

Sequence alignment

SEQ1 :

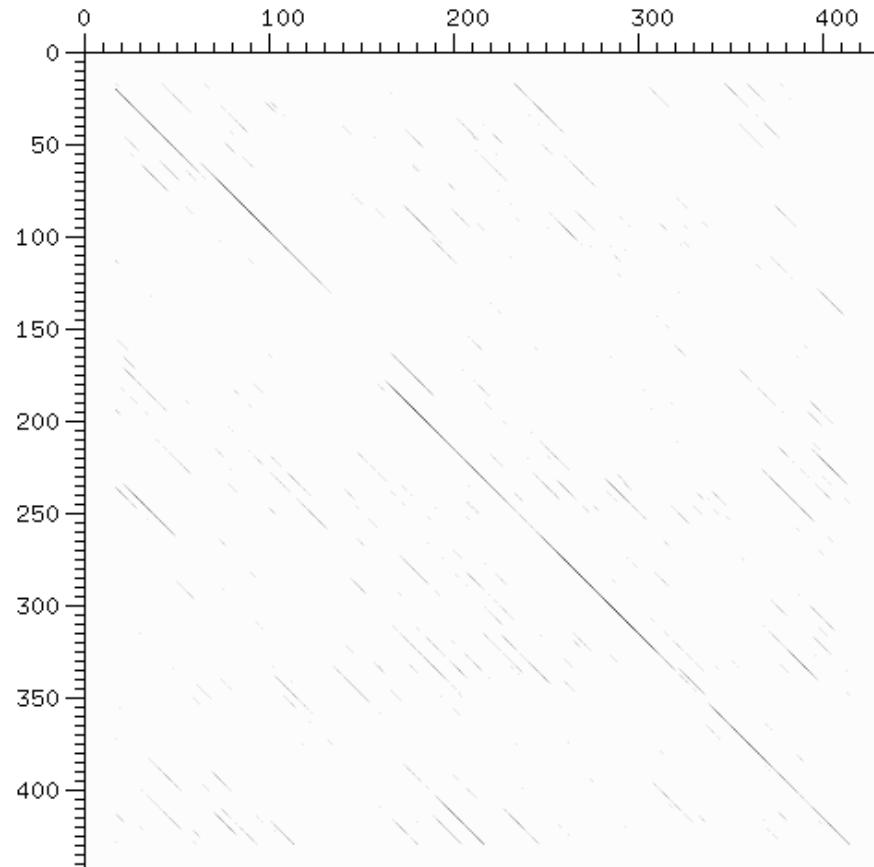
VLSPADKTNVKAAGKVGAAHAGEYGAEALEMFLSFPTTKTYFPHFDSLHGSAQVKGHGKK
VADALTNAVAHVDDPNALSALSDLHAHKLRVPVNFKLLSHCLLVTAAHLPAEFTPAVHA
SLDKFLASVSTVLTSKYR

SEQ2 :

VHLTPEEKSAVTALWGKVNVDEVGGEALGRLLVVYPWTQRFFESFGDLSTPDAVMGNPKVK
AHGKKVLGAFSDGLAHLDNLKGTATLSELHCDKLHVDPENFRLLGNVLVCVLAHHFGKEF
TPPVQAAYQKVVAGVANALAHKYH

Are these two proteins alike?

Dotplots: visual sequence comparison



1. Place two sequences along axes of plot
2. Place dot at grid points where two sequences have identical residues
3. Diagonals correspond to conserved regions

Alignment scores: match vs. mismatch

Simple scoring scheme (too simple in fact...):

Matching amino acids: 5

Mismatch: 0

Scoring example:

K	A	W	S	A	D	V
:	:	:	:	:		
<u>K</u>	D	W	S	A	E	V

5+0+5+5+5+0+5 = 25

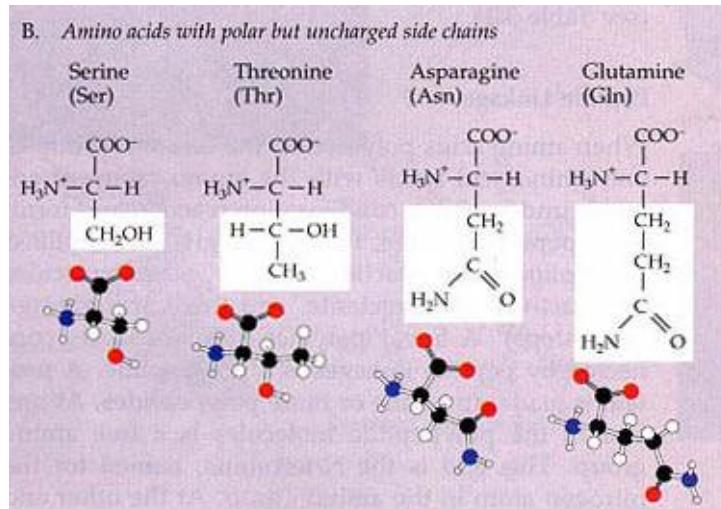
Pairwise alignments: conservative substitutions

43.2% identity;

Global alignment score: 374

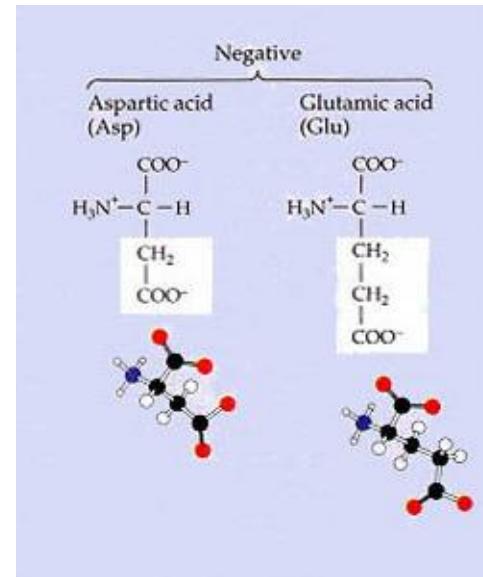
	10	20	30	40	50	
alpha	V-LSPADKTNVKAAGKVGAGHAGEYGAELERMFLSFPTTKTYFPHF-DLS-----HGSA					
	: : : : : . . . : : . . . : . . . : : : : . . .					
beta	VHLTPEEKSAVTALWGKV--NVDEVGGEALGRLLVVYPWTQRFESFGDLSTPDAMGNP					
	10	20	30	40	50	
	60	70	80	90	100	110
alpha	QVKGHGKKVADALTNAVAHVDDMPNALSALSDLHAHKLRDPVNFKLLSHCLLVTIAAHL					
 : : : : . . . : : : . . .					
beta	KVKAHGKKVLGAFSDGLAHLDNLKGTFATLSELHCDKLHVDPENFRLLGNVLVCVLAHHF					
	60	70	80	90	100	110
	120	130	140			
alpha	PAEFTPAPVHASLDKFLASVSTVLTSKYR					
	. . . : : : . . .					
beta	GKEFTPPVQAAYQKVVAGVANALAHKYH					
	120	130	140			

Amino acid properties



Serine (S) and Threonine (T) have similar physicochemical properties

- => Substitution of S/T or E/D occurs relatively often during evolution
- => Substitution of S/T or E/D should result in scores that are only moderately lower than identities



Aspartic acid (D) and Glutamic acid (E) have similar properties

Protein substitution matrices

A	5																				
R	-2	7																			
N	-1	-1	7																		
D	-2	-2	2	8																	
C	-1	-4	-2	-4	13																
Q	-1	1	0	0	-3	7															
E	-1	0	0	2	-3	2	6														
G	0	-3	0	-1	-3	-2	-3	8													
H	-2	0	1	-1	-3	1	0	-2	10												
I	-1	-4	-3	-4	-2	-3	-4	-4	-4	5											
L	-2	-3	-4	-4	-2	-2	-3	-4	-3	2	5										
K	-1	3	0	-1	-3	2	1	-2	0	-3	-3	6									
M	-1	-2	-2	-4	-2	0	-2	-3	-1	2	3	-2	7								
F	-3	-3	-4	-5	-2	-4	-3	-4	-1	0	1	-4	0	8							
P	-1	-3	-2	-1	-4	-1	-1	-2	-2	-3	-4	-1	-3	-4	10						
S	1	-1	1	0	-1	0	-1	0	-1	-3	-3	0	-2	-3	-1	5					
T	0	-1	0	-1	-1	-1	-2	-2	-1	-1	-1	-1	-2	-1	2	5					
W	-3	-3	-4	-5	-5	-1	-3	-3	-3	-2	-3	-1	1	-4	-4	-3	15				
Y	-2	-1	-2	-3	-3	-1	-2	-3	2	-1	-1	-2	0	4	-3	-2	-2	2	8		
V	0	-3	-3	-4	-1	-3	-3	-4	-4	4	1	-3	1	-1	-3	-2	0	-3	-1	5	
	A	R	N	D	C	Q	E	G	H	I	L	K	M	F	P	S	T	W	Y	V	

When Blast works!

