

**Figure S1. ClustalW alignment of 10 EPOR amino acid sequences**

<i>G. dobula</i>	MTKMRNSRLKIVF-MLCLAS--VTTGGQYFESKVAQLLQDETEDIKCFVEE-KDLTCFWE	56
<i>P. kaznakovi</i>	MTKMRNSRLKIVF-MLCLAF--VTTGGQYFESKVAQLLQDETEDIKCFVEE-KDLTCFWE	56
<i>S. gongshanensis</i>	MTKMRSDRLKIVF-MLCLAF--VTTGGQHFEVKVAQLLQDETEDIKCFVEG-KDLTCFWE	56
<i>S. nukiangensis Tsao</i>	MTKMRSDRLKIVF-MLCLAF--VTTGGQHFEVKVAQLLQDETEDIKCFVEG-KDLTCFWE	56
<i>S. prenati</i>	MTKMRSDRLKIVF-MLCLAF--VTTGGQHFEVKVAQLLQDETEDIKCFVEG-KDLTCFWE	56
<i>D. rerio</i>	MTKMRSDRLKIVF-MLCLAF--VTSGGKYFESKVAQLIRDETEDIKCFVEGKKDFTCFWE	57
<i>C. auratus</i>	MTKMRSNRLKIIIF-MLCLAC--VTTGGQHFEVKVAQLLQDETEHKCFVEG-KDLTCFWE	56
<i>C. carpio</i>	MTKM-SDRLKIVF-MLCVAF--VTTGGQHFEVKVAQLLQDETEDIKCFVEL-KDLTCFWE	55
<i>C. semilaevis</i>	MNAAHNRRAAALYALLCVV--SVLGGKTWEEKFSIMRRDPENPKCFAEDRMNLTCFWE	58
<i>E. lucius</i>	MTNDHLKKLLVYC-MLCCTQKIIIVQGAQTFQTKVALLRHVEPNNPKCFAGIRDLTCFWE	59
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<i>G. dobula</i>	[REDACTED]	116
<i>P. kaznakovi</i>	[REDACTED]	116
<i>S. gongshanensis</i>	[REDACTED]	116
<i>S. nukiangensis Tsao</i>	[REDACTED]	116
<i>S. prenati</i>	[REDACTED]	116
<i>D. rerio</i>	[REDACTED]	117
<i>C. auratus</i>	[REDACTED]	116
<i>C. carpio</i>	[REDACTED]	113
<i>C. semilaevis</i>	[REDACTED]	117
<i>E. lucius</i>	[REDACTED]	118
	FN3 super family domain (EPOR ligand binding domain)	
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<i>G. dobula</i>	DGRILYSRSLNAENLVLDPPRNLTVMSGKEQLNVSW[REDACTED]LYDS[I]YEVRYAVE	176
<i>P. kaznakovi</i>	DGRILYSRSLNAENLVLDPPRNLTVMSGKEQMNVSW[REDACTED]LYDS[I]YEVRYAVE	176
<i>S. gongshanensis</i>	DGRILYSRRLNAENFLLDPPRNLTVMSGKEQLNVSW[REDACTED]LYDS[I]YEVRYAVE	176
<i>S. nukiangensis Tsao</i>	DGRILYSRRLNAENFLLDPPRNLTVMSGKEQLNVSW[REDACTED]LYDS[I]YEVRYAVE	176
<i>S. prenati</i>	DGRILYSRRLNAENFLLDPPRNLTVMSGKEQLNVSW[REDACTED]LYDS[I]YEVRYAVE	176
<i>D. rerio</i>	DGQMLYNRSLNVENILLTDPPRNTVWSSGKEQLNVSWLPPAVKYIDDS[I]YEVRYAVE	177
<i>C. auratus</i>	DGQKLYTRSLNAENFLLDPPRNLTVLSSGKEQLNVSWLPPQLKYMDDSM[I]YEVRYAAE	176
<i>C. carpio</i>	DGRILYSRSLNAENYLFLDPPRNLTVLSSGKEQLNVSWLPPLLKYMDDSM[I]YEVRYAVE	173
<i>C. semilaevis</i>	KGLIHNRRTLFDVMFLLDPPANVTVRTEKQDQLNVSWAPKLKYMDDSMYQIRYSSA	177
<i>E. lucius</i>	DGLIHNRRLHIELFLLDPPVNLTVKRTEKQGQLKASWMPPLKYLDDSM[I]YEVSYVIA	178
	FN3 domain	
	. * : : * * : : * *** * : * : * : : * : * : : * : : * : :	
<i>G. dobula</i>	GSNMGKVEVIKVSTMLVLRLGLQSDTRYKVV[REDACTED]KPDGVSYKGWSAWTEPVFGVTPPSDV	236
<i>P. kaznakovi</i>	GSNMGKVEVIKVSTMLVLRLGLQSDTRYKVV[REDACTED]KPDGVSYKGWSAWTEPVFGVTPPSDV	236
<i>S. gongshanensis</i>	GSNMGKVEVMKVSTMLVLLGLQSDTRYKVV[REDACTED]KPDGVSYKGWSAWTEPVFGVTPPSDM	236
<i>S. nukiangensis Tsao</i>	GSNMGKVEVMKVSTRLVLLGLQSDTRYKVV[REDACTED]KPDGVSYKGWSAWTEPVFGVTPPSDM	236
<i>S. prenati</i>	GSNMGKVEVMKVSTMLVLLGLQSDTRYKVV[REDACTED]KPDGVSYKGWSAWTEPVFGVTPPSDM	236
<i>D. rerio</i>	DSHMKGKVEETKASTMLVLRLGLQPDTRYKVV[REDACTED]KPDGVSYKGWSSWTSPVIAVTPPGSM	237
<i>C. auratus</i>	GSDLGKVEVIKVSTMLVLGLGESDTRYKVV[REDACTED]KPDGVAYKGWSAWTEPVFGVTPPSDV	236
<i>C. carpio</i>	GSNMGKVEVIKVSTMLVLGLQSDTRYKVV[REDACTED]KPDGVAYKGWSAWTEPVFGVTQPSDL	233
<i>C. semilaevis</i>	DGLETQVTESLATEKILTNLRPGVKYEVQ[REDACTED]KLDGISYDGYWSAWSTGVFMETPPAEF	237
<i>E. lucius</i>	GTQMGQVEEVQASSEVILLGLQPGAKYSVR[REDACTED]KLDGVSYNGYWSAWTDPVLMATLPGDL	238
	FN3 domain	
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<i>G. dobula</i>	DPLIVLLALFIVLILCLLSLTGFLSQHKFLLKKLWPDIPTPEHKFPGLFSVYKGDFKEWM	296
<i>P. kaznakovi</i>	DPLIVLLVLFIVLILCLLSLTVFLSQHKFLLKKLWPDIPTPEHKFPGLFTVYKGDFKEWM	296
<i>S. gongshanensis</i>	DPLIVLLVLFIGLILCLLTTLTVFLSQHKFLLKKLWPDIPTPEHKFPGLFTVYKGDFKEWM	296
<i>S. nukiangensis Tsao</i>	DPLIVLLVLFIGLILCLLTTLTVFLSQHKFLLKKLWPDIPTPEHKFPGLFTVYKGDFKEWM	296
<i>S. prenati</i>	DPLIVLLVLFIGLILCLLTTLTVFLSQHKFLLKKLWPDIPTPEHKFPGLFTVYKGDFKEWM	296
<i>D. rerio</i>	DPLIVLLVVFIIILCLLSMTVILSHHKFLLKKLWPDIPTPEHKFPGLFTVYKGDFKEWM	297
<i>C. auratus</i>	DPLIVLLVLSIGLILCLLSLTVILSHHKFLLKKLWPDIPTPEHKFPGLFTVYKGDFKEWM	296
<i>C. carpio</i>	DPLIMVLVLFIGLILCLLSMTVILSHHKFLLKKLWPDIPTPEHKFPGLFTVYKGDFKEWM	293
<i>C. semilaevis</i>	DPLILCLTLIISFILLVFSLTILLSHRRFLTKKIWPVVPPTENKFPGFLFSVYGGDFQEWL	297
<i>E. lucius</i>	DPLIVSLTLIISLTLSSLTVLLSHRRFLLNKIWPVIPNPESKFQGLFTVYGGDFEEWL	298
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<i>G. dobula</i>	SQNNGYMWGRSVHVYTEELPSPLEVLSVLT----SHGTSQREERKPAEEEEKKPAEE	352
<i>P. kaznakovi</i>	SQNNGNMWMGRSVHVYTEELPSPLEVLSVLT----SHGTSQREERKPAEEEE-----	345
<i>S. gongshanensis</i>	SQNNGNMWMGRSVHVYTEELPSPLEVLSVLT----SHGTSQREERKPAEEEE-----	345
<i>S. nukiangensis Tsao</i>	SQNNGNMWMGRSVHVYTEELPSPLEVLSVLT----SHGTSQREERKPAEEEE-----	345
<i>S. prenati</i>	SQNNGNMWMGRSVHVYTEELPSPLEVLSVLT----SHGTSQREERKPAEEEE-----	345
<i>D. rerio</i>	SQNSGSMWARSVQMYTEELPSPLEVLSVLS----P----LDERKLVR-----	338
<i>C. auratus</i>	SHNNSGMWRSVHVYTEELPSPLEVLSVLT----SHDTSHREEVKPAEEDE-----	345
<i>C. carpio</i>	SQNNSSMWGRSVQVYTEELPSPLEVLSVLT----SHGTSQREERKPVVEDE-----	342
<i>C. semilaevis</i>	GHTRGGLWVAADYLFSERPSILLEVISEFRLGPPLVSPLPPKAPRGLTLGREE----N	352
<i>E. lucius</i>	GYSGAGGLCLRPAYFYSEELPAPLEVLSVSLDP--SSPLPPKAKQAPGED-----	347

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<i>G. dobula</i>	ERESQRSDSGLTDWRREPPQAHWLMQLRALQENPESRPQSALLQSHDTYVTLNQ---GD	409
<i>P. kaznakovi</i>	ERESERSDSGLTDWRREPPQAHWLMQLRALQENPESRPQSSLLQSHDTYVTLNQ---GD	402
<i>S. gongshanensis</i>	ERESERSDSGLTDWRQEPPQVHWLMQLRHLQENPESRSQSSILQSHDTYVTLNQ---GD	402
<i>S. nukiangensis Tsao</i>	ERESERSDSGLTDWRREPPQAHWLMQLRHLQENPESRSQSSILQSHDTYVTLNQ---GD	402
<i>S. prenati</i>	ERESERSDSGLTDWRREPPQAHWLMQLRHLQENPESRSQSSILQSHDTYVTLNQ---GD	402
<i>D. rerio</i>	-DEDQRSDSGLTE----PPH WLMEQLRALQENPESLRSRSTLLQSHDTYITLNH---SS	388
<i>C. auratus</i>	EKESERSDGSGLTDWRWPDPHPHAHWLMDQLRALQENPESHSQSSLQSHDTYVTLNQ---GD	402
<i>C. carpio</i>	EKESERSDGSGLTDWRREPPQAHWLMQLRALQENPQQLSQSLLVQSHDTYVTLNQNSQGD	402
<i>C. semilaevis</i>	QEVLERSASTV-----HDPWLMDRV-----LHQ TALLESQDPYVTLSN---PP	393
<i>E. lucius</i>	-EKLKRID-SLMERWRETPHEHWLMDQLRVLHQLPGP--QCSQLESQDAYVTNAQN-HS	402

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<i>G. dobula</i>	IEQQVDDVFEETLPLQLTLFSTAGTSSLSTSHSDLGSLQSSGSGRLSSQSSFEYPNHTWP	469
<i>P. kaznakovi</i>	IEQQVDDVFEETLPLQLTLFSTAGTSSLSTSHSDLGSLQSSGSGRLSSQSSFEYPNHTWP	462
<i>S. gongshanensis</i>	NEQQVDDVFEETLPLQLTFTTAGTSSLSTSHSDLGSLQSSGSGRLSSQSSFEYPNHTWP	462
<i>S. nukiangensis Tsao</i>	NEQQVDDVFEETLPLQLTFTTAGTSSLSTSHSDLGSLQSSGSGRLSSQSSFEYPNHTWP	462
<i>S. prenati</i>	NEQQVDDVFEETLPLQLTFTTAGTSSLSTSHSDLGSLQSSGSGRLSSQSSFEYPNHTWP	462
<i>D. rerio</i>	GGQREDDVFEETLPLQLTFTSAGTSSLNASHSDLGSLRQSSASGRLSSQSSFEDPNHPWP	448
<i>C. auratus</i>	IEQQQADDVFEETLPLQLTFTSAGTPSLSISSHSDLGSLQSSGSGRLSSQSSLEYPNHTWP	462
<i>C. carpio</i>	GEREVDDVFEETLPLQLTFTTGTSLSTSHSDLGSLQSSGSGRLSSQSSFEYPNQTWP	462
<i>C. semilaevis</i>	DGEQVDELLEESLPLEVLFHP-GVTALCESHSDLGSGQSSGS-RMSSQSSFEYPNQAWT	451
<i>E. lucius</i>	GEEP LDDILEETLPLQALFASGRT--CSESHSDLGSLQSSGSGRLSTQSSFEYPNHTWP	460

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<i>G. dobula</i>	PKPGPYAYMAVADSGVSMDYSPMSSSRIAEGVRHSMYTNDYKNEIV--WPLSGQYVKSG	527
<i>P. kaznakovi</i>	PKPGPYAYMAVADSGVSMDYSPMSSSRIAELGRHSMYTNDYKNEIV--WPLSGQYVKSG	520
<i>S. gongshanensis</i>	PKPGPYAYMAVADSGVSMDYSPMSSSKIAELGRHRMYTNDYKNEFLHKWPLSGQYVKSG	522
<i>S. nukiangensis Tsao</i>	PKPGPYAYMAVADSGVSMDYSPMSSSKIAELGRHRMYTNDYKNEFLHKWPLSGQYVKSG	522
<i>S. prenati</i>	PKPGPYAYMAVADSGVSMDYSPMSSSKITELGRHRMYTNDYKNEIFLHNWPLSGQYVKSG	522
<i>D. rerio</i>	PKPGPYAYMAVADSGVSMDYSPMSSRIIEIGKRSFYANEYKNEIFGYKWPFSAQQVQSG	508
<i>C. auratus</i>	PKPGPYAYMAVADSGVSMDYSPMSSSRIAELGRHRIYTNDYKNEIFPHKWPPLSGQYIKSG	522
<i>C. carpio</i>	PKPGPYTYMAVADSGVSMDYSPMSSSRITELGRHRMYTNDYKNEIFLHKWPFLGEHNKSG	522
<i>C. semilaevis</i>	RKPGPYTYMAVADSGVSMDYSPMS--RVEDVGRVPVYANDYKNDIMSQKTLFLQRQRADV	509
<i>E. lucius</i>	PKPGPYTYMAVADSGVSMDYSPMSSRIDYMKGKVITYTNEYRNEIPAHROPIPFAF----	516

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<i>G. dobula</i>	SESYTGAVW 536
<i>P. kaznakovi</i>	SESYTGAVW 529
<i>S. gongshanensis</i>	SESCGPVW 531
<i>S. nukiangensis Tsao</i>	SESCGPVW 531
<i>S. prenati</i>	SESCGPVW 531
<i>D. rerio</i>	Y----- 509
<i>C. auratus</i>	----- 522
<i>C. carpio</i>	SESCMGPVW 531
<i>C. semilaevis</i>	DND----- 512
<i>E. lucius</i>	----- 516