

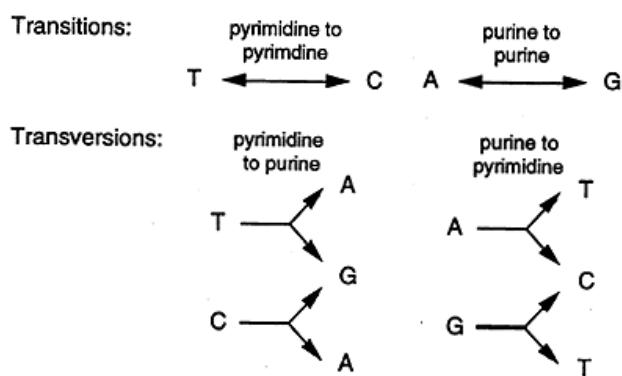
## Hands-On Four Multiple Sequence Alignment

The figures on the next page represent part of the alignment of DNA and protein sequences of the BRCA1 gene of different organisms. The top figures give the DNA sequence alignment, while the bottom figure gives the alignment of the BRCA1 protein sequences for the same region of the gene.

- 1) Examine more carefully the first twelve columns of the DNA sequence alignment and the first 4 columns of the protein sequence alignment. What can we conclude about the kind of substitutions that have occurred in the DNA sequences? Have they had any effect on the first 5 amino acids of Figure 2?
- 2) Examine more carefully the columns of the DNA sequence alignment. Are there more transitions or transversions?

**Transition:** this occurs when a purine is substituted with another purine or when a pyrimidine is substituted with another pyrimidine.

**Transversion:** when a purine is substituted for a pyrimidine or a pyrimidine replaces a purine.



- 3) Examine more carefully columns 8 and 12 in the amino acid sequence alignment. What can you conclude from the substitutions in each column?
- 4) Examine more carefully the gaps that are found in both figures.
  - a) Can you explain in your own words the gap produced in columns 38 to 46 in the DNA sequence alignment figure (or equivalently, columns 13, 14 and 15 in the protein sequence figure)?
  - b) Can you explain in your own words the gap produced in columns 108 to 113 in the DNA sequence alignment figure (or equivalently, columns 37 and 38 in the protein sequence figure)?

## Multiple Sequence Alignment

	10	20	30	*	*	*	*	*	*		
Wombat	:	AAAGTTAATGACTGGTTATCCACAAGTAGTGACATTTCAGCTCTGATA	ACTC	AAACGGTACCA	CCAT	GACGCCAG	CCAG	GCAG	CCAG	:	83
Opossum	:	AAAGTTAATGACTGGTTATTCAAGAAGTAATGACGTTTAGCCCCAGATTACTCAAGTGT	TAG	CACCATGAACAGA	ATGCAGA					:	83
Armadillo	:	AAAGTTAACGAGTGGTTTCCAGAGGTGATGACATATTAACTTCTGATGACTCACACGATAGGGGGTCTGAAT	AAATGCAGA							:	83
Sloth	:	AAAGTTAATGAGTGGTTTCCAGAAGTGATGACATACTAACTTCTGATGACTCACACAATGGGGGTCTGAATCAAATGCAGA								:	83
Dugong	:	AAAGTTAATGAGTGGTTTCAAGAAGTGATGACCTG-----GATGACTTGCGATGATAAGGGGTCTGAGTC	AAATGCAGA							:	74
Hyrax	:	AAAGTTAATGAGTGGTTTCCAGAAGTGACACCTA-----AGTGAATTTCACCTAG	GAGGGG	TCTGAAT	AAATGGAAA					:	74
Aardvark	:	AAAGTTAATGAGTGGTTTCCAGAAGTGATGCCG-----GATGGCTCACATGATGAAGGGTCTGAATCAAATGCAGA								:	74
Tenrec	:	AAAGTTAACGAGTGGTTTCCAAAGCCACGCCG-----GGTGCATTCGCGATGCCGCTGAGTCAGGCCAGA								:	74
Rhinocer	:	AAAGTTAACGAGTGGTTTCCAGAAGTGATGAAATATTAACTTCTGATGACTCACATGATGGGGGCTGGAATCAAATACTGA								:	83
Pig	:	AAAGTTAACGAGTGGTTTCCAGAAGTGATGAAATGTTAACTTCTGACGACTCACAGGACAGAGGTGTAATCAAATACTGG								:	83
Hedgehog	:	AAAGTGAATGAATGGCTTCCAGAAGTGATGAACTGTTAACTTCTGATGACTCATATGATAAGGGATCTAAATCAAATACTGA								:	83
Human	:	AAAGTTAATGAGTGGTTTCCAGAAGTGATGAACTGTTAACTTCTGATGACTCACATGATGGGAGCTGTAATCAAATGCCAA								:	83
Rat	:	AAAGTGAATGAATGGCTTCCAGAAGTGTTGAAATGTTAACTTCTGACAATGCGATCTGACAGGAGGCCGCGTCAAATGCAGA								:	83
Hare	:	AAAGTTAACGAGTGGTTCCAGAAGTAATGAAATGTTAACTTCTGATGACTCACCTGACCCGGTCTGAATCAAATGCCAA								:	83

	*	100	*	120	*	140	*					
Wombat	:	GGTGCCTAGTGCCTTA	GAACAT	GGGCA	ATCCAGATACC	CGCAGAGGAA	AT	TCTACCGT	TCTGACAA	GACTGAC	:	156
Opossum	:	GGCAACCAATGCTT	TAGAA	TAT	GGGCATGTAGAGACA	--GATGGAAAT	TCTACCATTTCTG	AAAAAGACTGAT			:	153
Armadillo	:	AGTAGCTGGTGCA	TGAAAGTT	-----	TCAAAAGAAGTAGATGAAAT	TCTAGTTTCTCAGAGAAAGATAGAC					:	150
Sloth	:	AGTAGTTGGTGCA	TGAAAGTT	-----	CCAATGAAGTAGATGGATATTCTGGTTCTCAGAGAAAGATAGAC						:	150
Dugong	:	AGTAGCTGGTGCTT	AGAAGTT	-----	CCAGAAGAAGTACATGGATATTCTAGTTCTCAGAGAAAATAGAC						:	141
Hyrax	:	AGTGGCTGGTCCAGT	AAAACTT	-----	CCAGGTGAAGTACATAGATAATTCTAGTTTCTCAGAGAAACATAGAT						:	141
Aardvark	:	AAATAGGTGGTGCATT	AGAAGTT	-----	TCAAATGAAGTACATAGTTACTCTGGTTCTCAGAGAAAATAGAC						:	141
Tenrec	:	CGTAGCTGTAGCCT	TGAAAGTT	-----	CCAGACGAAGCATGTGAATCTTATAGTTCTCAGAGAAAACAGAC						:	141
Rhinoceros	:	AGTAGCTGGTGCA	TGAAAGTT	-----	CAAATGAAGTAGATGGATATTCTGGTTCTCAGAGAAAATAGGC						:	150
Pig	:	GGTAGCTGGTGCAGCAGAGTT	-----	CCAAATGAAGCAGATGGACATTGGTTCTCAGAGAAAATAGAC							:	150
Hedgehog	:	AGTAACTGAAACAGAAAGTT	-----	CCAAATGCAATAGATAAGTTTCTGGTTCTCAGAGAAAATAAAC							:	150
Human	:	AGTAGCTGATGTAGGAGTT	-----	CTAAATGAGGTAGATGAAATTCTGGTTCTCAGAGAAAATAGAC							:	150
Rat	:	AGCTGCTGTGTTAGAAGTT	-----	CTAAATGAAGTGGATGGATGTTCTAGTTCTCAGAGAAAATAGAC							:	150
Hare	:	AGTGGCTGGTGCATTAGAAGTC	-----	CCAAAGGGAGGTAGATGGATATTCTGGTTCTACAGAGAAAATAGAC							:	150

Part of the alignment of the DNA sequences of the BRCA1 gene

	*	20	*	40	*					
Wombat	:	KVNEWLSRSSDILASDN	NGRSHEQSAEV	P	SALEDGHPD	TAEGNSSVSEKTD	:	52		
Opossum	:	KVNEWLFRSNDVLAPDYS	SVRSHEQNAEATNALEY	GH	VET-DGNSS	I	SEKTD	:	51	
Armadillo	:	KVNEWFSRGDDILTSDDSH	DRGSELN	AEV	EDGHPD	T	EYSSF	SEKID	:	50
Sloth	:	KVNEWFSRSDDILTSDDSH	HNGGSESNAEV	VGAL	KV--PNEVDG	YGS	SEKID	:	50	
Dugong	:	KVNEWFFRSDGL	---	DDLDHKGS	ESNAEV	GALEV	--PEEVHG	YSSSEKID	:	47
Hyrax	:	KVNEWFSRSRNLD	---	SDSPSEGSEL	NGKVAGPVKL	--PGEVHRY	YSSFPENID		:	47
Aardvark	:	KVNEWFSRSRNLD	---	DGSHDEGSE	SNAEIGGALEV	--SNEVHSY	SGSSSEKID		:	47
Tenrec	:	KVNEWFSKSHGL	---	GDSRDGRPESG	ADVAVAFEV	--PDEACESY	SSPEKTD		:	47
Rhinoceros	:	KVNEWFSRSDEILTS	DDSSH	DGGPESNT	EVAGAVEV	--QNEVDG	YGSSEKIG		:	50
Pig	:	KVNEWFSRSDEM	LTSDDS	QDRRSE	SNTGVAGAAEV	--PNEADGHL	GSSEKID		:	50
Hedgehog	:	KVNEWLSRSDELLT	SDDS	YDKGSKSKTE	VTTEV	--PNAIDXF	FGSSSEKIN		:	50
Human	:	KVNEWFSRSDELL	LGSDDSH	DGESESNAK	VADVLDV	--LNEVDEY	SGSSSEKID		:	50
Rat	:	KVNEWFSRTGEM	LTSDNA	SDRRPASA	EEAVVLEV	--SNEVDGF	CSSSKKID		:	50
Hare	:	KVNEWFSRSNEM	TPDDS	LDRRSE	SNAKVAGALEV	--PK	EVHGYSGS	TEKID	:	50

Alignment of BRCA1 protein sequences for the same region on the gene

From “Bioinformatics and Molecular Evolution” by Paul Higgs and Teresa Attwood