

Bioinformatic Analysis of NEIL1 in Human

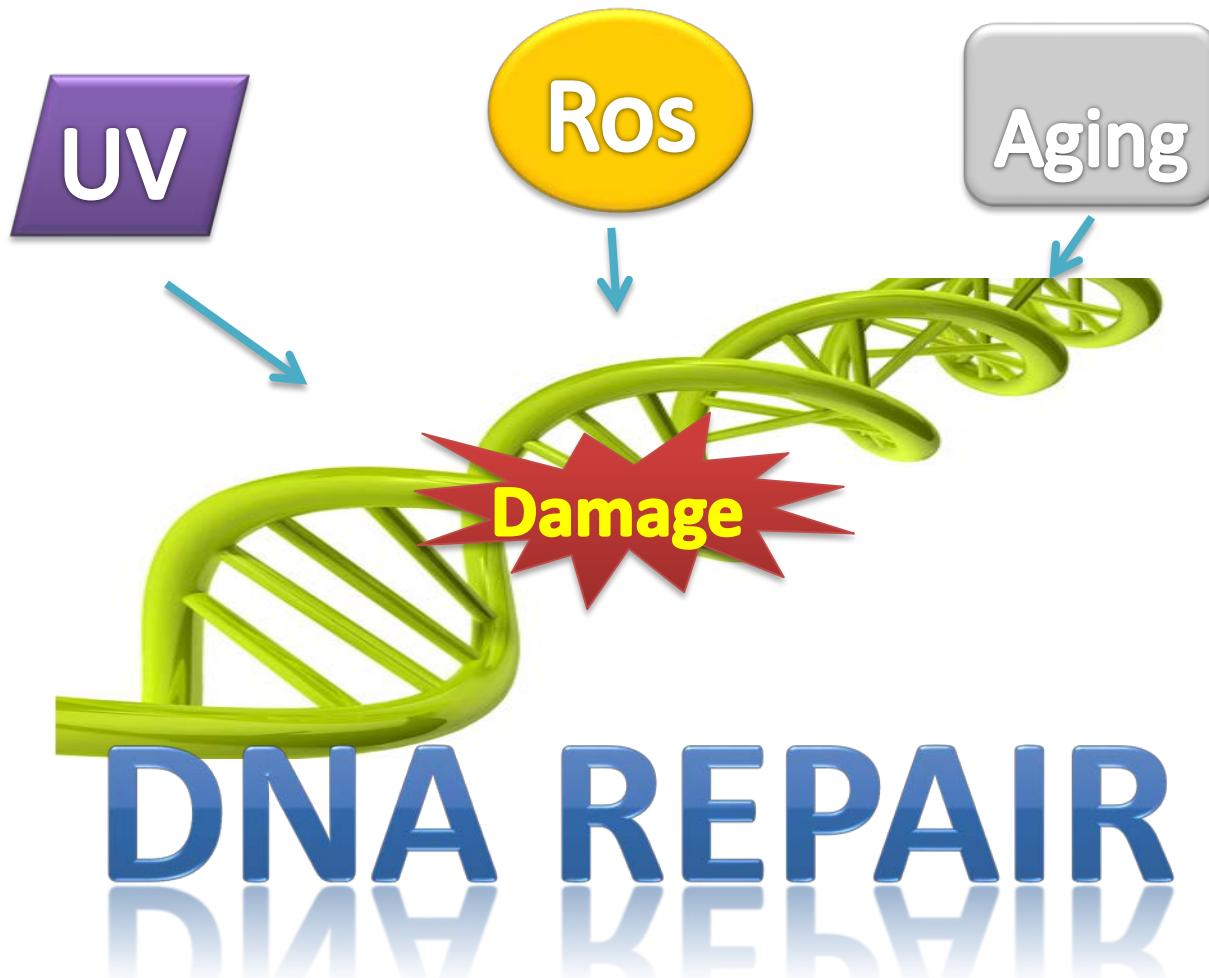
人NEIL1蛋白的相关生物信息学分析

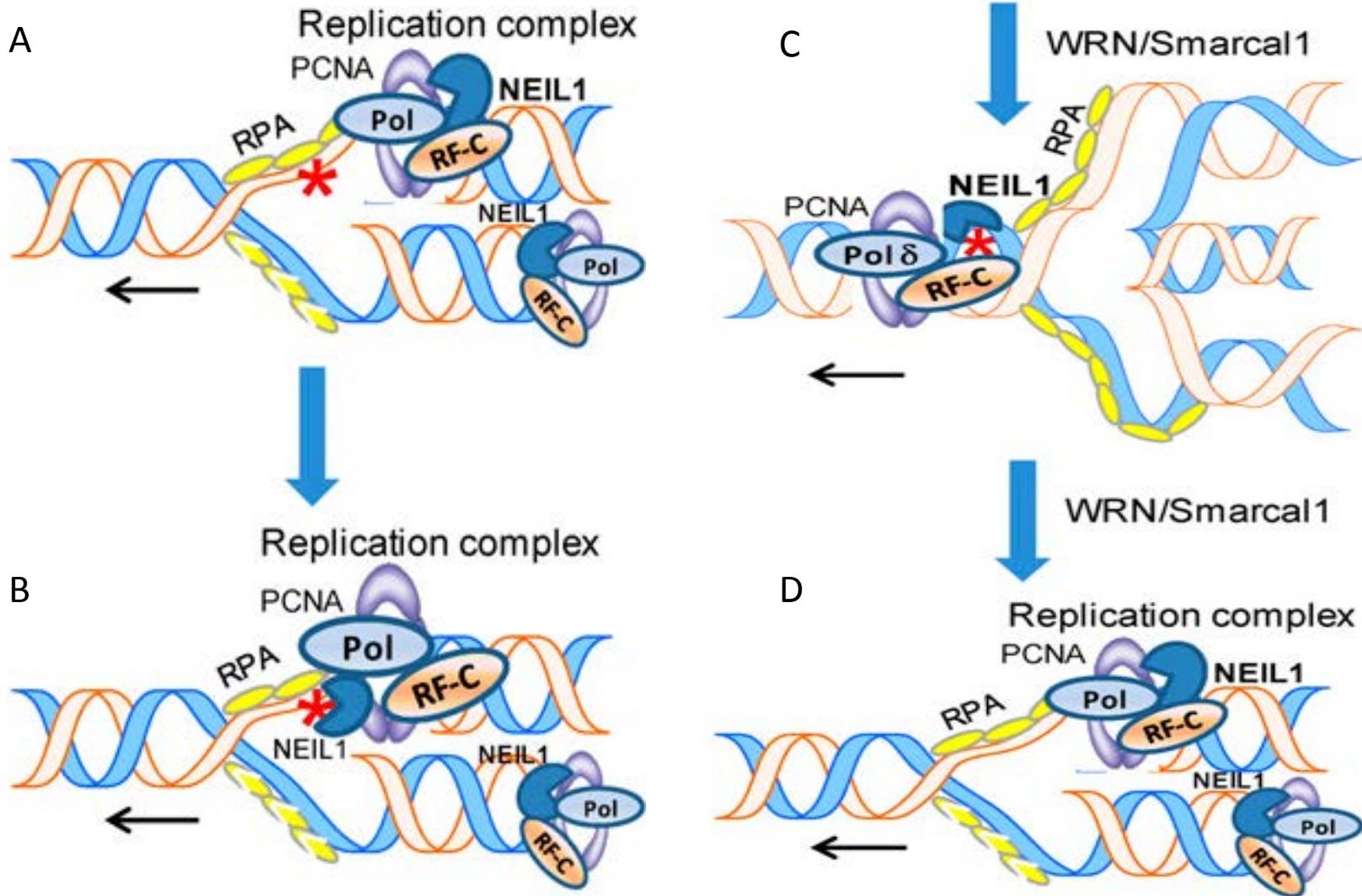
G09: 王奕蓉 田明洁 杨秋华 郑良珺

Outline

- Background
- Sequence analysis
- Structure analysis
- Molecular pathway

NEIL1- new component of BER system





Basic information

- Name: nei Endonuclease VIII-like 1
- Gene ID: 79661
- Sequence length: 389 AA
- Family: FPG family
- Orthologous: NEI (E.coli)
- Paralogous: NEIL2, NEIL3
- Ligand: DNA-binding
- Catalytic activity: Removes damaged bases from DNA, leaving an abasic site.



T-coffee

NEIL1	1	PEGPELHLASQFVNEACRALVFGGCVEKSSVSRNPEV	-PFESSAY	4 4	
NEIL2	1	PEGPLVRKFHHLVSP-F	--VGQQQVVKTGGSSKKLQ	-PASLQLS	3 9
NEIL3	1	VEGPGCTLNGEKIRARVL	--PGQAVTG	--VRGSALR-SLQGRAL	3 9
END8_	1	PEGPEIRRRAADNLEAA	--IKGKPLTDVWFAFPQLK	-PYQSQLI	4 0
FPG_E	1	PELPEVETSRRGIEPHL	--VGATILHAVVRNGRLRWPVSEEIY		4 1
	1	* * : :	*	: .	4 5
NEIL1	4 5	R-----	-ISASARGKELRL		5 7
NEIL2	4 0	WLQ-	--DTQVHGKKLFL		5 3
NEIL3	4 0	RLAASTVVVSPQAAALNNNDSSQNVLSLFNGYVYSGVETLGKEFM			8 4
END8_	4 1	G---QH	--VTHVETRGKALLT		5 6
FPG_E	4 2	RLSDQP	--VLSVQRRAKYLLL		6 0
	4 6		. . * *		9 0
NEIL1	7 2		LALVFRFGMSGSFQLV		8 7
NEIL2	9 9	GSSRSAELVPQGEDDSEYLERDAPAGDAG	RWLRVSFGFLFGSVWVN		14 3
NEIL3	9 0		ALRIHFGMKGFIIMIN		10 4
END8_	6 2		LTLYSHNQLYGVWRVV		7 7
FPG_E	6 6		WIIHLGMMSGSLRIL		8 0
	13 6		: : * : :		18 0
NEIL1	12 8	-WQPGRGPCVLQEYQQFRENVLRN	--LADKAFDRPICEALLDQRF		16 9
NEIL2	18 6	SPVVTPTCDILSE-K	--FHRGQALEALGQAQPVCYTLLDQRY		22 4
NEIL3	14 7	RMMKEL-DVCSP-EFS	--FLRAESEVKKQKGRMLGDLVMDQNV		18 5
END8_	12 0	PFLQRVGPDVLDP	--NLTPEVVKERLLSPRFRNRQFAGLLLDQAF		16 2
FPG_E	12 3	NVLTHLGPPEPLSD	--DFN--GEYLHQKCAKK-KTAIKPWLMNDNKL		16 2
	22 6		* : * : *		27 0
NEIL1	17 0	FNGIGNYLRAEILYRLKIPPEKARSVLEALQQHRSPELTLSQK		21 4	
NEIL2	22 5	FSGLGNIIKNEALYRAGIHPLSLGSVLSASRREVLV		26 0	
NEIL3	18 6	LPGVGNIIKNEALFDSDLHPAVKVCQLTDEQIHHLM		22 1	
END8_	16 3	LAGLGNYLRVEILWQVGLTGNHKAKDLNAAQLDALA		19 8	
FPG_E	16 3	VVGVGNIYASESLFAAGIHPDRLASSLSLAECCELLA		19 8	
	27 1	. * : ** * * : : : : : *			31 5

Sequence logo

MEME

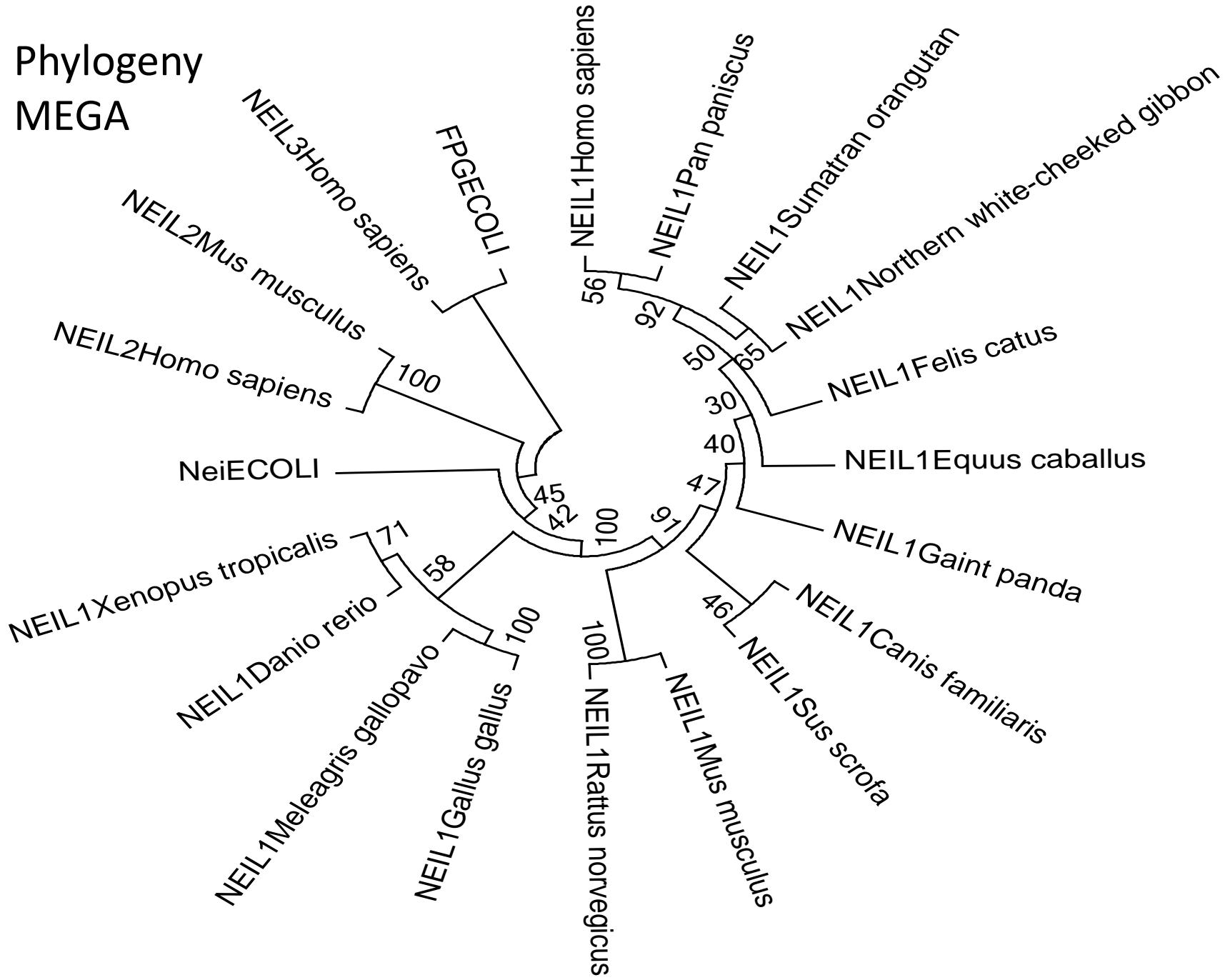


Helix-two turns-helix motif

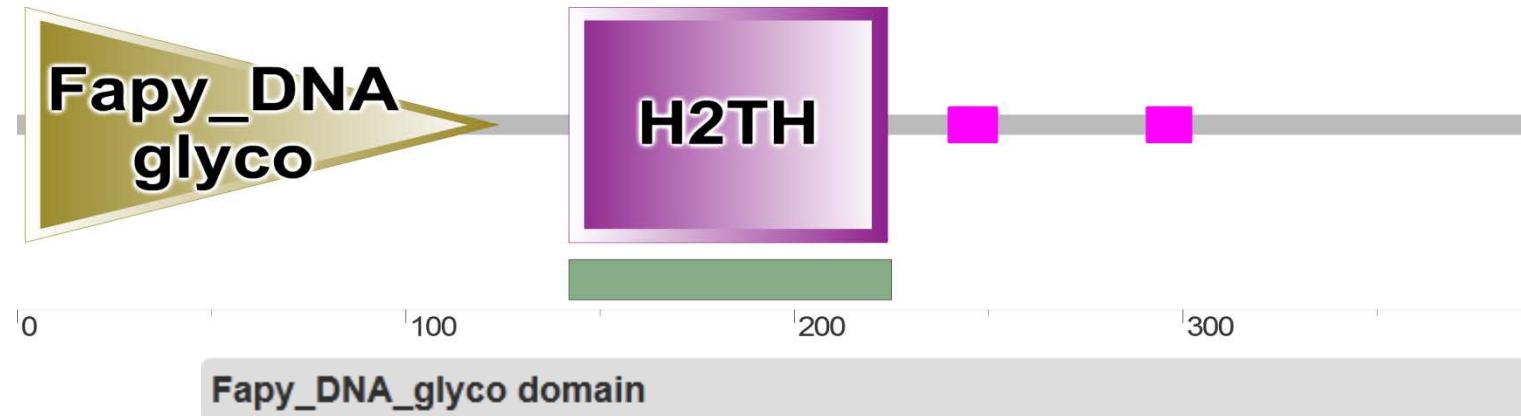


Phylogeny

MEGA



SMART



This is a SMART **Fapy_DNA_glyco** domain ([full annotation](#)).

Position: 2 to 124

E-value: 5.12274901095688e-17 ([HMMER2](#))



H2TH domain

This is a SMART **H2TH** domain ([full annotation](#)).

Position: 142 to 224

E-value: 0.139273012466126 ([HMMER2](#))



- Low complicity region

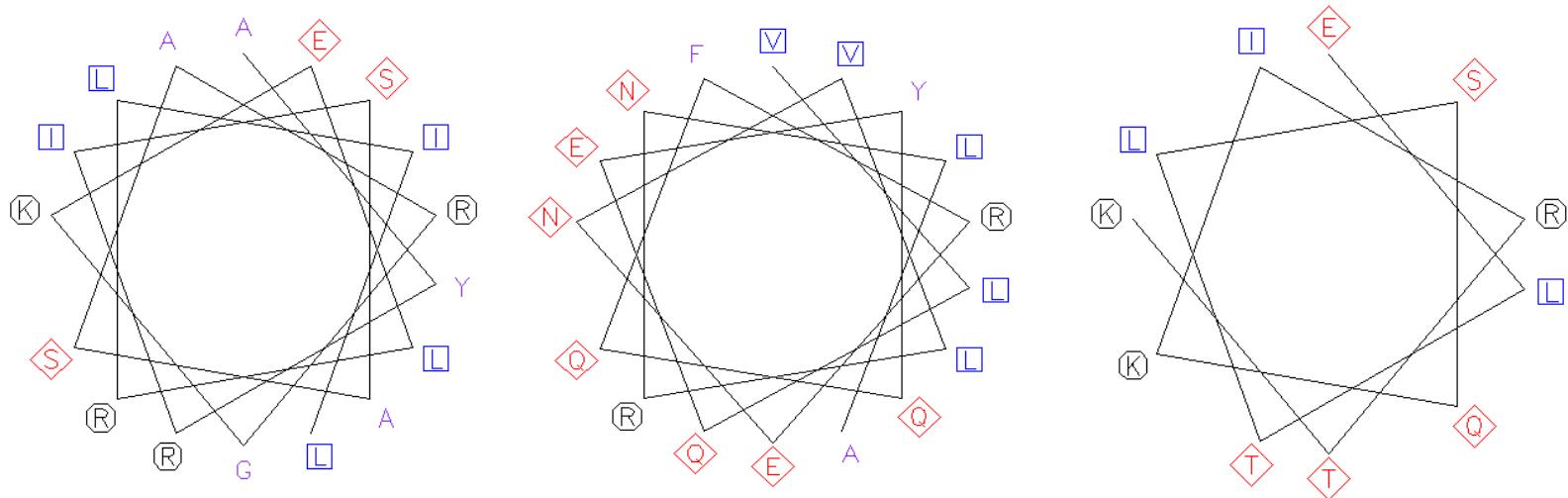
ExPASy GOR

10 20 30 40 50 60 70

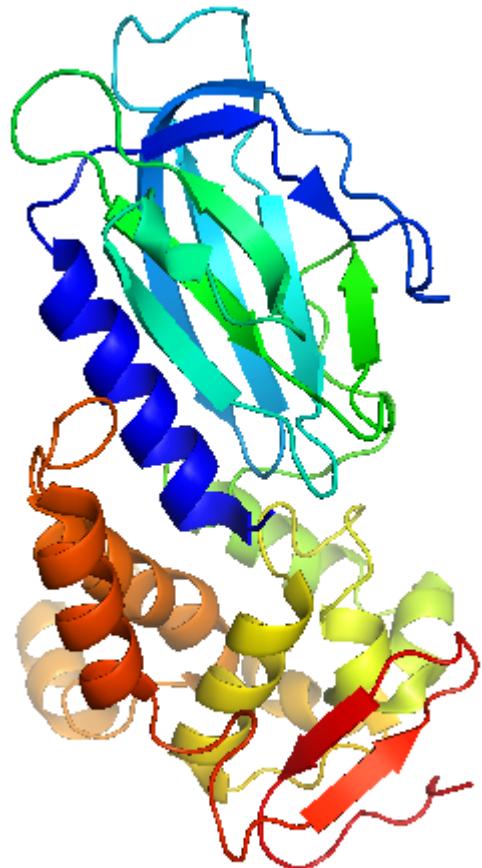
PEGPELHLASQFVNEACRALVFGGCVEKSSVSRNPEVPFESSAYRISASARGKEILRLIILSPLPGAQPQQE
ccccccccc hhhhhhhh c e e e e c
PLALVFRFGMSGSQLVPREELPRAHRLFYTAPPGPRLALCFVDIIRFGRWDLGGKWQPGRGPCVLQEVY
h h h h h h h c c c c c c e e e c
QQFRENVLRLNLADKA FDRPICEALLDQRFFNGIGNYLRAEILYRLKIPPF E K A R S V L E A L Q Q H R P S P E L T
h h h h h h h h h h h h h c c c c c h h h h h h h h h c c c c c c c c c c c c c c c
LSQKIRTKLQNPDLLELCHSVPKEVVQLGGKGYGS E S G E E D F A A F R A W L R C Y G M P G M S S L Q D R H G R T I W F
h h h h h h h c c c c c e e e c
QGDPPGLAPKGRKSRKKSKATQLSPEDRVEDALPPSKAPSRTTAKRDLPKRTATQRPEGTSLQQDPEA
e c c c c c c c c c h h h c
PTVPKKGRRKGRQAASGHCRPRKVKA DIPSLEPEGTSAS
c e e e c

Sequence length : 389

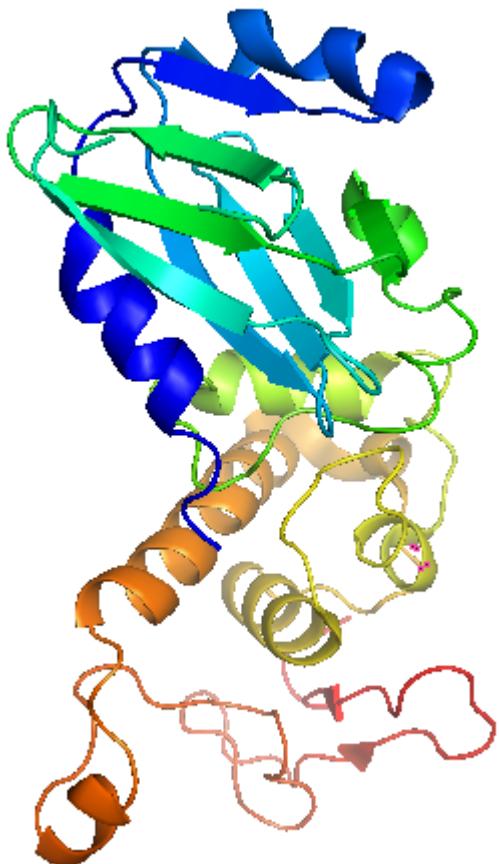
Pepwheel



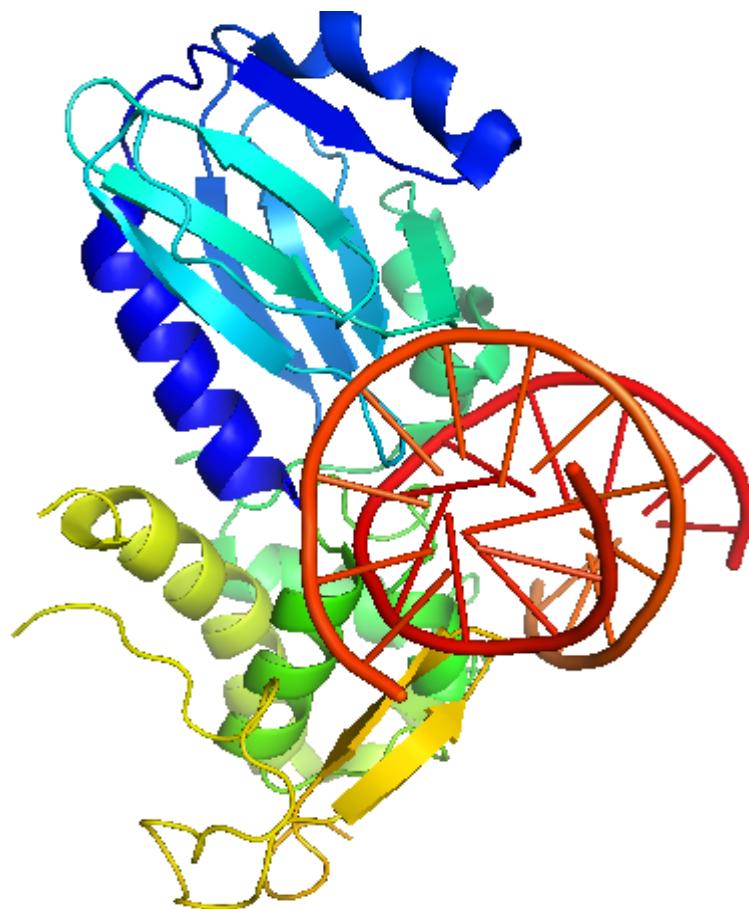
Pymol



NEIL1 (human)
1TDH



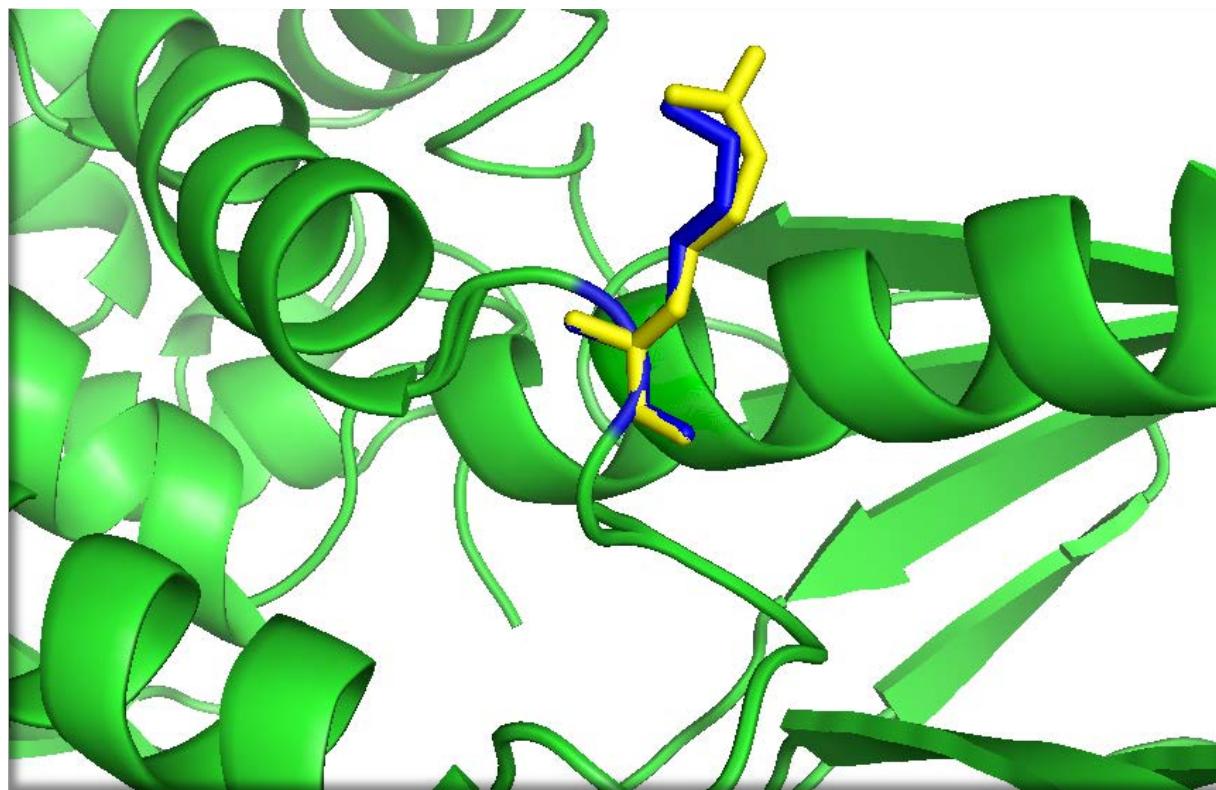
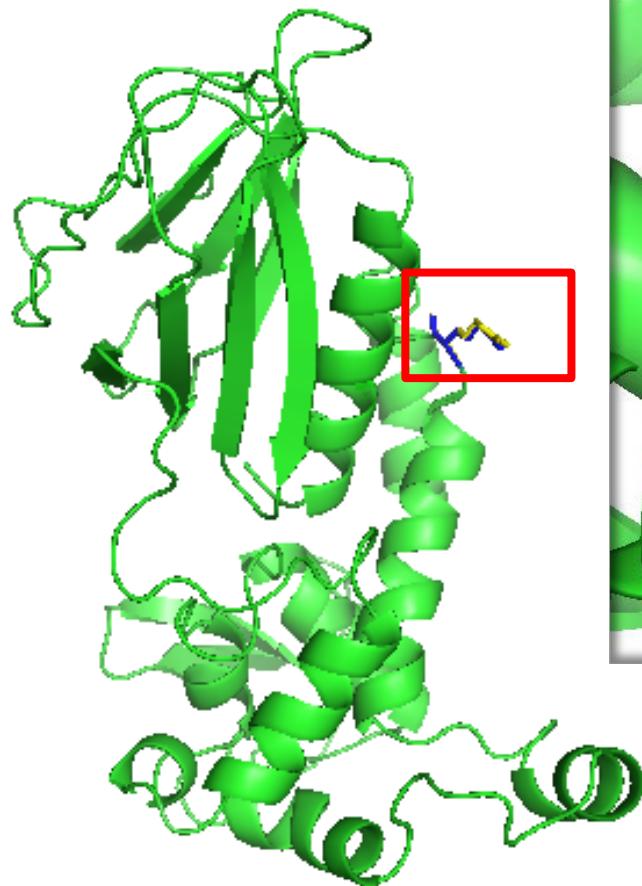
NEI (E.coli)
1Q39



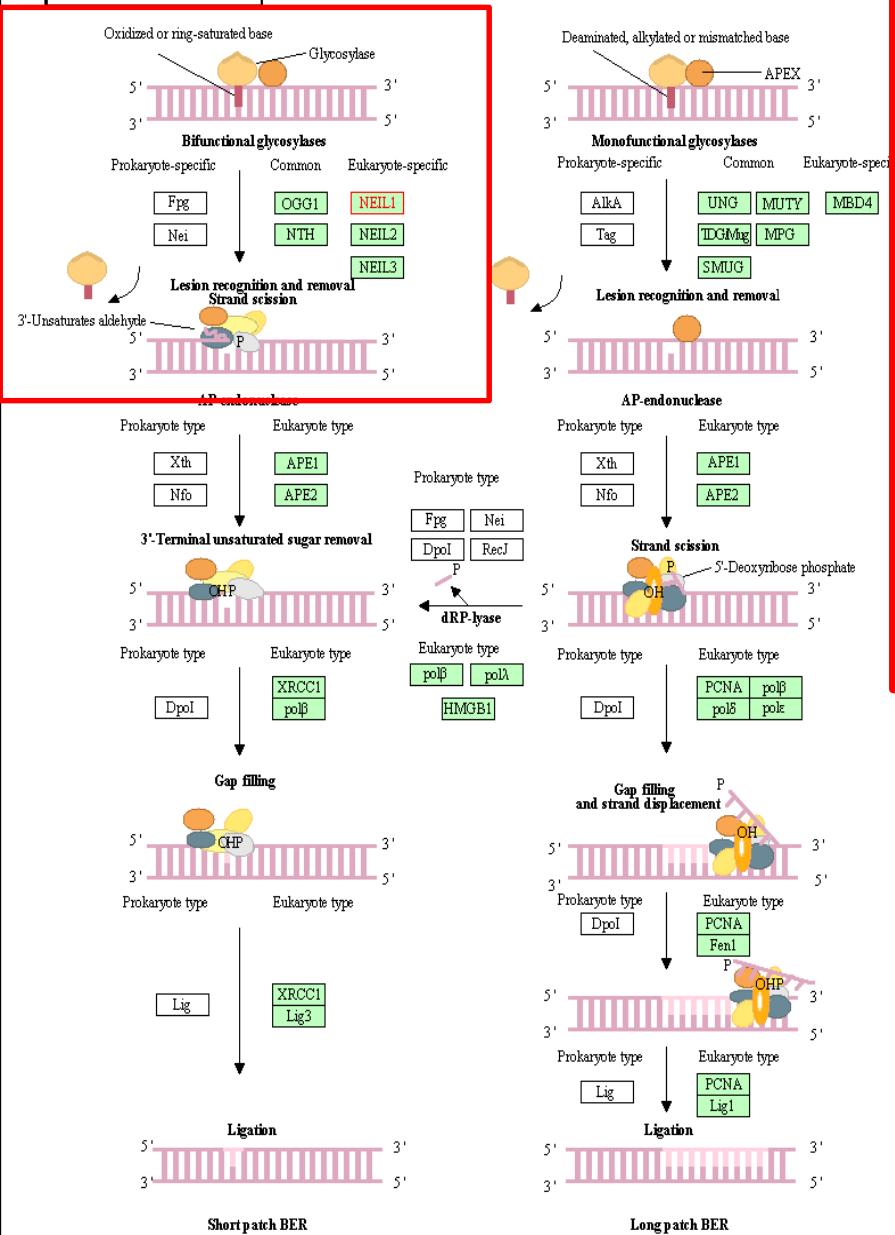
NEI binding with DNA (E.coli)
2EA0

Swiss-Model

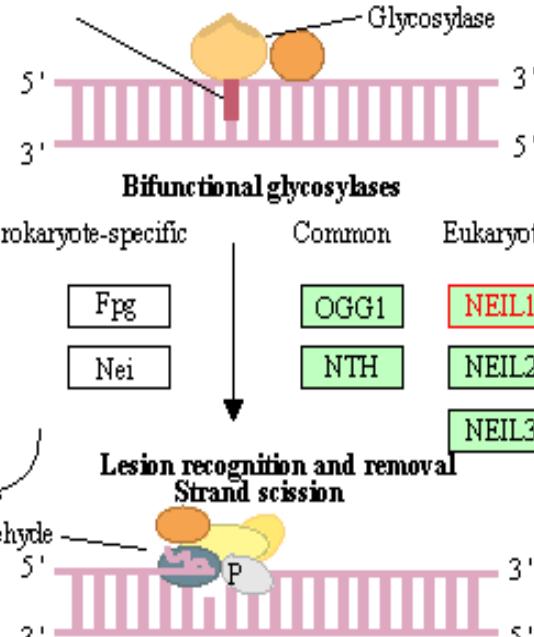
RNA editing: K242—R242



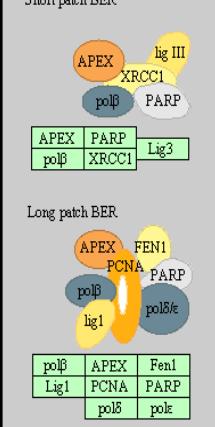
BASE EXCISION REPAIR



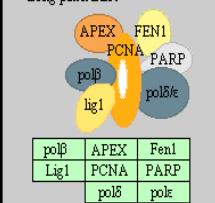
Oxidized or ring-saturated base



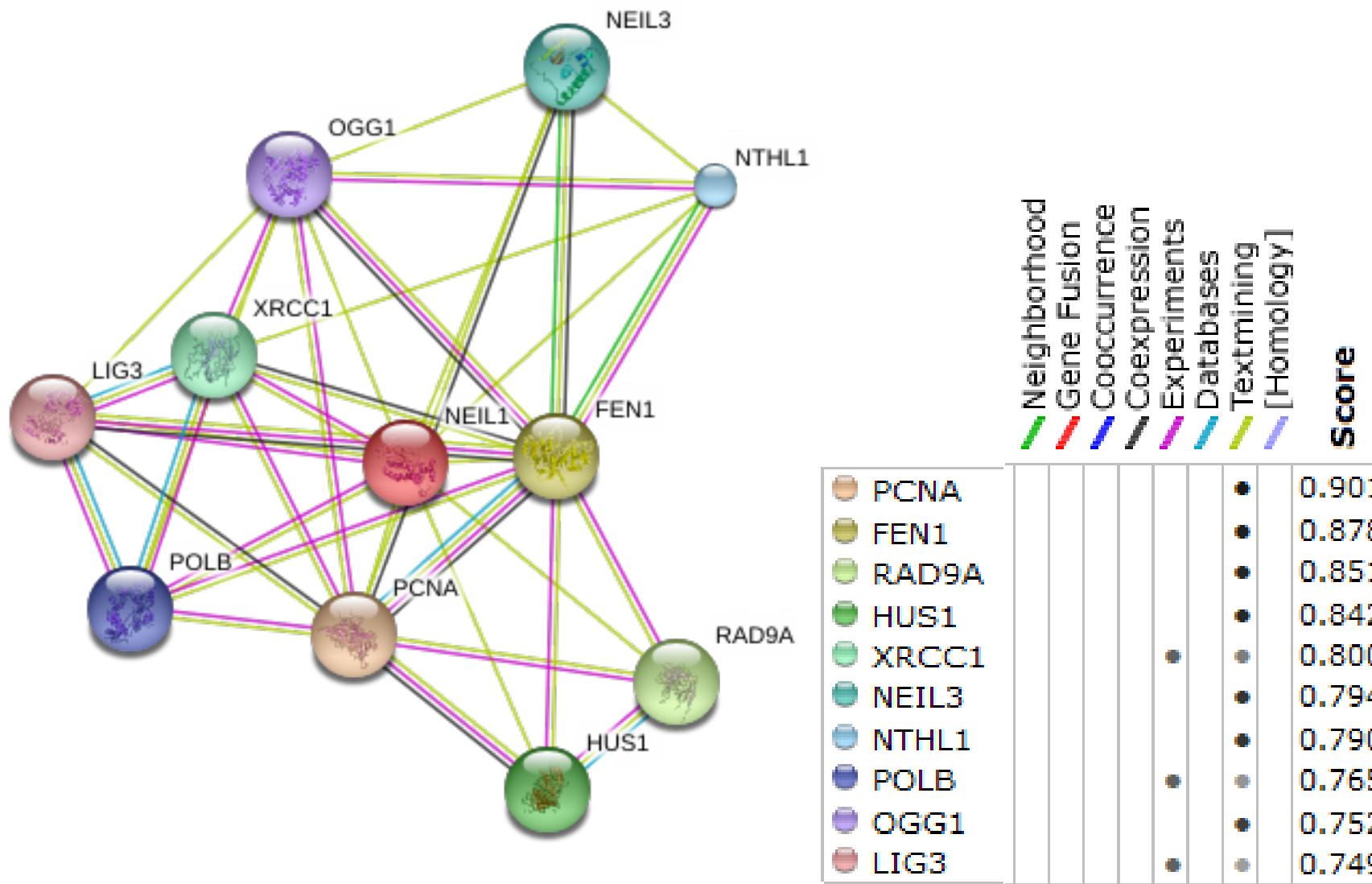
BER complex Short patch BER

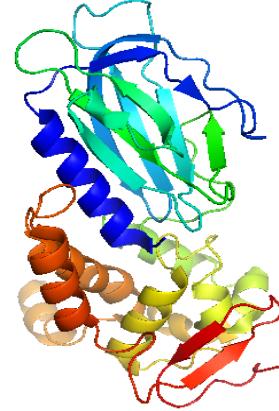
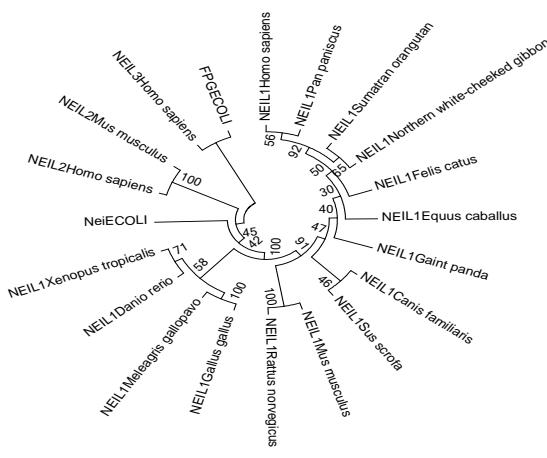
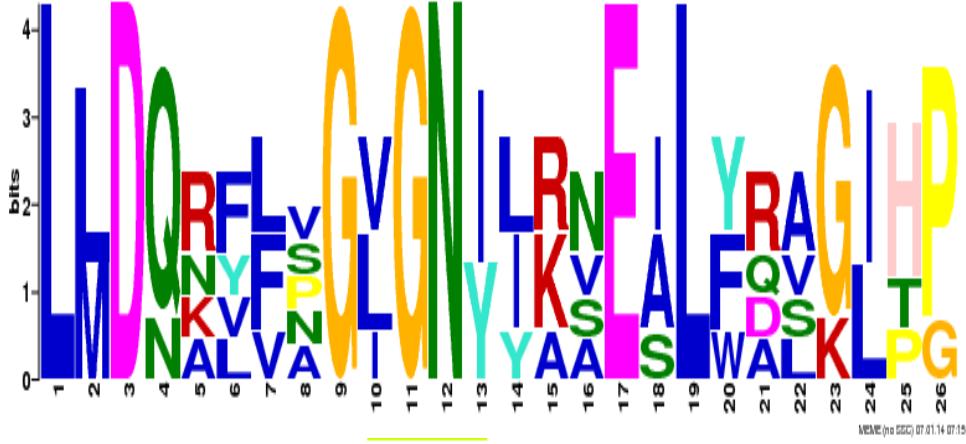


Long patch BER

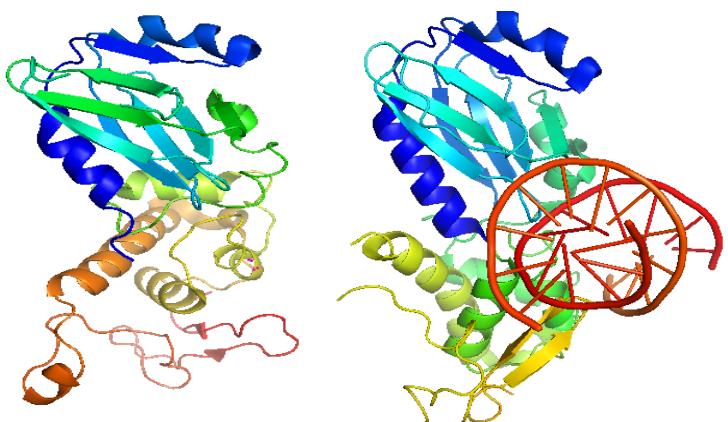


Protein-Protein Interaction STRING





Integrate all clues and make a plan for bench works



NEIL1	128	-WOPGRGPGCVLQEYQOFRENVLRN--LADKA FDRPICEALLDQRE	169
NEIL2	186	S P V V T P T C D I L S E --K-----F H R G Q A L E A L G Q A Q P V C Y T L L D Q R Y	224
NEIL3	147	R M M K E L - D V C S P -E F S --F L R A E S E V K K Q K G R M L G D V L M D Q N V	185
END8_	120	P F L Q R V G P D V L D P -N L T P E V V K E R L L S P R F R N R Q F A G L L L D Q A F	162
FPG_E	123	N V L T H L G P E P L S D -D F N --G E Y L H Q K C A K K -K T A I K P W L M D N K L	162
	226		
NEIL1	170	F N G I G N Y L R A E I L Y R L K I P P F E K A R S V L E A L L Q Q H R F S P E L T L S Q K	214
NEIL2	225	F S G L G N I I K N E A L Y R A G I H P L S L G S V L S A R R E V L V -----	260
NEIL3	186	L P G V G N I I K N E A L F D S G L H P A V K V C Q L T D E Q I H H L M -----	221
END8_	163	L A G L G N Y L R V E I L W Q V G L T G N H K A K D L N A A Q L D A L A -----	198
FPG_E	163	V V G V G N I Y A S E S L F A A G I H P D R L A S S L S L A E C E L L A -----	198
	271		

Reference

- **Grin IR, Zharkov DO.** Eukaryotic endonuclease VIII-like proteins: new components of the base excision DNA repair system. *Biochemistry (Mosc)*. 2011 Jan;76(1):80-93.
- **Yeo J, Goodman RA, et al.** RNA editing changes the lesion specificity for the DNA repair enzyme NEIL1. *Proc Natl Acad Sci U S A*. 2010 Nov 30;107(48):20715-9.
- **Hegde ML, Hegde PM, et al.** Prereplicative repair of oxidized bases in the human genome is mediated by NEIL1 DNA glycosylase together with replication proteins. *Proc Natl Acad Sci U S A*. 2013 Aug 13;110(33):E3090-9.

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