

Supplementary Material for:

Targeted metagenomics: finding rare tryptophan dimer natural products in the environment.

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Supplementary Material Table of Contents

Figure S1. CLUSTALW-based phylogenetic tree of TD environmental sequence tags.	S2
Figure S2. Annotation of the <i>hys</i> gene cluster.	S3
Figure S3. Annotation of the <i>red</i> gene cluster.	S4
Figure S4. NMR spectra of compound 1.	S5
Figure S5. ORTEP diagram of compound 2.	S6
Figure S6. NMR spectra of compound 2.	S7
Table S1. PCR primer list.	S8
Table S2. Tryptophan dimer sequence tag list.	S9

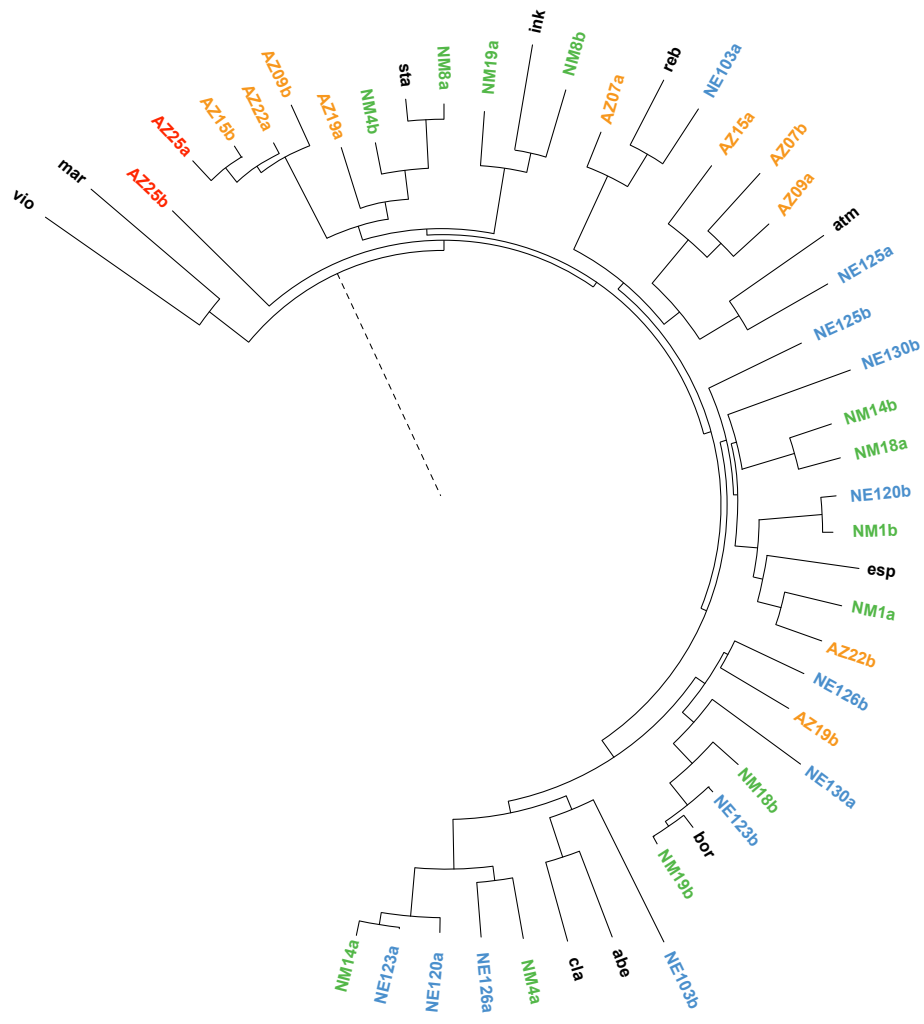
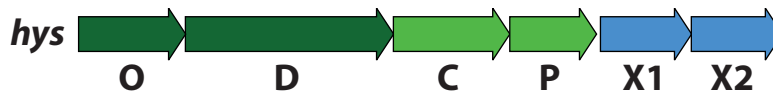
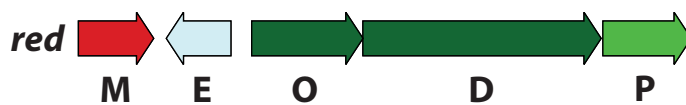


Figure S1. CLUSTALW-based phylogenetic tree of TD environmental sequence tags. CPAS gene sequence tags from Arizona (AZ; orange), New Mexico (NM; green), and New England (NE; blue) soil samples are aligned with the knowns (black). The two tags (AZ25a, AZ25b) investigated in this study are highlighted in red.



N	Gene	Size (aa)	Homolog	ID/SM (%)	Origin	Accession Number
1	orf-25	645	sensor kinase	48/62	Rhodococcus wratislaviensis NBRC 100605	GAF48609.1
2	orf-24	225	LuxR family transcriptional regulator	77/88	Rhodococcus opacus	WP_025433290.1
3	orf-23	322	dehydrogenase	78/85	Pseudonocardia acaciae	WP_028921473.1
4	orf-22	134	transcriptional regulator	71/84	Pseudonocardia dioxanivorans	WP_013677698.1
5	orf-21	286	RNA ligase	78/84	Streptomyces viridochromogenes	WP_003995917.1
6	orf-20	140	archease	59/66	Verrucospora maris	WP_013733957.1
7	orf-19	173	transporter	35/52	Nitrolancea hollandica	WP_008477262.1
8	orf-18	372	MFS transporter	65/79	Myxococcus xanthus	WP_020478826.1
9	orf-17	166	endonuclease	71/80	Pseudonocardia asaccharolytica	WP_028931995.1
10	orf-16	191	alcohol dehydrogenase	32/41	Mycobacterium cosmeticum	CDO08914.1
11	orf-15	198	reductase	74/81	Actinokineospora enzanensis	WP_018687140.1
12	orf-14	402	FAD-dependent oxidoreductase	72/81	Amycolatopsis mediterranei	WP_013226545.1
13	orf-13	119	hypothetical protein	60/75	Nocardia brasiliensis NBRC 14402	GAJ84167.1
14	orf-12	215	hypothetical protein	54/63	Micromonospora sp. ATCC 39149	WP_007071715.1
15	orf-11	123	NADH-quinone oxidoreductase	77/87	Pseudonocardia dioxanivorans	WP_013678831.1
16	orf-10	331	hypothetical protein	40/51	Saccharomonospora xinjiangensis	WP_006240734.1
17	orf-9	249	NADH dehydrogenase	79/86	Pseudonocardia dioxanivorans	WP_013678829.1
18	orf-8	188	NADH-ubiquinone oxidoreductase	71/78	Streptomyces afghaniensis	WP_020275586.1
19	orf-7	100	NADH-ubiquinone oxidoreductase	74/90	Actinomadura madurae	WP_021596654.1
20	orf-6	610	NADH dehydrogenase	64/72	Streptomyces sp. NRRL WC-3641	WP_031120050.1
21	orf-5	502	oxidoreductase	73/81	Streptomyces sp. NRRL F-2305	WP_030835849.1
22	orf-4	476	NADH-quinone oxidoreductase	60/70	Streptomyces sp. NRRL S-37	WP_030871967.1
23	orf-3	223	none			
24	orf-2	287	RNA polymerase sigma factor	56/74	Kutzneria albida	WP_025359630.1
25	orf-1	260	methyltransferase	37/49	Microclunatus phosphovorus	WP_013864415.1
26	O	490	indole-3-pyruvic acid imine synthase StaO	51/64	Streptomyces sp. NRRL WC-3719	WP_031128187.1
27	D	954	chromopyrrolic acid synthase StaD	52/62	uncultured bacterium	AHE14639.1
28	C	533	FAD-binding monooxygenase StaC	67/77	Streptomyces sp. TP-A0274	BAF47693.1
29	P	397	cytochrome P450 StaP	57/68	Actinomadura mellioura	ABC02792.1
30	X1	417	cytochrome P450 hydroxylase	46/61	uncultured bacterium	AHE14835.1
31	X2	417	cytochrome P450 hydroxylase	47/64	uncultured bacterium	AHE14835.1
32	orf+1	140	cyclase	50/63	Frankia sp. QA3	WP_009739180.1
33	orf+2	175	cyclase	58/68	Streptosporangium amethystogenes	WP_030923309.1
34	orf+3	379	alcohol dehydrogenase	76/86	Streptosporangium amethystogenes	WP_030923305.1
35	orf+4	243	cytochrome C oxidase	77/91	Actinoalloteichus cyanogriseus	WP_030104074.1
36	orf+5	364	membrane protein	30/50	Xenococcus sp. PCC 7305	WP_006512167.1
37	orf+6	440	dehydratase	42/55	Oceanicola granulosus	WP_007253650.1
38	orf+7	266	dehydratase	34/55	Mesorhizobium ciceri	WP_024505168.1

Figure S2. Annotation of the *hys* gene cluster. Tryptophan dimer gene cluster found in clone P204 (GenBank accession: KP274855). The annotation is based on BLASTP (NCBI) homology search of translated genes predicted by FGENESB.



N	Gene	Size (aa)	Homolog	ID/SM (%)	Origin	Accession Number
1	orf-2	110	flavodoxin	75/85	Streptomyces filamentosus	WP_006129488.1
2	orf-1	107	multidrug transporter	71/84	Amycolatopsis nigrescens	WP_020667160.1
3	M	344	methyltransferase	64/75	Streptomyces mobaraensis	WP_004943262.1
4	E	295	imine reductase (6-phosphogluconate dehydrogenase)	66/74	Streptomyces mobaraensis	WP_004937742.1
5	O	509	indole-3-pyruvic acid imine synthase StaO	66/73	Streptomyces mobaraensis	WP_004942121.1
6	D	1091	chromopyrrolic acid synthase StaD	57/70	uncultured bacterium	AHE14826.1
7	P	388	cytochrome P450 StaP	65/73	Streptomyces mobaraensis	WP_004942127.1
8	T	399	ABC transporter	57/67	Saccharopolyspora erythraea	WP_011874711.1
9	orf+1	209	tryptophan 2,3-dioxygenase	42/51	Amycolatopsis nigrescens	WP_020673272.1
10	orf+2	221	putative membrane protein	78/84	Streptomyces aurantiacus	WP_016638623.1
11	orf+3	139	ketosteroid isomerase	87/92	Streptomyces afghaniensis	WP_020273417.1
12	orf+4	448	putative glucarate dehydratase	93/96	Streptomyces ambofaciens ATCC 23877	CAJ89466.1
13	orf+5	156	gluconate permease	58/63	Streptomyces ambofaciens ATCC 23877	CAJ89467.1
14	orf+6	92	hypothetical protein	77/86	Streptomyces chartreusis	WP_010040441.1
15	orf+7	587	peptidase S8	84/88	Streptomyces venezuelae	WP_015038177.1
16	orf+8	439	protease	71/80	Streptomyces niveus NCIMB 11891	EST31045.1
17	orf+9	692	ABC transporter	90/95	Streptomyces sp. CNB091	WP_018956698.1
18	orf+10	396	transcriptional regulator	57/71	Streptomyces niveus NCIMB 11891	EST33937.1
19	orf+11	67	hypothetical protein	63/67	Streptomyces niveus	WP_023536416.1
20	orf+12	582	asparagine synthase	55/64	Streptomyces niveus	WP_031228349.1
21	orf+13	214	hypothetical protein	49/60	Streptomyces niveus	WP_031228351.1
22	orf+14	43	none			
23	orf+15	274	phosphoesterase	71/77	Streptomyces sp. Mg1	WP_008737016.1
24	orf+16	651	diacylglycerol kinase	47/59	Streptomyces griseus	WP_003967271.1
25	orf+17	312	ribonuclease	53/67	Streptomyces sp. FxanaC1	WP_018090524.1
26	orf+18	210	hypothetical protein	40/59	Cellulomonas massiliensis	WP_019136883.1
27	orf+19	288	G-D-S-L family lipolytic protein	70/80	Streptomyces pratensis	WP_014152362.1

Figure S3. Annotation of the *red* gene cluster. Tryptophan dimer gene cluster found in clone P121 (GenBank accession: KP274854). The annotation is based on BLASTP (NCBI) homology search of translated genes predicted by FGENESB.

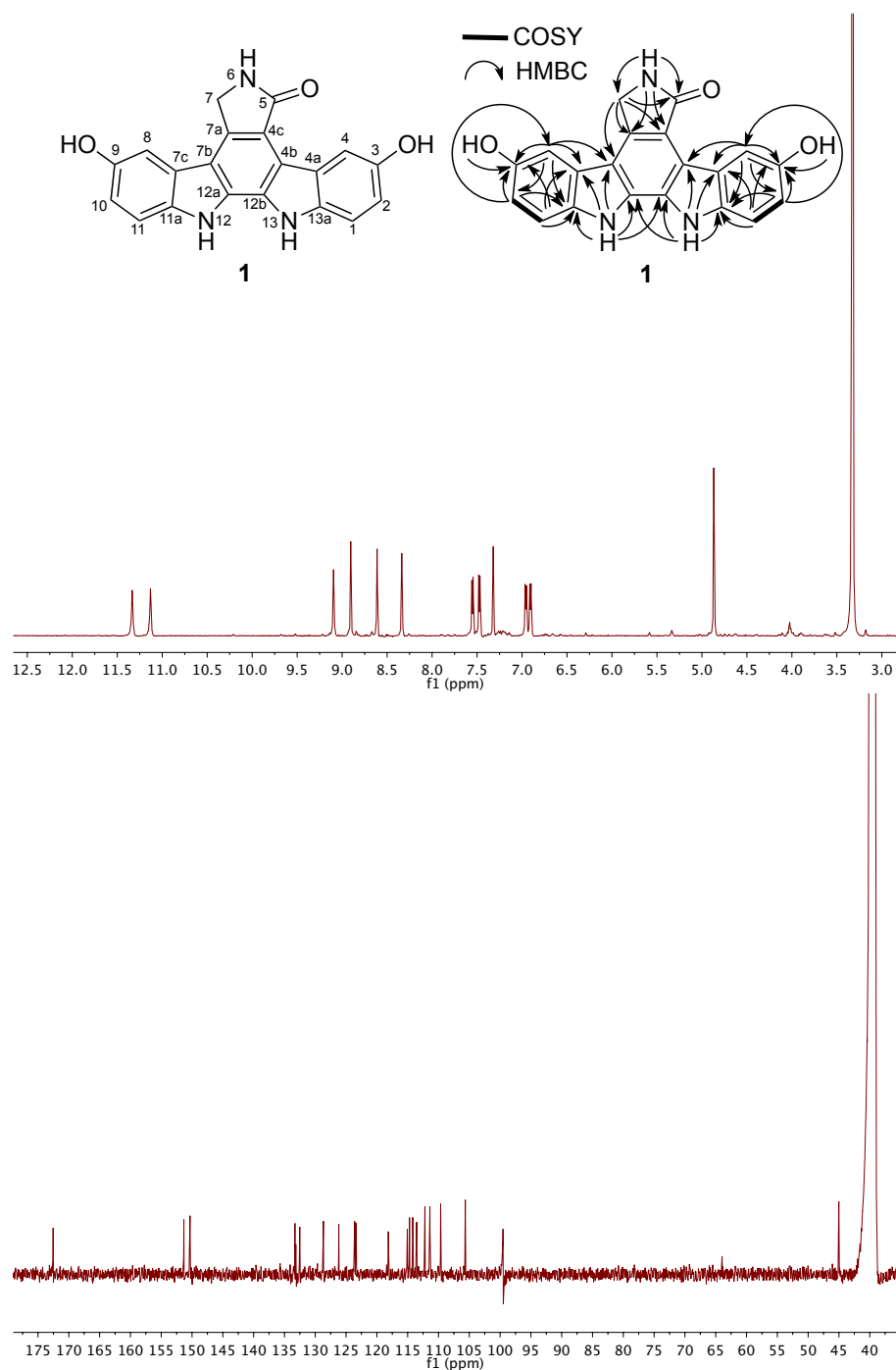


Figure S4. NMR spectra of compound 1. Numbering scheme (top left), the observed 2-D NMR correlations (top right), ^1H NMR (top, 600 MHz) and ^{13}C NMR (bottom, 150 MHz) in $\text{DMSO-}d_6$.

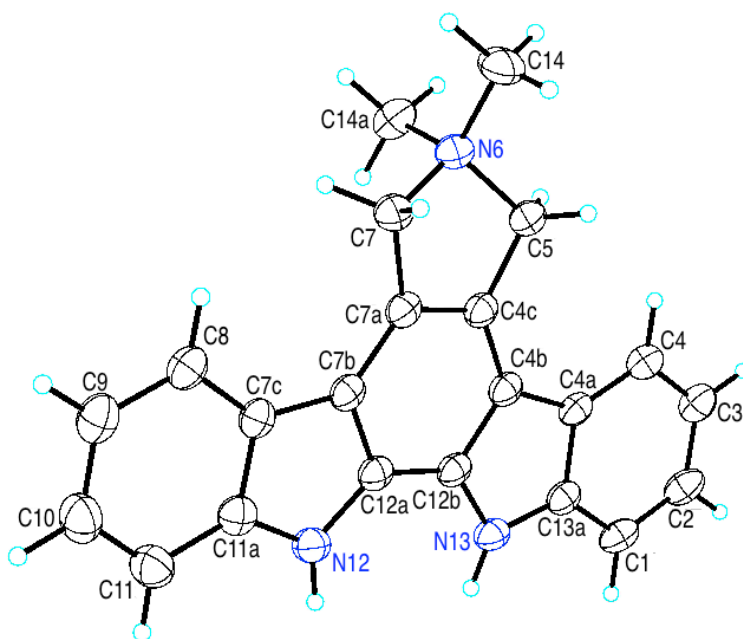


Figure S5. ORTEP diagram of compound 2. CCDC deposition number: CCDC 1043291. Formula: $C_{22}H_{20}N_3^+$, $C_2F_3O_2^-$ (trifluoroacetate salt). Space group: C 2/c. Cell: $a=28.3226(19)$, $b=10.5984(7)$, $c=16.3567(8)$; $\alpha=90$, $\beta=113.507(3)$, $\gamma=90$. Temperature: 203 K. Z: 8. R (reflections): 0.0486(3644).

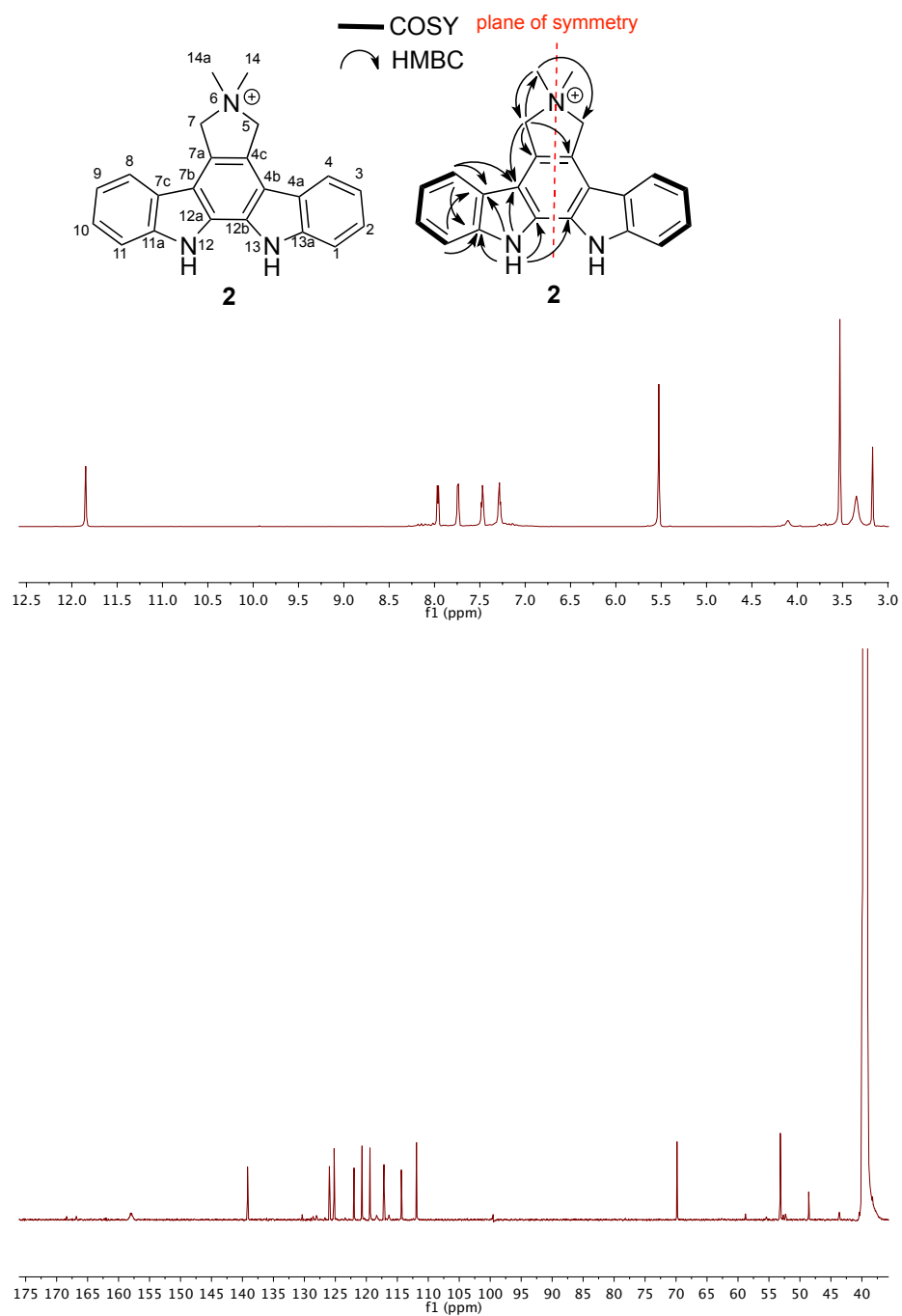


Figure S6. NMR spectra of compound **2**. Numbering scheme (**top left**), the observed 2-D NMR correlations (**top right**), ^1H NMR (**top**, 600 MHz) and ^{13}C NMR (**bottom**, 150 MHz) in $\text{DMSO-}d_6$.

Table S1. PCR primer list.

Name	Sequence
StaDVF	GTSATGMTSCAGTACCTSTACGC
StaDVR	YTCVAGCTGRTAGYCSGGRTG
AZ25aF	ACTGGGTCTGTGGAACGAG
AZ25aR	TCCCTTCTCCACGACGAAG
AZ25bF	ACGCTGCATGACGGGATG
AZ25bR	CCTTCTCCACCAGAAACAGG
EspOF	GAGACCATGGCAATGAGGTGGGATGAGGCG
EspOR	GAGAGCGGCCGCCACTCATCACGACACCTCC
EspDF	GAGACATATGAGTGTTTTTGATCTGCCCC
EspDR	GAGACAATTGCGTCATTGTTTCGTCTCTCGG
EspPF	GAGATCATGATCATGACGCAGCGCGGTACA
EspPR	GAGAAAGCTTCGTTACAGCGGATCGAC
RedOF	GAGACATATGGCCGAGATCCCACACG
RedOR	GAGACAATTGCGAATACGCTCATCACGGGG
RedDF	GAGAACATGTCTATGAGCGTATTCGACCTGCCG
RedDR	GAGAAAGCTTGCATCACGTACCTTCCGCG
RedPF	GAGACCATGGCAATGGACCCGTA CTCTTTTACC
RedPR	GAGAAAGCTTGATCTCGAATGTCCGTCACAG
RedMF	GCGCCCATGGCAATGAGTGACACGTCTCCCG
RedMR	GCGCAAGCTTGTC ACTTGGTGGCGATCAC
RedEF	GCGCCATATGGGTGCCAAGGTGACAGT
RedER	GCGCCAATTGGGTA CT CAGATGGGGCTG

(a) Underlined sequences indicate the restriction sites added for ligation.

Table S2. Tryptophan dimer sequence tag list.

AZ07a:

GTCATGATGCAGTACCTCTACGCGGCCTACTCGATTCCCACCTACGGCGCGGGCGAAGAGCTGGTGCGGC
GCGGTGACTGGACGCCAGAGCAGCTGACTCTGATGTGCGGGCAGCGGTGGCGAGACCCGCGAGGGCGGCC
TGCGGGCGAGCCTGATGGACGTCGCACGCGAGGAGATGATCCACTTCCTGGTCATCAACAACGTGCTGA
TGGCCACTGGTGAACCGTTCTACGTGCCGGACATCGACTTCGGCACGCTGAACGAGCAGCTGCCGGTGCC
GCTGGACTTCTGCCTCGAAGCGTTCAACCTCGGCAGCGTGCAGCGGTTTCATCCACATCGAGCAGCCGGAG
GGCCTCGTGGGCGCGGTCCGGCTGGGCGATCTGCCGTCCTCATCGGACAGTACGACTACGCGTCGCTGA
GCGAGCTGTACGGGGACATCAGGGAAGGCCTGCAACGGGTTCCGGACCTGTTCTGGTAGACCAGGGCC
GTGGCGGTGGCGAGCACC GGCTGTTTCATGCGCGAGTCGGTCAACGCCCGGCATCCCGACTACCAGCTCG
AG

AZ07b:

GTGATGATGCAGTACCTGTACGCAGCGTTCTCAATCCCGACCTACGGTGC CGGCCTGGAATATGTGCGAC
GCGGCGAGTGGACAGTCAACAGTTGCGGCTGGTATGCGGCAACGGGGGAGAAACACGCGACAAGGGG
ATACGCAGCAGCCTGCTACCGTAGCACGGGAGGAGATGATCCACTTCCTGATCGTCAACAACATCATC
ATGGCCATTGGCGAACCATTTACATCCCGGCCATCGACTTCGGCACTATCAACAACCAGCTGCTGGTAC
CGTTGACTTTTCACTCGAGCGCCTGAACATTGGCAGCGTGGCACGGTTTATCGCCATCGAGCAGCCAGA
GGGCTCCTCGGTGAGGTCAGGCGTGAGGACATCACAAGAAGTTCGGACTCTCCCGCGGAGGACCACCT
CTACAGCTCACTTAGCGAGCTTTACGAAACATCCGCGAAGGGCTACAACGCGTCCCGAATATCTTCATG
GTAGACAAGGGCAGAGGGCGGCGGAGAACATCACCTCTTCTCCGAGAATCGATCAACGCAGTCCATCCC
GACTACCAGCTGGAG

AZ09a:

GTGATGCTGCAGTACCTCTACGCGGCGTTCTCGATCCCGACTTACGGCGCAGGCGTGGAGTACGTGCGAC
GCAACGAGTGGACCGCCGACCAGTTGCGGCTGCTATGCGGGCAGCGGGGAGAGACGCGCGACAACGGC
ATGCGGAGCAGCCTGCTACCGTGGCCCGCGAGGAAATGATCCACTTCCTCCTCGTCAACAACATCATCA
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AGCTCACTGAGCGAGCTCTACGCGAACATCCGCGAAGGGCTACAACGCGTTCAGATCTCTTCATGGTGG
ACAAGGGAAGAGGCGGTGGAGAGCACCACCTCTTCTCCGAGAATCGATCAACGCGGTCCATCCCGGCT
ACCAGCTCGAG

AZ09b:

CGCCAGTGTGATGGATATCTGCAGAATTCGCCCTTGTCATGATGCAGTACCTGTACGCGGCCTACTCGGT
GCCACGCACGGCGCTGGCTTGAGTACGTGCGCCTGGGTCTGTGGA ACTCCAGGCAGCTCGCTGTGGCC
TGCGGTGACGGCGGTGAGTCGCTGGACACCGGAATCCGCACCACGTTGCTGTCCATAGCGCGCGAGGAG
ATGATGCATTTCTGATGGTCAACAACGTCCTGATGGCCATCGGAGAACCGTTACGGTGCCGCGGATCG
ACTTCGGCACGATCAACGACGAGTTGCCGGTGGCGCTGGATTTCTCCCTCGAGGGACTGGGGCTTGGTAG
CGTCCAGCGTTTCATCGGATCGAGCAGCCGAGAACGCGGTCGGCGATGTCCAGCTCGGGACGTTGAT
GGCGACGAGAACCAGCCGCTTGC GAATTATGCCGGGCTGAGCGAGCTGTACGGCGATATCCGCGACGG
GCTCGAGCGGATTCCGGGCCTTTCGTGGTTCGAGAAGGGACGCGGGCGGCGAGCACCATCTGTTCT
GCGTCGCTCCATCGACCTGGACCACCGGGCTACCAGCTTGAAAAGGGCGAATTCCAGCACACTGGCGG
CC

AZ15a:

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ATGCGAGGCAGCTTGCTACTGTGGCGCGGAGGAAATGATCCACTTCTGGTCGTCAACAACATCATCA
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ACCAGCTGGAG

AZ15b:

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CACCACGTTGCTGTCCATAGCCCGCAGGAGATGATGCACTTCTTGATGGTCAACAACGTCCTGATGGCC
ATCGGGGAACCTTTCACGGTGCCGCGGATCGACTTCGGCACGATCAACGACGAGCTGCCGGTTGCGCTG
GACTTCGCACTCGAGGGGCTGGGGCTCGGCAGCGTCCAGAGGTTTCATCGAGATCGAGAAGCCCGAGAGT
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TGAG

AZ19a:

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CACCTACGGCGCCGGCGCCGAGTACGTGCGGCTCGGCCTGTGGACCCCGAGCAGCTCCAGCTCGCCTG
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AZ19b:

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AZ22a:

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AZ22b:

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TGCGCATGGGTGAGCCGTTCTTCGTGCCGGCCGTCGACTTCGGCACGGTCAACAACACCCTGCCACTGCC
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TACATACGGTTTCGCTCAGCGGCCTGTACGCCGACATCCGGGAAGGACTGCAGCGGGTGCCGGACCTGTT
CATGGTGAAGAAGGGCCGTGGCGGAGGCGAGCACCACCTGTTCTCCGTGAGGGCGTCAACGCGGTCCA
CCCGGACTACCAGC

AZ25a:

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GGCCATCGGAGAACCGTTCACGGTGCCGCGGATCGACTTCGGCACGATCAACGACGAGTTGCCGATCGC
GCTGGAGTTCTACTCGAGGGGCTGGGGCTCGGCAGCGTCCAAAGGTTTCATCGAGATCGAGAAGCCCGA
GAGTGCGGTCGGCGATGTACATCTCGGCGACGTCAGGGCGACCGGACTCCGGCCCGCTTCAGTTATGC
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AAGGGACGCGGCGGGCGGAGCACCCTTGTTCCTGCGCCGCTCCATCGACATGGATCACCCGGACTAC
CAG

AZ25b:

CTCCAGTACCTGTACGCGGCCTTCTCCGTACCGACCCACGGGGCAGGGCTCACGTACGTGCGCCGGGGCC
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AAGGGGCGCGGCGGGCGGTGAACACCATCTTTCTGCGGGAGTCGGTCAACAGGAACCATCCGGACTAT
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CGGCATCGGCAGCGTCCAGCGGTTTCATCGCCATCGAACGCCACCCACGCCAGGACGGCGAGATCCAGAT
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CTACGCCGACATCCGGGAAGGACTGCAGCGGG

NM1b:

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CATCCGCGAGGCCTGCAACGGGTCCCCGACGTC

NM4a:

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GGACGCGGCGCCCTGGAGCGGTTTCATCGCGCTGGAGCGGCCCTACGACCTGGTCAGGGACATCGCCGGG
AACGACGCGCCGGCCGGCGGCGGTGCCCGAGGGGCGCGCCCCGTACGGTACGGCTCGCTGAGCGAGCTGT
ACTCGGCCATCCGCGAAGCCGTCAGGCCATCC

NM4b:

GCCTATTCGGTGCCACCCACCGTGGCCGCTGGAGTACGTGCGGCGGGGCACGTGGACACCCGAGCAG
CTCCGGCTCGCCTGCGGCGACGGCGGGGAGACCCTAGACGAGGGCATCCGACGATGCTGCTGAACATC
GCCCCGAGGAGATGATCCACTTCTCCTCGTCAACAACATCCTCACCGCGGTGGGCGAGCCGTTCCACG
TGCCCCGCTCGACTTCGGCACCGTCAACCACGAGCTGCCGGTACCCTGGACTTCTCCCTGGAGAGGCT
GGGGCCCCGCGACGTCGAACGGTTCGTGCAGATCGAACGGCCGGAAGACCTCGTCCACGACGTACGGCG
GGGCGACACCCGGCGGCCCGCCCGACGACCGGCACCCCGTACGCCTCGCTGAGCGAGCTGT
ACACGGACATCCGCGAGGGCCTGCACCGCGTCCC

NM8a:

GCCTACTCGGTACCCACCCACGGCGCCGGCGCGGAGTACGTACGCCGGGGCCTGTGGACGCCCGACCAA
CTGCGGCTCGCGTGCAGTACGGCGGGGAGACCCTCGACGAGGGCATCCGCAGCATGCTGCTGACCATC
GCCCCGAGGAGATGATCCACTTCCTCCTCGTCAACAACATCCTCATGGCGGTGGGCGAACCCTTCCACG
CGCCCCGATCGACTTCGGCACCGTCAACCGACAGCTGGCCGTCCCGTGGACTTCGCCCTGGAGCGCCT
GGGGCCCGCAGCGTGGAGCGGTTCTGACAGATCGAACGCCCGAGGACCTCGTCGACGAGGTACGGCG
CGGCGACGCTCCGGCGCCCCCGCCGCTACGACGAGCGGCACCCGTACGCTCGCTGAGCGAGCTGTAC
GCGGACATCCGGGAAGGGCTGGAGAGCATCCCG

NM8b:

GCGTACTCGGTCCCGGCGCACGGCGCCGGCCAGGAGTACGTCCGCACCGGCGACTGGACCGAGGAACAG
CTCCGGACCGCCTGCGGTGACGGGGGAGAGACGCTCGACGGCGGCATCCGGGGCGTCTGGTCCGGGTC
GCGCGCAGGAGATGATCCATTTCTGGCGGTCAACAACATCCTCACGGCCATCGGTGAGCCGTTCCACG
TCCCGACGTCGACTTCGGCACCCCTGAACGCGACACTGCCCGTCCACTGGACCTCTGCCTGGAACGCCT
GAACCTCGCCAGCGTCGAGCGGTTCTGTCGAGATCGAGCGGCCCGAGGGCCTGACCGGCGACGTGCGGCT
GGGGGAGCGGGCGGGGCGCGGTGAGCTCGACCGGCACACCTACTCGTCGATCAGCGAGCTGTACGGCGA
CATCAGGCAGGGCCTGCAGGACATCCCGG

NM14a:

CTGTGGTCCATCCCCACCCACTCGGCCGGGATCGAGTACGTGCGGCGACGCGAGTGGACGCCCGGGCAG
CTCCGGCTCATGTGCGGCGAGGGCCCGCACAGCCTCGACGGCGGCGTCCGGCAGGACCTGTTCGCCGTC
GCCCGCAGGAAATGATCCACTTCCTGCTGATAAACAACATCATCATGGCCATGGGGGAGCCGTTCCACC
TGCCCCGATCGACTTCGGCACAGTGAACGGCGAGCTCCCGTCCCATCGACCTGTGCCTGGAGCCCTTCG
GGCGGGCAGCTGCAACGGTTCGCCGCGCTGGAACGGCCCTACGATCTGGTCCGCGACCTCGCCGCCGA
CCGGCCACCGGTGACCGCGTGCACCCGTATCCCTACGGTTCGCTCAGCGAGCTGTACAAGGCCATCCCG
CAGGCGATCCAGGGACATCCCGGACGT

NM14b:

GCGTACTCCATCCCGACCCACGGCGCCGGGGTTCGAGCACGTCCGCCGCGGCCTGTGGACCCCGCAGCAG
CTGGAGCTGGCCTGCGGCGACGGCGGCCACCACCGCCGGCGGGTTCGCGGCATGCTGCTCGGTGTG
GCCCGCAGGAGATGATCCACTTCCTGCTGGTCAACAACATCATCATGGCCATGGGGGAGCCGTTCCAC
GTGCCGGTGGTTCGACTTCGGCACCGCCAACACCACCCTGCCGGTGGCGCTGGACGTCAGCCTGGAGGGC
CTAACCTCGGCAGCGTCCAGCGGTTTCATCGCCATCGAGCGGCCGGACTGCGAGGTGCGCGAGCTGCGC
CGGGCGCCGGGACCGGAGACCCGCTCGCCGCGCCGGCCGGTTCGCCGCGCTGCTACGGCACCGTCAGC
GAGCTGTAACGCCGAGACGTCCGGGGAGGGCC

NM18a:

GCGTACTCGGTGCCGACCCACGGCGCCGGGGCCGAGCTCGTCCGGCGCGGCCGGTGGACGCCCGAGCAG
CACCGGCTGGCCTGCGGCGACGGCGGTGAGACCACCGACGCCGGGGTTCGCGGCATGCTGCTGTCGGTG
GCCCGGAGGAGATGATCCACTTCCTGCTGATCAACAACATCATCATGGCCATGGGTGAGCCGTTCCACG
TGCCGGTGGTTCGACTTCGGCACCATCAACACCACCCTGCCGGTGGCGCTGGACGTCAGCCTGGAGGGCCT
GAACGTCGGCAGTGTGACGCGGTTTCATCGCGATCGAACGCCCGACACCAGATCGGCGAGCTGCACCCG
GCGGCCGGTACCGGCTACCCACCGGTCGTCGACCCGACCCGACCGCCGCGCAGCTACGGCACCCCTCCAG
CGAGCTGTACGCCGAGACTCCGCGACGGGCT

NM18b:

GCCTGGTTCGATCCCGACGGCCGGCGCCGCCGAGTTCGTCCGCCGCGGCGAATGGACGCCCGGAGCAG
CTGCGGCTGGCGTGCAGGCGACGGCGGACCCACCCTCGACTACGGGATGCGCGGCACGCTGCTCAACGTG
GCCCGCAGGAAATGATCCACTTCCTGGTTCATCAACAACATCATCACCGCGACCGGCGACGCGTTCCAC
GTGCCGGCGATCGACTTCGGCACCCCTCAACGAGCAGCTGCCGGTGGCGCTGGACTTCAGCCTGGAGGGC
TTCGGGCTGGGGCCGCTGCAACGCTTCATCGCCATCGAACGGCCGGACGACCAGACCGTCGAGTTCGCC
GGACCGACACGCTGCTCGACCGGGGCGACGCGCTGTACCCGTACGGCTCGCTCAGCGAGCTCTACGCG
GCCATCCCGCAGGGGCCATTCCAGCCGGGT

NM19a:

GCGTACTCGGTGCCGACGTACGCGGCGGCCGAGGCCTTCGTCCACCGCGGTCTGTGGACCCCGAGCAG
CTGCGGGTTCGCCCTGGGTGACGGCGGAGAGACCCTCGACGGCGGCGTCCGGGGAAAGCTGGTGGAGATC

GCGCGGAGGAGATGATCCATTTCTCCTCGTCAACAACATCCTCATGGCTCTCGGTGAGCCCTTCTGCG
TCCCCGCGCTGGACTTCGGCGCGCTCGGCACCGACCTGCCGGTGGCCCTGGACCTGTGCCTCGAAGGGCT
CGACATCGCCACCGTGGCGCGGTTTCATCGCGATCGAGCAGCCCGCGGTGGGCACGCCGAGGTACGGCG
ACCCGACCTGCCACCACCAACGGCGCGGGTTCGGGGCCGGTACGAGACGTTGAGCGAGATGTACGC
GCGGATCCGTCAGGGGCTGCAGGACGTCCCG

NM19b:

GCCTGGTCCATCCCGACGGCCGGTGGCGGGGCGGAACTCGTCCGCCGTGGCGAGTGGACCCCGGAGCAG
TTGCAGCTGGCCTGCGGGACGGCGGTCCGACGCTCGACTACGGGATGCGCGGCACGCTGCTCAACGTG
GCCCCGGAGGAGATGATCCACTTCTGGTTCATCAACAACATCATCACCGCCACCGGGGACAGCTTCCAC
CTGCCGGCGATCGACTTCGGCACCCCTCAACGAGCAGCTGCCGGTGGCCCTGGACTTCAGCCTGGAGGGT
CGGGATCGGGCCGTTGCAGCGGTTTCATCGCCATCGAGCAGCCGGACGACCTGACCGTCGAGTTCGCCGG
GACCGACACCCGTGCGTCGACCGGGGCGACGCCATCGTACCCGTACGGCTCGCTCAGCGAGCTGTACGC
GGCGAGTCCGGGAGGCCGTCCGAGCCGGGGC

NE103a:

GCGTTCTCGATCCCCACCTACGGCGCGGGTTCAGCAGCTGGTGGCGCGGGTACTGGACACCGGAACAG
CTGCGGCTCATGTGCGGTGACGGGGTGAGACGGCGGACGGCGGCGTGGGGGCGCGCTGCTCGGCGTC
GCCCCGAGGAGATGATCCACTTCTGGTGGTCAACAACGTCTCGTGGCCATCGGCGAGCCGTTCCACC
TGCCCGACATCGACTTCGGCACGATCAACGACAACCTCATGGTGGCGCTCGACTTCTCGCTCGAGGCGCT
CGGGCTCGGCAGCGTACAGCGTTCATCCAGATCGAGCAGCCGGACGGGCTCGTGGCGCCGTGCGGGT
CGGCGACCTCCCGCCCCGTTGAGTCCGAACAGGACTACGACTACGCCTCGCTCAGCGAACGTGTACGGT
CGGACGACTACACGGAAGGGGTTAGGTTCGACC

NE103b:

TGTGGTTCGCTGCCGACTACCGTATGGGGCCGCGTGGTCCACCAGGGCGAGTGGACCGAGGACCAGTA
CACGCTGGTCTGCGGGCGCGGGCCGGCGACGGCTGACGGGGGATCCGGGGCGCTCTGTTTCGGGGTGGC
CCGAGAGGAGATGATTCATTTCTCGTTCATCAACAACATCATCATGGCCACGGGTACGCCCTTCCACGTG
CCCGACATCGACTTCTCTCGTCAACGCGCAGATCGACCTGCCGATGGACTTCTGCCTGGAGCGGTTTCG
GCCTGTCTCGCTGAGCAGTTTCGTCGAGTTTGAAGGCGTTTTCGCTCACCCTCGAGCCAGCGCCGGC
ACCGGGAGGCCCGCAGCGAAAGGGTGGCCGTTACGGGTCGCTGAGCGAGCTCTACGCGTCGATCCGGGA
CGCCTCGCCCCGTTCCCGAGGCCTTCT

NE120a:

CTGTGGTCCATCCCCACCCACTCGGCCGGGACCGAGTACGTGCGGCGGGGCGAGTGGACGCCCGGGCAG
CTCCGGCTCATGTGCGGGACGGCGCCGGAGCCGCGACGGCGGTATCCGGCAGGGCCTGTTTCGCCGTC
GCCCCGAGGAAATGATCCACTTCTGCTGATCAACAACATCATCATGGCGACCGGCCAGCCGTTCCACC
TGCCCCGGATCGACTTCGGCACGGTGAACGGCGAACTCCCCGTCCCCTCGACCTGTGCCTGGAGCCCTTC
GGGCGGGGACGCTGCAACGGTTTCGCCGCGTGGAAACGACCCACGACCTGACCCGCGACCTCGCCGAG
GACCGGCCGCGGTGACCACGCGGACCCGTACCCCTACGGTTCGCTCAGCGAGCTGTACGCCGCCATCC
GCCAGGCCGTCCAGGACATCCCGGACGTCTTCC

NE120b:

GCGTACTCGGTCCCGGCCCTACGGGGCGGGGAGGAGTACGTCCGGCGGGGGCTGTGGACCCCCGAGCAGC
TGCGGCTCGCTGCGGGGACGGCGCCGGAGCCGCGACGGGGGATCCGCGGCACGCTGCTCGGCATCG
CCCGGAGGAGATGATCCACTTCTGATCGTCAACAACATCATCATGGCGATGGGCGAACCCCTTCCACGT
CCCCGACGTCGACTTCGGCACGATCAACAACACCCTGCCGGTGGCGTGGACTTCGCCCTGGAGCCCTTC
GGCGTGGGACGCTCCAGCGGTTTCATCGCGATCGAACGGCCGGAGGACCAGGTCGGCGAGCTCCACCCC
CCGGTCCGGTACGGCTCCACGCCGGACCCGGGACCCGTACGCCTCGCTCAGCGAGCTGTACGGCGAC
ATCCGCGAGGGCCTGCAACGGGTCCCCGAGTC

NE123a:

CTGTGGTCCATCCCCACCCACTCGGCCGGGACCGAGTACGTGCGGCGAGGCGAGTGGACGCCCGGGCAG
CTCCGGCTCATGTGCGGGACGGCGCCGGAGCCGCGACGGCGGCGTCCGGCAGGACCTGTTTCGCCGTC
GCCCCGAGGAAATGATCCACTTCTGCTGATCAACAACATCATCATGGCGACGGGTTCAGCCGTTCCACC
TGCCCCGGATCGACTTCGGCACGGTGAACGGCGAGCTCCCCGTCCCCTCGACCTGTGCCTGGAGCCCTTC
GGGCGGGGACGCTGCAACGGTTTCGCCGCTTGGAAACGGCCCTACGACCTGGTCCGCGACCTGGCCGCC

GACCGGCCACCGGTGACCGCGCGGACCCGTACCCCTACGGTTCGCTCAGCGAGCTGTACGGGGCCATC
CGCCAGGCGATCCAGGACATCCCGGACGTCTTTC

NE123b:

GCCTGGTCCATCCCGACGGCCGGTGCCGGGGCGGAACTCGTCCGCCGTGGCGAGTGGACCCCGGAGCAG
TTGCAGCTGGCCTGCGGGACGGCGGTCCGACGCTCGACTACGGGATGCGCGGCACGCTGCTCAACGTG
GCCCCGGAGGAGATGATCCACTTCTGTCATCAACAACATCATCACCGCCACCGGGGACAGCTTCCAC
CTGCCGGCGATCGACTTCGGCACCCCTCAACGAGCAGCTGCCGGTGCCCTGGACTTCAGCCTGGAGGGTT
CGGGATCGGGCCGTTGCAGCGGTTTCATCGCCATCGAGCAGCCGGACGACCTGACCGTCGAGTTCTCCGG
GACCGACACCCTGCTCGACCGGGGGCGACGCCATGTACCCGTACGGCTCGCTCAGCGAGCTGTACGCGC
GATCCGGGAGGCATCCGAGCGGGTGCCCGACC

NE125a:

GCCTTCTCCATCCCGGCCTACGGGGCCGGAACCGAGTACGTGCGTTCGTGGTGTGTGGACACCCGGGCAGT
TGCGGCTCGCTTGTGGCAGTGGCGGCGAGAACCAGGACGACGGCATGCGTGGCGCCCTGCTACCGTGG
CGCGTGAGGAGATGATCCACTTCTGTTGGTCAACAACATCATCATGGCCACCGGGGAGACGTTCTACGT
CCCGTCCATCGATTTCCGGACGATCAACAACGAGCTGCCGGTACCGCTGGACTTCTCCCTCGAACC GTTC
GGGATCGGCAGTGTCCAGCGGTTTCATCGCGATCGAGCAACC CGCGGGCCTGGCCGGGGAGATCCGGCGC
GGTGACGTCCGGGGTGACGACGCGGAGGTCGCCCGCACCCGTACGGCTCACTCAGCGAGCTCTACGCG
AGCATCCGGGAAGGTCTGTGCCGGGTGCCCGGT

NE125b:

GCCTACTCCATCCCGACGTACAGCGCGGGCCGGCGGCACGTGCGGCACGGGCTGTGGACGCCGGAGGAG
CTGTGGCTCGTGTGCGGGACGGCGGCGAGACGACGAGGGTGGCATGCGCGGCAGGCTGCTGACCGTC
GCCCCGAGGAGATGATCCACTTCTGTCATCAACAACATCATCATGGCGATCGGCGAGCCGTTCTACG
TGCCCGACGTGACTTCGGCACGGTTCGACGCGGTCCTGCCGGTGCCGCTCGACCTCGCGCTCGAGGCCTT
CGGCCTGGGCAGCGTGCAGCGGTTTCATCGCCCTCGAGCGGCCCGACGCGCAGGTCGGCGAGGTGCGCCG
CGCCGGCGCCGCCCGACCGCCGTGTCCGACGGAGCCGGCGCGCACGTA CTCCCCGCCAAGCGCGCGTGT
ACGCGACCGATCCGGTACGGGGCGGTGAAGGCGC

NE126a:

CTGTGGTCCGTCCCCACCTACTCGGCGGGGGTCCGCCACGTGCGAACCGGGCGAGTGGACGCCGGAGCAG
CTGCGCCTGATGTGCGGTGCGGGGCCGACGGCCTCGACGGCGGGCTCCGGGGGAGCCTGCTGGGCGTG
GCCCCGGAGGAGATGATCCACTTCTGTCATCAACAACCTGATCATGGCGACCGGCCAGCCCTTCCACC
TGCCCGACATCGACTTCGGCACGGTGAACAGCGTGTGCCCCGTCCCGGTCGACTTCTGCCTGGAGCCGTT
CGGCATCGCCGCCCTGCAGCGGTTTCATCGCCCTGGAGCGGCCGTACGGCCTGGTCCGGGACTTCGCGGG
GAGCGGCGGGAACGGCGGGCGGGTTCGGCGACGGCGCGCCGCGAGGGGGAACCGCACCGGTACGGCT
CGCTGAGCGAGCTGTACGCGGCCATCCGCCGGGCG

NE126b:

GCCTGGTCCGTCCCGACCCACGGCGCGGGCCGGCGAGTACGTGCGCCGGGGCGCCTGGACCCCCCGGAG
CTCCGGCTCGCCTGCGGGACGGGGGCCGACCGGTGACTACGGCGCGCGCGCCTGCTGCTCGACGTG
GCCCCGGAAGAGATGATCCACTTCTGTCATCAACAACCTGATCATGGCGACCGGCCAGCCCTTCCACC
CTGCCCGGATCGACTTCGGCACCCCTCAACGCCGAGCTGCCGATCCCGCTGGACCTCTGCCTGGAGGGGT
TCGGGATCGGGCGCTGCAACGCTTCATCGCCGTGCAACAGCCCCGACGGGCTGACGCCGGGGCTCGCCG
GGGACCGGCCGGCGGGCGGGTGGCACCCGTACGGTTCGCTCAGCGAGCTCTACGCCGCGATCCGCG
AGGCGGTACCCGGATCCCCGACCTGTTCTGTTGTC

NE130a:

GCCTGGTTCGATCCCGCTGTACGGGGCGGGCCGGGAACTGGTACGGCGCGGGGAGTGGAGCGCCCGGAG
CTCCGCCTGGCGTGCGGCGACGGCGGGGAGACCCTCAGCAACGGCATGCGCGGCAGCCTGATGAACGTG
GCCCCGGAGGAGATGACGCACTTCTGTCATCAACAACGTCATCACCGCGGGCGGACTTCCCTTCCACC
TGCCCGGATCGACTTCGGGACCGTCAACAACGGCTGCCGGTGCCGCTCGATGTGAGCCTGGAGGGTTC
GGGCTGGGCGAGCGTGCAGCGGTTTCATCGCCCTGGAGCAGCCGACGGACTGAGCCCCGACTCGCCGAGG
AGACCGAGAACGCCGCCCCCGGGACGAGCGGCGACCCGTACCCGTACCGTACCGTACCGTACCGGACCTGTACGC
CGGATCCGGGAGGGCATCGCCACCATC

NE130b:

GCCTACTCGATCCCGACCCACGGTGCGGCCCCGGGAGTTCGTCCGCCGCGGTCTGTGGACGCCCCGACCAG
GCCCCGGCTGGCGTGCGGGGACGGCGGCCGCAGCCGCGACGCCGGGGTGCGCGGCCGGCTGCTGGAGGT
GGCCCGGAAGAGATGATCCACTTCCTCGCCGTCAACAACATCGTCATGGCGATGGGGGAGGCGTTCCA
CGTCCCCGACATCGCGTTCGGCCGGCTCAACGCCACCCTGCCGGTGGCGATGGACTTCTGCCTGGAACCC
TTCGGGCTGGGCACCCTCCAGCGGTTTCATCGCGATCGAGCGGCCCTGGACGCGGCACCGCCCCGCTCA
GCGCGGTGTCCGAGGACCGCCGGCGATGCCGGGACCCCGCCGCTACCAGGGCATCCCGGAGTTGTACG
AGGAGATCCGGCAGGGCCTGCACCGCGTCCCCGAC