ndx2]
    x2 = H_aedes[ndx2]
    if x1 > 0.0:
        if x2 == 0.0:
            Hx[ndx2] = x1

# Task[3] compute cumulative MI (cMIxx) in paired members of Human Exclusive
Subset

cMIxx = zeros(n)
for ndx1 in range(0, n, 1):
    if Hx[ndx1] > 0.0:
        seqx1 = make_seq_human(ndx1)
        seqx1 = seqx1.replace('X', '-') # 'X' represents gaps
        for ndx2 in range(0, n, 1):
            if Hx[ndx2] > 0.0:
                seqx2 = make_seq_human(ndx2)
                seqx2 = seqx2.replace('X', '-') # 'X' represents gaps
                pairs = zip(seqx1, seqx2)
                list_out2=[]
                M = zeros((20, 20))
                M = matrix(M, float)
                for ndx3 in range(0, 20, 1):
                    x1 = aa_list1[ndx3]

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for ndx4 in range(0, 20, 1):
    x2 = aa_list2[ndx4]
    ref_pair = (x1, x2)
    ref_pair2 = aa_list1[ndx3], aa_list2[ndx4]
    if ref_pair in pairs:
        cx=pairs.count(ref_pair)
        M[ndx3, ndx4] = float(cx)

sum_M =sum(M)
probs = M/sum_M
M2 = zeros((20, 20))
M2 = matrix(M, float)
for ndx5 in range(0, 20, 1):
    for ndx6 in range(0, 20, 1):
        x = probs[ndx5, ndx6]
        if x > 0.0:
            Hjoint = -x*log2(x)
            M2[ndx5, ndx6] = Hjoint
MI = Hx[ndx1] + Hx[ndx2] - sum(M2)
if MI>0.0:
    if ndx1 != ndx2: # correction for autocorrelation
        cMIxx[ndx1] = cMIxx[ndx1] + MI

```