

Engineering of chimeric polyketide synthases using SYNZIP docking domains

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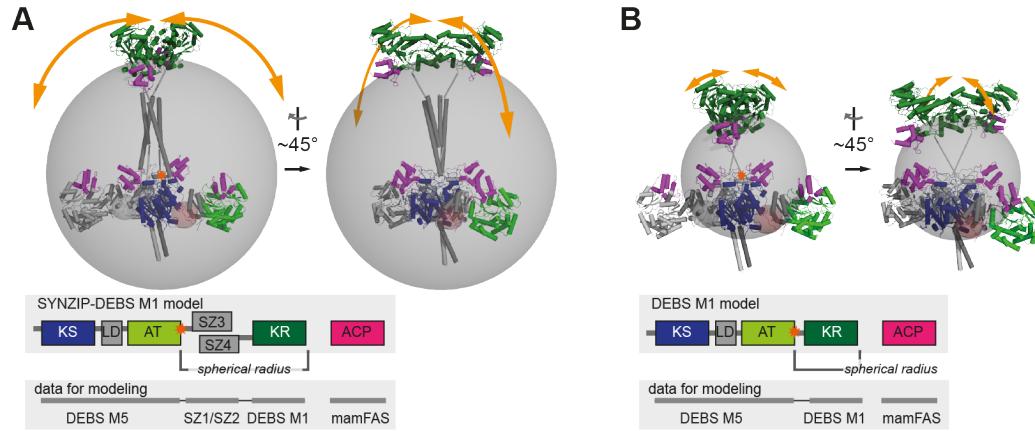


Figure S1. Structural model of DEBS M1 with and without SYNZIP docking domains. Structural models are based on X-ray crystallographic data of the KS-AT didomain of DEBS M5 (pdb 2HG4¹), of the KR domain of DEBS M1 (pdb 2FR0²), and of the coiled-coil pair SYNZIP1:SYNZIP2 (pdb 3HE5³). The coiled-coil pair SYNZIP3:SYNZIP4 was modeled regarding data provided by Thompson *et al.*⁴. Linkers connecting the domains are not shown, but dashed lines indicate distances bridged by linkers (linker lengths calculated with 3.6 Å per amino acid). ACP is shown in docking positions at KS, AT and KR as defined by Viegas *et al.* for mammalian FAS.⁵ Please note that the model shows only the docked ACPs, but does not contain the ACP C-terminally attached to the KR. The relative orientation of domains bridged by linkers is not defined, and the positions and orientations of those domains were arbitrarily chosen. A) SYNZIP-bridged M1 model. The coiled-coil pair induces an elongated structure. The SYNZIP3:SYNZIP4-KR-ACP part is presumably conformationally highly flexible, because of flexible linkers and missing dimerizing elements (flexibility indicated by orange arrows). The grey sphere indicates the maximum conformational space of the C-terminal ACP (linked to KR, see Figure 2A and B). For illustration of the conformational space (shown for one protomer of the dimer), the distance between the attachment site of the coiled-coil pair at the AT domain and the attachment site of the ACP at the KR domain was used as the radius of a sphere (shown in grey; center of sphere indicated by orange star), notwithstanding more complex structural properties, as for example induced by the KS-AT didomain that constrains a free motion of the SYNZIP3:SYNZIP4-KR-ACP part. The structural model illustrates that the ACP can essentially reach any position of the KS-AT didomain in any orientation, including a “translocation site” revealed in a previously reported study⁶. The translocation site is highlighted by a red sphere (grey for other protomer). Docking at the translocation site may allow for direct transacylation of the acyl moiety from KS to ACP, which is discussed as side reaction. B) M1 model. The model has been designed as described above, including construct specific properties. According to the model, the conformational space of ACP is limited, and access to the translocation site prevented.

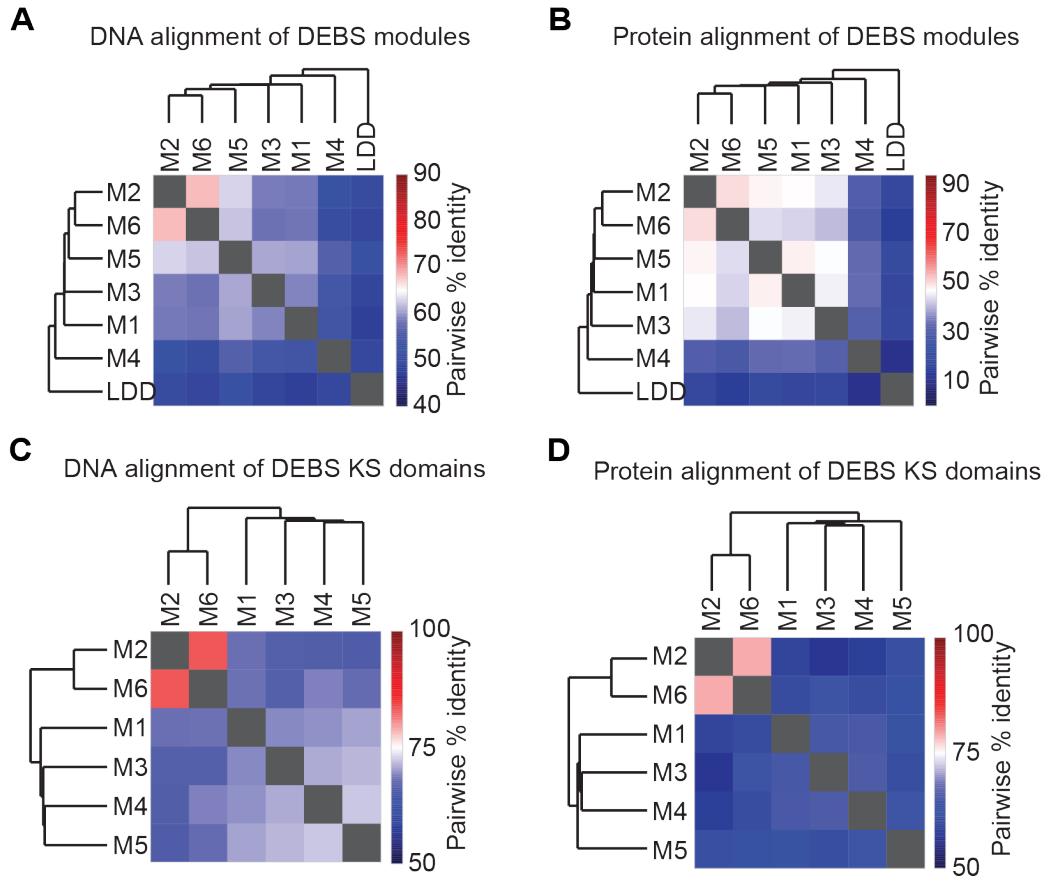


Figure S2. Sequence similarity analysis of DEBS modules and KS domains. Pairwise alignment of DNA and protein sequences of DEBS modules (A) and DEBS KS domains (B) are depicted as heatmaps.

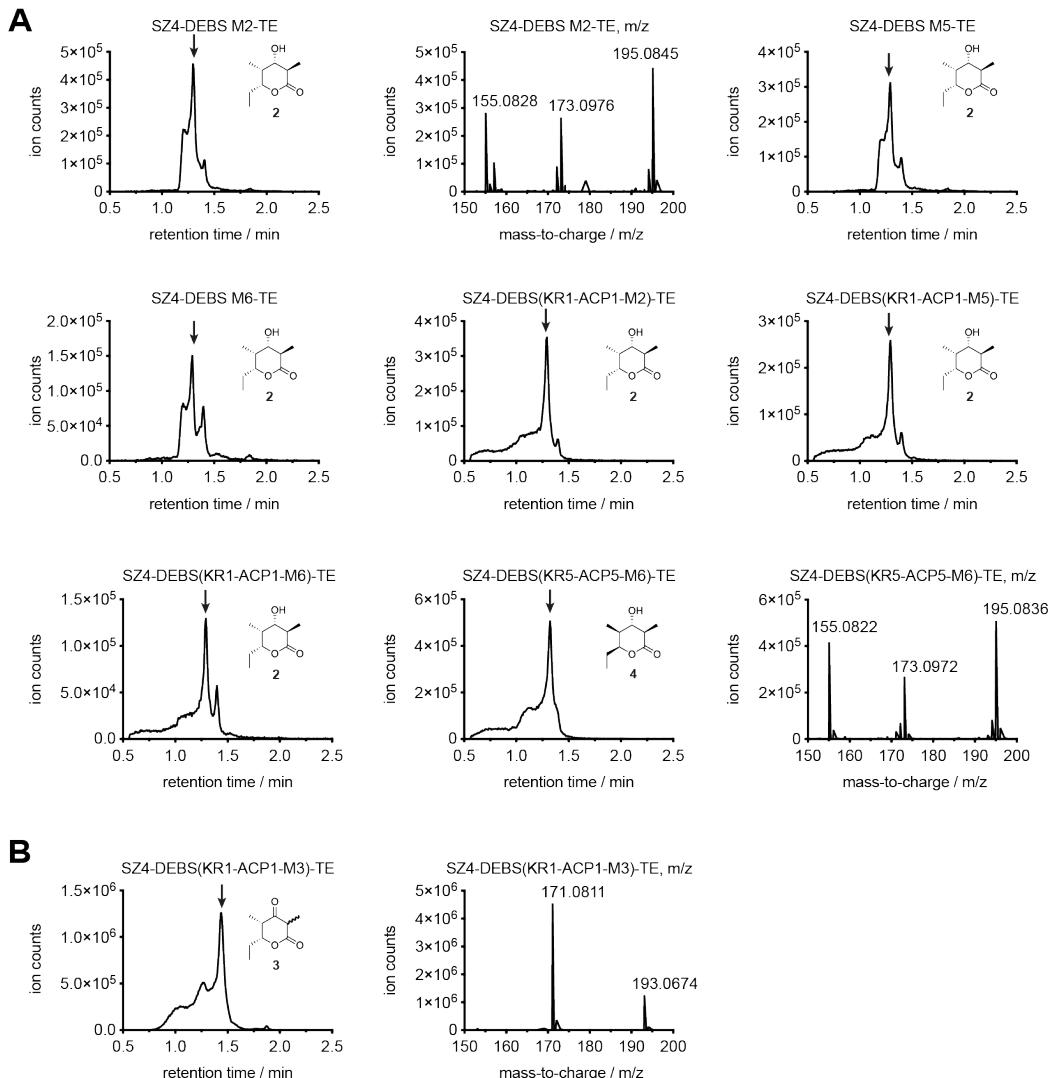


Figure S3. LC-MS analysis of products produced by bimodular chimeric PKSs newly generated in this study.
 A) Reduced triketide lactone products **2** and **4** ($C_9H_{16}O_3$, calculated MW 172.110) were detected in reaction mixtures containing SZ4-M2-TE, SZ4-M5-TE, SZ4-M6-TE, SZ4-KR1-ACP1-M2-TE, SZ4-KR1-ACP1-M5-TE, SZ4-KR1-ACP1-M5-TE, and SZ4-KR5-ACP5-M6-TE. B) Ketolactone **3** ($C_9H_{14}O_3$, calculated MW 170.090) was detected in reaction mixtures containing SZ4-KR1-ACP1-M3-TE. All PKS domain/modules are derived from DEBS. For all systems the extracted ion chromatograms were obtained by extraction of the $[M+Na]^+$ species and one chromatogram per compound is shown as an example. Labeled peaks from left to right correspond to $[M+H-H_2O]^+$, $[M+H]^+$, and $[M+Na]^+$ ions. The peak of interest is marked with an arrow based on its mass spectrum.

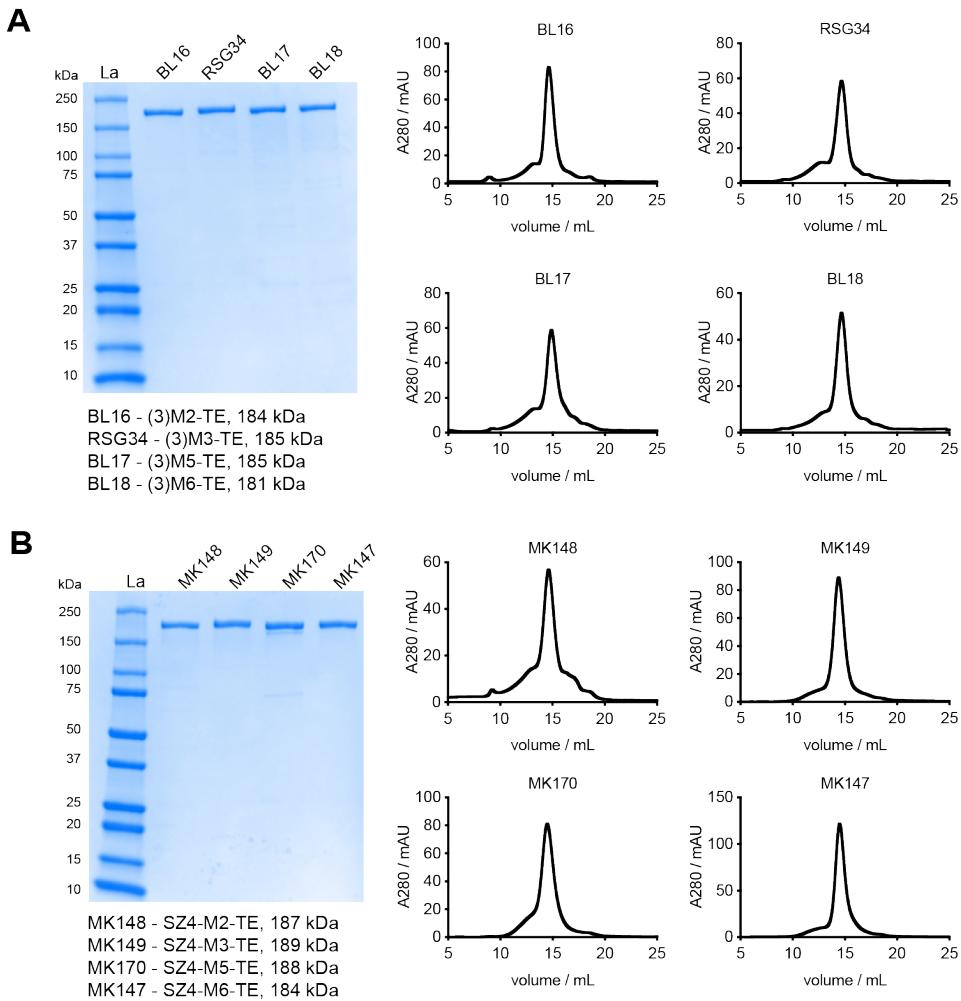


Figure S4. Analysis of protein purity by SDS-PAGE and SEC – single module acceptors. Acceptor proteins harboring a DEBS docking domain (A), and in comparison, a SZ domain (B). Protein abbreviations and their molecular weights (MW) are indicated. All proteins are pure and eluted in a predominately single peak from SEC.

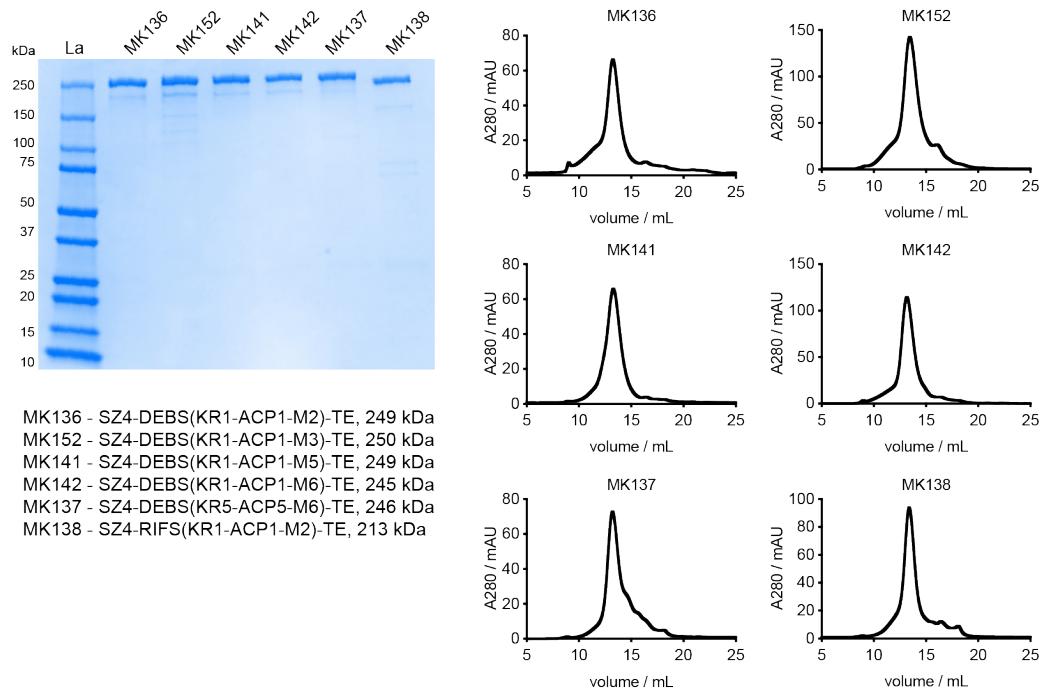


Figure S5. Analysis of protein purity by SDS-PAGE and SEC – covalent fusion acceptors. Acceptor proteins in which a KR-ACP fragment was fused to the acceptor module. Protein abbreviations and their MW are indicated. All proteins are pure and eluted in a predominately single peak from SEC.

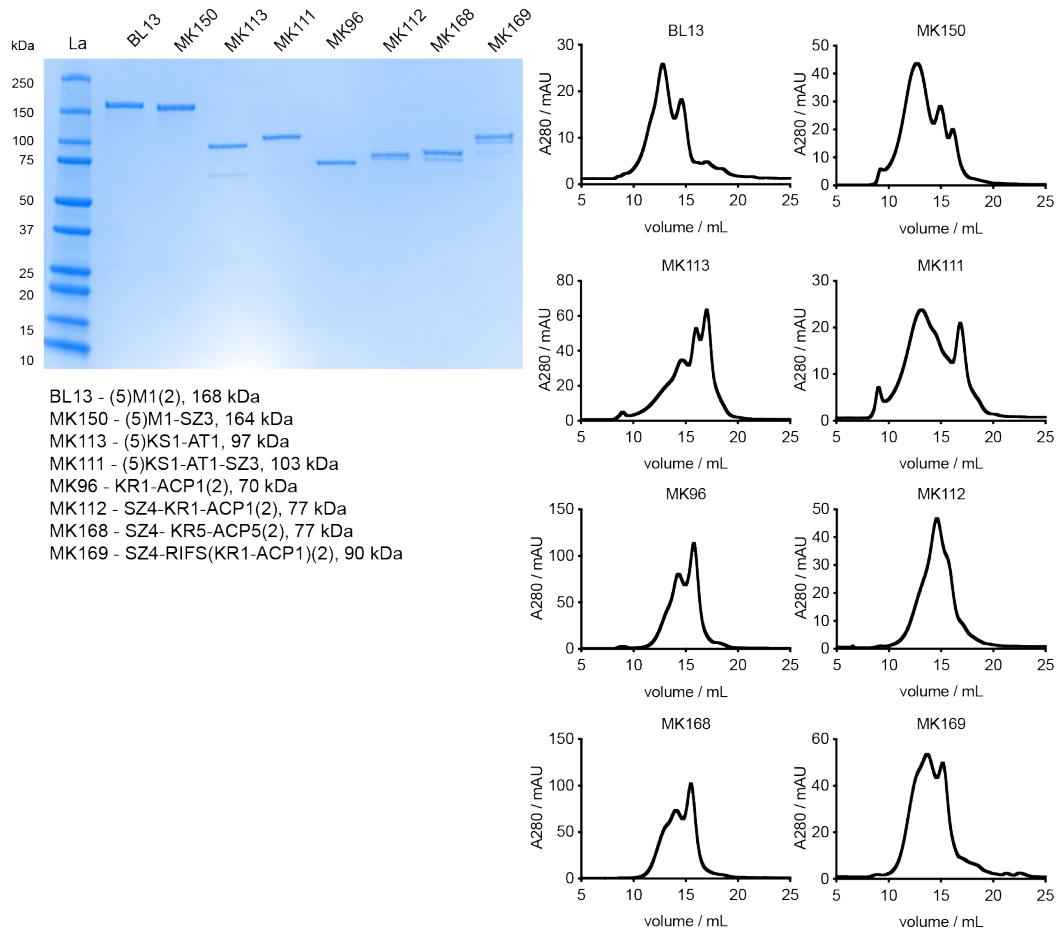


Figure S6. Analysis of protein purity by SDS-PAGE and SEC – DEBS M1 and its derivatives. DEBS M1 and its derivatives/fragments used in this study. If no indication is given the PKS domain/modules are derived from DEBS. Protein abbreviations and their MW are indicated. All proteins are pure, yet they appear as multiple oligomeric species based on SEC.

Table S1: Plasmids and primers used in this study. The cloning strategy is indicated for each construct. Some plasmids were only generated to function as intermediate constructs of the given cloning strategy and were not used for protein purification. If no indication is given the PKS domain/modules are derived from DEBS.

Plasmid Encoded Protein	Cloning Method	Cloning Fragments	Primer Name	Primer Sequence 5'-3'	Template
pMK80 - ACP1-M5-TE, used to generate pADD02	Infusion	80_V1	P-MK254	CGCCTGGCTGGCGGGAGGG	pBL17
			P-MK239	CATATGTATATCTCCTTCTAAAGTTAACAAAATTAA	
		80_V2	P-MK250	GAGCCGATCGCGATCGTCGGCATGGCGT	pBL17
			P-MK255	CACGCCCTGCGCGTACGCCTCGCC	
		80_V3	P-MK256	GCGTACGCCAGGGCGTG	pBL17
			P-MK253	CCCTCCGCCAGCCAGGCG	
		80_I	P-MK236	AAGAAGGAGATATACTATGCTGGCGTCGCTGCCG	pCK7
			P-MK242	CCGC GG GTGGCGCGCTCGAGCACCACCACCACTGAGATC	
pMK81 - ACP1-M6-TE, used to generate pMK93	Infusion	81_V1	P-MK257	GACCCGATCGCGATCGTCGGCATGGC	pBL18
			P-MK258	CACGCCGTGCACGTGCGCCCGG	
		81_V2	P-MK249	GCGCACGTGCACGGCGTG	pBL18
			P-MK239	CATATGTATATCTCCTTCTAAAGTTAACAAAATTAA	
		81_I	P-MK236	AAGAAGGAGATATACTATGCTGGCGTCGCTGCCG	pCK7
			P-MK246	CCGACGATCGCGATGGGTGACGGGGCCGTGGTC	
pMK83 - ACP1-M2-TE, used to generate pADD03	Infusion	83_V1	P-MK250	GAGCCGATCGCGATCGTCGGCATGGCGT	pBL17
			P-MK251	GACACCGCGCGTGTGCGCGTCGG	
		83_V2	P-MK252	CGCACACCGCGCGGTGTC	pBL16
			P-MK253	CCCTCCGCCAGCCAGGCG	
		83_V3	P-MK254	CGCCTGGCTGGCGGGAGGG	pBL16
			P-MK239	CATATGTATATCTCCTTCTAAAGTTAACAAAATTAA	
		80_I	P-MK236	AAGAAGGAGATATACTATGCTGGCGTCGCTGCCG	pCK7
			P-MK242	CCGC GG GTGGCGCGCTCGAGCACCACCACCACTGAGATC	
pMK91- (5)KS1-AT1, used as a template for other cloning strategies	Infusion	91_V	P-MK239	CATATGTATATCTCCTTCTAAAGTTAACAAAATTAA	pBL13
			P-MK262	TTTCCCGCGCTCGCGTACCGCCTCGAGCACCACCAC	
		91_I1	P-MK228	GTGCAGCCCGTGTGCGCGGTATGG	pBL13
			P-MK261	GTGGTGGTGGTGTGCTCGAGGCGGTAGCGCAGCGCGGAAACCTCGT	
		91_I2	P-MK229	CCATGACCGCGAACATCACGGCTGCAC	pBL13
			P-MK260	GAAGGAGATATACTATGAGCGGTGACAACGGCATGACCGAGGAAAA	

			G		
pMK93 - KR1-ACP1-M6-TE, missing an Asp in ACP1-KS2 linker	Infusion	93_V	digestion of pMK81 with NdeI		
		93_I	P-MK276	AAGGAGATATACATATGGACGAGGTTCCCGCGCTG	pAYC59
			P-MK277	AGCGACGCCAGCATATGCGGCCAACCGCGGTTTC	
pMK96 - KR1-ACP1(2)	Infusion	96_V	P-MK239	CATATGTATATCTCCTTCTTAAAGTTAACAAAATT	pBL13
			P-MK62	GGCACCGAGGTCCGGG	
		96_I	P-MK276	AAGGAGATATACATATGGACGAGGTTCCCGCGCTG	pBL13
			P-MK280	CCGGACCTCGGTGCCAGTTCGCGGCCAGGT	
pMK111 - (5)KS1-AT1-SZ3	Infusion	111_V	P-MK309	TCCGCCACCGGATCCGCCGAGCCAGACCGCGCTCGC	pMK91
			P-MK304	CTGGCACACAAAAAGCTCGAGCACCACCACAC	
		111_I	P-MK308	GGATCCGGTGGCGGATCCGGTAACGAAGTTACAACACTTGAGAATGAC	pfRSZ
			P-MK303	CTTTTGTGTGCCAGTCTATTCTCAAT	
pMK112 - SZ4-KR1-ACP1(2)	Infusion	112_V	P-MK239	CATATGTATATCTCCTTCTTAAAGTTAACAAAATT	pMK96
			P-MK311	TCCGGTGGCGGATCCGGTGACGAGGTTCCCGCGCTG	
		112_I	P-MK305	GGAGATATACATATGCAGAAAGTGGCTGAATTAAAAACAGA	pfRSZ
			P-MK310	GGATCCGCCACCGGATCCGCCCTCAGCAACATCGTTCTCCAATCTG	
pMK113 - (5)KS1-AT1 (shorter than pMK91)	Infusion	113_V	P-MK239	CATATGTATATCTCCTTCTTAAAGTTAACAAAATT	pMK91
			P-MK313	GAGCGCGTCTGGCTCCTCGAGCACCACCACAC	
		113_I	P-MK260	GAAGGGAGATATACATATGAGCGGTGACAACGGCATGACCGAGGAAAA	pMK91
			P-MK312	GAGCCAGACCGCGCTCGC	
pADD02 - KR1-ACP1-M5-TE, missing an Asp in ACP1-KS2 linker	Infusion	ADD02_V	digestion of pMK80 with NdeI		
		ADD02_I	P-MK276	AAGGAGATATACATATGGACGAGGTTCCCGCGCTG	pAYC59
			P-MK277	AGCGACGCCAGCATATGCGGCCAACCGCGGTTTC	
pADD03 - KR1-ACP1-M2-TE - missing an Asp in ACP1-KS2 linker	Infusion	ADD03_V	digestion of pMK83 with NdeI		
		ADD03_I	P-MK276	AAGGAGATATACATATGGACGAGGTTCCCGCGCTG	pAYC59
			P-MK277	AGCGACGCCAGCATATGCGGCCAACCGCGGTTTC	
pMK133 - KR1-ACP1-M2-TE, complete ACP1-KS2 linker	Quickchange	133_QC	P-MK347	GCGACCACGGCCCCCGTCATGAGCCGATCGCGATCGTC	pADD03
			P-MK348	GACGATCGCGATCGGCTCATCGACGGGGCCGTGGTCGC	

pMK134 - KR1-ACP1-M5-TE, complete ACP1-KS2 linker	Quickchange	134_QC	P-MK347	GCGACCACGGCCCCCGTCATGAGCCATCGCGATCGTC	pADD02
			P-MK348	GACGATCGCGATCGGCTCATCGACGGGGCCGTGGTCGC	
pMK135 - KR1-ACP1-M6-TE, complete ACP1-KS2 linker	Quickchange	135_QC	P-MK349	GCGACCACGGCCCCCGTCATGACCCGATCGCGATCGTC	pMK93
			P-MK350	GACGATCGCGATCGGCTCATCGACGGGGCCGTGGTCGC	
pMK136 - SZ4-KR1-ACP1-M2-TE	Infusion	136_V	digestion of pMK133 with NdeI		
		136_I	P-MK351	GAAGGAGATATAACATATGCAGAAAGTGGCTGAATTGAAAAAC	pMK112
pMK137 - SZ4-KR5-ACP5-M6)-TE	Infusion	137_V	P-MK247	GACCCGATCGCGATCGTC	pMK93
		137_V	P-MK239	CATATGTATATCTCCTTCTTAAAGTTAACAAAATTA	
pMK138 - SZ4-RIFS(KR1-ACP1-M2)-TE	Infusion	137_I1	P-MK351	GAAGGAGATATAACATATGCAGAAAGTGGCTGAATTGAAAAAC	pMK112
		137_I1	P-MK352	GCCGGTGGGGATGGGACCGGATCCGCCACCG	
		137_I2	P-MK353	CCCATCCCCACCGGGCG	pBL130
		137_I2	P-MK354	GATCGCGATCGGGTCGTCGGCATCCTCGGCAC	
pMK138 - SZ4-RIFS(KR1-ACP1-M2)-TE	Infusion	138_V	P-MK358	GAGATCGGCACCGCCCGGGCGAGGAGCCGATCGCGATCGTC	pAJ21
		138_V	P-MK239	CATATGTATATCTCCTTCTTAAAGTTAACAAAATTA	
pMK141 - SZ4-KR1-ACP1-M5-TE	Infusion	138_I1	P-MK351	GAAGGAGATATAACATATGCAGAAAGTGGCTGAATTGAAAAAC	pMK112
		138_I1	P-MK355	GGGCTCGGGCTCACGGATCCGCCACCG	
pMK142 - SZ4-KR1-ACP1-M6-TE	Infusion	138_I2	P-MK356	GAGCCCGCCGAGCCC	pAJ20
		138_I2	P-MK357	GGCGGTGCCGATCTGGCCGGCTCGCCGCCGATCCGAAGAGCTT GGCGCGCAG	
pMK141 - SZ4-KR1-ACP1-M5-TE	Infusion	141_V	digestion of pMK134 with NdeI		
		136_I	P-MK351	GAAGGAGATATAACATATGCAGAAAGTGGCTGAATTGAAAAAC	pMK112
pMK142 - SZ4-KR1-ACP1-M6-TE	Infusion	136_I	P-MK345	AGCGACGCCAGCATATGCACGCCACCGC	
		142_V	digestion of pMK135 with NdeI		
pMK146 - SZ4-MCS-H6, intermediate vector for various cloning strategies	Infusion	146_all	P-MK386	TGCCGCCCAAGCTTACCGGATCCGCCACCGG	pMK112
		146_all	P-MK387	AAGCTTGCAGGCCGCACTC	
pMK147 -	Infusion	147_V	digestion of pMK146 with HindIII		

SZ4-M6-TE		147_I	P-MK388	CGGATCCGGTAAGCTTGACCCGATCGCGATCGTC	pMK137
			P-MK389	GTGCAGGCCGAAGCTTCAATTCCCTCCGCCAG	
pMK148 - SZ4-M2-TE	Infusion	148_V	digestion of pMK146 with HindIII		
		148_I	P-MK390	CGGATCCGGTAAGCTTGAGCCGATCGCGATCGTC	pMK136
			P-MK389	GTGCAGGCCGAAGCTTCAATTCCCTCCGCCAG	
pMK149 - SZ4-M3-TE	Infusion	149_V	digestion of pMK146 with HindIII		
		149_I	P-MK391	CGGATCCGGTAAGCTTGACCCGATGCCATCGTC	pRSG34
			P-MK389	GTGCAGGCCGAAGCTTCAATTCCCTCCGCCAG	
pMK150 - (5)M1-SZ3	Infusion	150_V	P-MK394	ACGACCGCGGACCGGTT	pMK111
			P-MK389	GTGCAGGCCGAAGCTTCAATTCCCTCCGCCAG	
		150_I	P-MK392	GAACCGGTGCGGGTCTG	pBL13
			P-MK393	GCCGAGTTGGCGGGCC	
pMK152 - SZ4-KR1-ACP1- M3-TE	Infusion	152_V	P-MK478	GATGGCGATCGGGTCATCGACGGGGCCGTGG	pMK142
			P-MK153	AGCGGGACTCCCGCCC	
		152_I	P-MK398	GACCCGATGCCATCGTC	pRSG34
			P-MK399	GTCGAGCTGACTAGTGTGCTG	
pMK168 - SZ4-KR5- ACP5(2)	Infusion	168_V	P-MK402	GACGAGCCGCTCCAGGTA	pMK137
			P-MK291	TCGAGCTCCGTCGACAAGCT	
		168_I	P-MK457	CTGGAGCGGCTCGTGGCACCGAGGTCCGGG	pMK112
			P-MK458	GTCGACGGAGCTCGAATTGGATGCCGTGAGC	
pMK169 - SZ4-RIFS(KR1- ACP1)(2)	Infusion	169_V	P-MK459	GAAGAGCTGGCGCCAG	pMK138
			P-MK291	TCGAGCTCCGTCGACAAGCT	
		169_I	P-MK460	CGCGCCAAGCTTCCGGCACCGAGGTCCGGG	pMK112
			P-MK458	GTCGACGGAGCTCGAATTGGATGCCGTGAGC	
pMK170 - SZ4-M5-TE	Infusion	170_V	digestion of pMK146 with HindIII		
		170_I	P-MK390	CGGATCCGGTAAGCTTGAGCCGATCGCGATCGTC	pBL17
			P-MK389	GTGCAGGCCGAAGCTTCAATTCCCTCCGCCAG	

References for plasmids constructed elsewhere: pCK7⁷; pBP130⁸; pAYC59⁹; pAJ20 and pAJ21¹⁰; pBL13, pBL16, pBL17, and pBL18¹¹; pRSG34¹²; pfRSZ was a gift from Mislav Oreb, Goethe University Frankfurt. In addition pBL12 was used to produce LDD(4)¹¹.

Table S2: Amino acid sequences of proteins used in this study. If no indication is given the PKS domain/modules are derived from DEBS. SZ domains are shown in red, docking domains in green, and linker regions in gray. L₁₂ is the ACP1-KS2 linker used to covalently fuse ACP1 to a heterologous acceptor module.

Construct	Amino acid sequence
MK96 KR1-ACP1-(2)	MDEVSALRYRIEWRPTGAGEPARLDGTWLVAKYAGTADETSTAAREALESAGARVRELVVDARCGRDELAERLSRVGEV AGVLSLLAVDEAEPEEAPLALASLADTLSVQAMVSAELGCPLWTVTESAVATGPFERVRNAAHGALWGVGRVIALENPA VWGGLVDVPAGSVAELARHLAAVSGGAGEDQLRADGVYGRRWVRAAAPATDDEWKPTGTVLVTGGTGGVGGQIA RWLARRGAPHLLLVSRSGPADGAGELVAELEALGARTTVACDVTDRSREVRELLGGIGDDVPLSAVFHAAATLDDGTV TLTGERIERASRAKVLGARNLHELTRELDLTAFVLFSSASAFAFGAPGLGGYAPGNAYLDGLAQQRSSDGLPATAVAWTW AGSGMAEGPVADRFRRHGVIEMPPETACRALQNALDRAEVCPVIDVRWDRFLAYTAQRPTRLFDEIDDARRAAPQAAA PRVGALASLPAPEREKALFELVRSHAAAVLGHASAERVPADQAFELGVDSLSALELRNRLGAATGVRPLTTVFDHPDVR TLAAHAAEL GTEVRGEAPSALAGLDALEAALPEVPATEREELVQRLERMLAALRPVAQAADASGTGANPSGDDLGEAGV DELLEALGRELDGDPN SSVDKLAAALEHHHHHH*
MK111 (5)KS1-AT1-L-SZ3	MSGDNGMTEEKLRRYLKRTVTELDSVTARLREVEHRAPEPVAVVAMACRLPGGVSTPEEFWELLSEGRDAVAGLPTDRG WDLDSLFHPDPTRSGTAHQRGGGFLTEATAFDPAFFGMSREALAVDPQQRLMLELSWEVLERAGIPPTSLQASPTGVFVGL IPQEYGPRLAEGEGEGVEGYLMTGTTSVASGRIAYTLGLEGPVDTACSSLVAVHLCQSLRRGESSLAMAGGVTVMP PGMLVDFSRMNSLAPDGRCKAFSAGANGFGMAEGAGMLLERLSDARRNGHPVLAVLRTAVNSDGASNGLSAPNGRAQ VRVIQQLAESGLGPADIDAIVEAHGTGTRLGDPIEARALFEAYGRDREQPLHLSVKSNLGHTQAAAGVAGVIKMLAMR AGTLPTLHASERSKEIDWSSGAISLLDEPEPWAGARPRAVGSSFGISGTNAHIIIEAPQVVEGERVEAGDVVAPWVLSA SSAEGLRAQAARLA AHLREHPGQDP RDIA YSLATGRA ALPHRA AFAPVDE SAAL RVL DGL ATGNADGAA VGT SRA QQRAV FVFPQGWQWAGMAV DLLTSPVFAA ALREC A DALE PHL DF E V IPFL RAE A AR QD A LSTER DV V QP VM F AV M VSL ASMWRAHGVEPAAVIGHSQGEIAAACVAGALS LDDA ARV VAL SRV IAT MPGNKG M ASIAA P AGE VR A RIG DR V EA AV N GPRSVVVAGDSDELDRLVASCTTECIRAKRLAVDYASHSSH VETIRDALHAELGEDFHPLPGFVPF STVGRWTQPDELDA GYWYRNLRRTVRFADA VR ALAEQGYRTFLEVAHPI LTAAIIEIGD GSGADLSAIHSRRGDGS LADFG E ALS RAFAAGVA VDWESVHLGTGARRVPLPTYPFQRERVWLGGSGGGSG NEVTTLENDAAF IENENAYLEKEIARLKEKAALRNRLAHKKL EHHHHHH*
MK112 SZ4-L-KR1-ACP1(2)	M QKV AELKNRVAVKLN RNEQLKNKVEELKNRNAYLKNE LATLENEVARLENDVAEGGSGGGSGDEV SALRYRIEWRPTG AGEPARLDGTWLVAKYAGTADETSTAAREALESAGARVRELVVDARCGRDELAERLSRVGEVAGVLSLLAVDEAEPEEAP LALASLADTLSVQAMVSAELGCPLWTVTESAVATGPFERVRNAAHGALWGVGRVIALENPAVWGLVDVPAGSVAELA RHLAAVSGGAGEDQLRADGVYGRRWVRAAAPATDDEWKPTGTVLVTGGTGGVGGQIARWLARRGAPHLLLVSRS PDADGAGELVAELEALGARTTVACDVTDRSREVRELLGGIGDDVPLSAVFHAAATLDDGTVDTLTGERIERASRAKVLGA RNLHELTRELDLTAFVLFSSASAFAFGAPGLGGYAPGNAYLDGLAQQRSSDGLPATAVAWTWAGSGMAEGPVADRFRRH GVIEMPPETACRALQNALDRAEVCPVIDVRWDRFLAYTAQRPTRLFDEIDDARRAAPQAAAEP RVGALASLPAPEREKAL FELVRSHAAAVLGHASAERVPADQAFELGVDSLSALELRNRLGAATGVRPLTTVFDHPDVRTLAHAAEL GTEVRGE APSALAGLDALEAALPEVPATEREELVQRLERMLAALRPVAQAADASGTGANPSGDDLGEAGVDELLEALGRELDGDPNS SSVDKLAAALEHHHHHH*
MK113 (5)KS1-AT1	MSGDNGMTEEKLRRYLKRTVTELDSVTARLREVEHRAPEPVAVVAMACRLPGGVSTPEEFWELLSEGRDAVAGLPTDRG WDLDSLFHPDPTRSGTAHQRGGGFLTEATAFDPAFFGMSREALAVDPQQRLMLELSWEVLERAGIPPTSLQASPTGVFVGL IPQEYGPRLAEGEGEGVEGYLMTGTTSVASGRIAYTLGLEGPVDTACSSLVAVHLCQSLRRGESSLAMAGGVTVMP PGMLVDFSRMNSLAPDGRCKAFSAGANGFGMAEGAGMLLERLSDARRNGHPVLAVLRTAVNSDGASNGLSAPNGRAQ

	VRVIQQALAESGLGPADIDAVEAHGTGTRLGDPIEARALFEAYGRDREQPLHLGSVKSNLGHTQAAAGVAGVIKMLAMRAGTLPTLHASERSKEIDWSSGAISLLDEPEPWPGARPRRAGVSSFGISGTNAHAIIEAPQVVEGERVEAGDVVAPWVLSASSAEGLRAQAARLA AHLREHPGQDPDRDIASLATGRAALPHRAAFAPVDESAALRVLDGLATGNADGAAVGTSRAQQRAVFVFPQGWQWAGMAVDLLTSPVFAAALRECACADEPHLDFEVIPFLRAEAARREQDAALSTERVDVVQPVMFAVMVSLASMWRAHGVEPAAVIGHSQGEIAAACVAGALSDDAARVVALRSVIATMPGNKGMASIAAPAGEVRARIGDRVEIAAVNGPRSVVVAGDSDELDRLVASCTECIRAKRALAVDYASHSSHETIRDALHAELGEDFHPLPGFVFFSTVTGRWTQPDELDAGYWYRNRLRTVRFADAVRALAEQGYRTFLEVAHPILTAIIIEIGDGSGLADSAIHSLRGDGLADFGEALSRAFAAGVAVDWEVHLLGTGARRVPLPTYPFQRERVWLLEHHHHH*
MK136 SZ4-L-KR1-ACP1-L₁₂-M2-TE	MQKVAELKNRVAVKLNRNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAE GGSGGGSGDEVSALRYRIEWRTGAGEPARLDGTWLVAKYAGTADETSTAAREALESAGARVRELVDARCGRDELAERLRSVGEVAGVLSLLAVDEAEPEEAPLALASLADTLSVQAMVSAELGPLWTVTESAVATGPFERVRNAAHGALWGVGRVIALENPAVWGLVDVPAGSVAELARHLAAVVGSGAGEDQLALRADGVYGRWRVRAAAPATDDEWKPTGTVLVTGGTGGVGGQIARWLARRGAPHLLVSRSGPDADGAGELVAELEALGARTTVACDVTDRRESVRELLGGIGDDVPLSAVFHAAATLDDGTVDTLTGERIERASRAKVLGA RNLHELTRELDLTAFVLFFFASAFGAPGLGGYAPGNAYLDGLAQQRSDGLPATAVAWGTWAGSGMAEGPVADRFRRH GVIEMPPETACRALQNLDRAEVCPVIDVRWDRFLAYTAQRPTRLFDEIDDARRAAPQAAEPRVGAHMLASLPAPEREKALFELVRSHAAAVLGHASAERVPADQAFaelGVDSLSELRNRLGAATGVRLPTTVFDHPDVRTLAALHAAELGGAT GAEQAA PATTAPVDEPIAIVGMACRLPGEVDSPERLWEITSGRDSAAEVPPDRGWPDELMASDAAGTRRAHGNFMAGA GDFDAAFFGISPREALAMPDQQRQAETTWEALESAGIPPETLRGSDTGVFVGMSHQGYATGRPRPEDGVGYLLTNTVASGRIAYVLEGPAUTVDTACSSLVALHTACGSLRDGDCGLAVAGGVSVMAGPEVFTFSRQGALSPDGRCKPFSDEADGFGLEGSAFVVLQRLSDARREGRRVLGVVAGSAVNQDGASNGLSAPSGVAQQRVIRRAWARAGITGADVAVVEAHGTGTRLGDPVEASALLATYGKSRGSSGPVLLGSVKSNIGHAQAAAGVAGVIKVLGLERGVVPPMLCRGERSGLIDWSSGEIELADGVREWSPAADGVRRAGVSAFGVSGTNAHVIIAEPPEPEPVPQPRRMLPATGVVPVVLARTGAALRAQAGRLADHLAAHPGIAPADVSWTMARARQHFEERAAVLAADTAEAVHRLRAVADGAVVPGVVTGSASDGGSVFVFPQGAQWEGMAREL LPVPVFAESIAECDAVLSEVAGFSVSEVLEPRPDAPSLERVDVVQPVLFAMVSLARLWRACGAVPSAVIGHSQGEIAAAVAGALSLEDGMRVVARRSRAVRAVAGRGMSLVRGGRSDVEKLLADD SWTGRLEVAAVNGPDAVVVAGDAQAAREFLEYCEGVGIRARAIPVDYASHTAHVEPVRDELVQALAGITPRRAEVFFFSTLTGDFLDGTEL DAGYWYRNLRHPVEFHSQVQALTDQGYATFIEVSPHPVLASSVQETLDDAESDAAVLGLTLERDAGDADRLTALADAHTRGAVADWEAVLGRAGLVDLPGYFQGKRFWLLPDRTTPRDELDGWFYRWDWTEVPRSEPAALRGRWLVVPEGHEEDGWTVEVRSALAEAGAEPEVTRGVGLVGDCAVGVSLLALEGDGAQVQLVRELDAEGIDAPLWTFTGAVDAGSPVARPDQAKLWGLQVASLERGPRWTGLV DLPMPDPELRGRLTAVLAGSEDQAVRADA VRARRLSPAHTATSEYAVPGGTILVTGGTAGLAEVARWLAGRGAELALVSRGPDTEGVGLTAELTRLGARVSVHACDVSSREPVELVHGLIEQGDVVRGVVHAAGLPOQVAINDMDEAAFDEVVAAKAGGAVHLDELCSDAELFLFSSGAGVWGSARQGAYAAGNAFLDAFARHRRGRLPATSAWGLWAAGGMTGDEEAVSFLRERGVRAAMPVPRALAALDRVLASGETAVVVDWPAFAESYTAARPRPLLDIVTTAPSERAGEPETESLRDLAGLPRAERTAELVRLVRTSTATVLGHDDPKAVRATTPFKELGFDLSA VRLRNLLNAATGLRLPSTLVFDHPNASAVAGFLTSELGSGTPAREASSALRDGYRQAGVSGRVRSYLDLLAGLSDFREHDGSDGFSLDLVDMADGPGEVTVICCAGTAAISGPHEFTRLAGALRGIAPVRAVPQPGYEEGEPLPSSMAAAVAVQADAVIRTQGDKPFVVAAGHSAGALMAYALATELLDRGHPPRGVVLIDVYPPGHQDAMNAWLEELTATLFDRETVMDDTRLTALGAYDRLTGQWRPRETGLPTLLVSAGEPMGPWPDDS WKPTWPFEHDTVAVPGDHFTMVQEHAADIARHIDAWLGGGNSSVDKLAALHHHHHH*
MK137 SZ4-L-KR5-ACP5-	MQKVAELKNRVAVKLNRNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAE GGSGGGSGPIPTGGRARDEDWWRYQVVWREAESASLAGRVLLVTGPVSELSDAIRSGLEQSGATVLTCDVESRSTIGTALEAADTDLSTVVSLSRDGE

M6-TE	AVDPSLDALALVQALGAAGVEAPLWVLTRNAQVADGELVDPAQAMVGLGRVVGIEQPGRWGLVLDLVDADAASIRS LAAVLADPRGEEQVAIRADGIKVARLPAPARAARTRWSRGTVLVTGGTGGIAHVARWLARSGAEHLVLLGRRGADAP GASELREELTALGTGVTIAACDVADRARLEAVLAERAEGRTVSAMHAAGVSTPLDDLTEAEFTIADVVKVRTVNLD ELCPDLDASFVLFSSNAGVGSPGLASYAAANAFLDGFARRRSEGAAPVTSIAWGLWAGQNMADEGGYLRSQGLRAMD PDRAVEELHITLDHGQTSVSVMDMRRRFVELFTAARHRPLFDEIAGARAEARQSEEGPALAQRALSTAERREHLAHLIR AEVAAVLGHGDDAAIDRDRAFRDGLFDMSMTAVDLRNRLAAVTGVREAATVVFDPHTITRLADHYLERLVGAAEAEQAPA LVREVPKDADDPIAVGMACRFPGGVHNPGELWEFIVGGDAVTEMPTDRGWLDALFDPDPPQRHGTSYSRHGAFLDGAA DFDAAFFGISPREALAMDPPQQRQVLETTWELFENAGIDPHSLRGSDTGVFLGAAYQGYGQDAVVPEDSEGYLLTGNSAVV SGRVAYVLGLEGPATVDTACSSLVALHSACSLRDGDCGLAVAGGVSMAGPEVFTEFSRQGGLAVDGRCKAFSAEAD GFGFAEGVAVVLLQLRSLDARRAGRQVLGVVAGSAINQDGASNGLAAPSVAQQRVIRKAWARAGITGADAVVVAEHTG TRLGDPVEASALLATYGKSRGSSGPVLLGSVKSNIHGAAQAAAGVAGVIKVVLGLNRGLVPPMLCRGERSPLIEWSSGGVEL AEAVSPWPPAADGVRRAGVSAFGVSGTNAHVIIAEPPEPEPLPEPGPVGLAAANSVPVLLSARTETALAAQARLLESAVDD SVPLTALASALATGRAHLPTRAALLAGDHEQLRGQLRAVAEGVAAPGATTGTASAGGVVFVPGQGAQWEGMARGLLSV PVFAESIAECDAVLSEVAGFSASEVLEQRPDAPSLERVDVVQPVLSVMVSLARLGACGVSPSAVIGHSQGEIAAAVAVG VLSLEDGVRVVALRAKALRALAGKGGMVSLAAPGERARALIAPWEDRISVAAVNSPSSVVVSGDPEALALVARCEDEGV RAKTLPVDYASHSRHVEEIRETILADLDGISARRAAIPLYSTLHGERRDGADMGRYWDNLRSQVRFDEAVSAAVADGHA TFVEMSPHPVLTAAVQEIAADAVAIIGSLHRDTAEEHLIAELARAHVHGAVDWRNVFPAPPVALPNPFEPQRYWLAPEV SDQLADSRYRVDWRPLATTPDLEGGFLVHGSAPESLTSAVEKAGGRVVPVASADREALAAALREVGEVAGVLSVHTGA ATHLALHQSLGEAGVRAPLWLVTSAVALGESEPVDPEQAMVWGLGRVMGETPERWGLVLDLPAEPAPGDGEAFVACL GADGHEDQVAIRDHARYGRRLVRAPLGTRESSWEPAWTALVTGGTGALGHVARHLARCGVEDLVLVSRRGVDAAGAAE LEAELVALGAKTTITACDVADREQLSKLLEELRGQGRPVRTVVTAGVPESRPLHEIGELESVCAAKVTGARLLDECPDAE TFVLFSSGAGVWGSANLGAYSAANAYLDALAHRRRAEGRATSVAWGAWAGEGMATGDLEGLTRRGLRPMAPERAIRA LHQALDNGDTCVSIADVWERFAVGFTAAPRPLLDELVTPAVGAVPAVQAAPAREMTSQUELEFTHSHVAAILGHSSPDA VGQDQPFTELGFDSTAVGLRNQLQQATGLPATLFEHPTVRLADHGQQLDSGTPAREASSALRDGYRQAGVSGRVR SYLDLLAGLSDFREHFDGSDGFSDLVDMADGPGEVTICCACTAAISGPHEFTRLAGALRGIAVPAVPQPGYEEGEPLSS MAAAVAAVQADAVIRTQGDKPFVVAAGHSAGALMAYALATELLDRGHPPRGVVLIDVYPPGHQDAMNAWLEELTATLFDRE TVRMDDTRLTALGAYDRLTGQWRPRETGLPTLLVSAGEPMGPWPDDSWKPTWPFEHDTVAVPGDHFTMVQEHAIAIRH IDAWLGGNSSSVDKLAALEHHHHH*
MK138 SZ4-L-RIFS (KR1-ACP1-M2)-TE	MQKVAELKNRVAVKLNNEQLKNKVEEKNRNAYLKNELATLENEVARLENDVAEGGSGGGSGEPAEPASAGDPLLGT VSTPGSDRLTAVAQWSRRAQPWAVDGLVPNAALVEAAIRLGDLAGTPVVGELVVDAPVVLPRGSREVQLIVGEPEQRR RPIEVFSREADEPWTRHAHTLAPAAAAPVPEAAAGDATDVTVAGLRDADRYGIHPALLDAAVRTVVGDDLLPSVWTGVS LLASGATAVTVPTATGLRLTDPAQPVLTVESVRGTPVAEQGTTDALFRVDWPEIPLTAETADFLPYEATSAEATLSAL QAWLADPAETRLAVVTGCTEPGAAAIWGLVRSAQSEHGRIVLADLDDPAVLPAAVVASGEQPVVRVRNGVASVPRLTRVT PRQDARPLDPEGTVLITGGTGLALTARHLVTAHGVRLVLVSRRGEAPELQEELTALGASVIAACDVADRAQLEAVLR AIPAEHPLTAVIHTAGVLLDGVTTELTPDRLATVRRPKVDAARLLDELTREADLAFFLSSAAGVLGNPGQAGYAAANAE LDALARQRNSLDPAVSIAWGYWATVSGMTEHLDGADLRRNQRIGMSGLPADEGMALLDAAIATGGTLVAAKFDVAALR ATAKAGGPVPLRGLAPLPRRAAKTASLTERLAGLAETEQAAALLDLVRRHAAEVLGHSGAESVHSGRFTFKDAGFDST AVELRNRLAAATGLTLSLSPAMIFDYPKPPALADHLRAKLFDSAANRPAEIGTAAAEEPIAIVAMACRFPGGVHSPEDLWRLVA DGADAVTEFPADRGWDTDRLYHEDPDHEGTTYVRHAGFLDDAAGFDAFFGISPNEALAMDPPQQLLLETSWELFERAAI DPTTLAGQDIGVAGVNSHDYSMRMHRAAGVEGFRLTGGSASVLSGRVAYHFGVEGPATVDTACSSLVALHMAVQAL

	QRGECSMALAGGVMVMGTVETFVEFSRQRGLAPDGRCKAFADGADGTGWSEGVLGGLLVERLSEAQRRGHQVLA VVRGS AVNSDGASNLTAPNGPSQQRVIRKALAAAGLSTDVDAVEAHGTGTLGDPIEAEALLATYQQNRETPLWLSVKSNLG HTQAAAGVAGVIKMVMAMRHGVLPRTLHVDRPSSYVDWSAGA VELLTEARDWVSNGHPRRAVGSSFGIGGTNAHVLE EVAAPITTPQPEPAEFLVPVLVSARTAAGLRGQAGRLAAFLGDRTDVRVPDAAYALATTRAQLDHRAVVLASDRAQLCAD LAAFGSGVTGTPVDGKLA VLFTGQGSQWAGMGRELAEFTFPVFRDAFEAACEAVDTHLRERPLREVVFDDSA LLDQTMYT QGALFAVETALFRLFESWGVPGPLL A GHSIGELAAAHSVGVL DLA DAGE LVAARGRLMQALPAGGMVAVQATEDEVAP LLDGTVCVAAVNGPDSVLSGTEAAVLA VADELAGRGRKTRRLAVSHAFHSPLMEPM LDDFRAVAERLTYRAGSLPVVST LTGE LA ALDSPDYWVGQVRNAVRFSDAVTALGAQASTFELGP GGALAAMALGTLGGPEQSCVATLRKNGAEVPDVLT ALAELHVRGVGVWDTTVLDEPATAVGTVLPTYAFQHQRFWVDVDETAASVTPPPAEPIVDRPVQDVLELVRESAAVVLG HRDAGSF DLDRSFKDHGFDSLSAVKLRNRLRDFTGVELPSTLIFDYPNP AVLADHLRAELLSGT PAREASSALRDGYRQAGV SGRVRSYLDLLAGLSDFREHFDGSDGFSLDLVDMADGPGEVTVICCACTAISGPHEFTRLAGALRGIA PVRA VPQPGYEEG EPLPSSMAAAVAAVQADA VIRTQGD KPFVVAGHSAGALMAYALATELLDRGHPPRGVVLIDVYPPGHQDAMNAWLEELTA TLF DRET VRMDDTRLTALGAYDRLTGQWRPRTGLPTLLVSAGEPMGPWPDDSWKPTWPFEHDTVAVPGDHFTMVQEHA DAIARHIDA WLGGNSSSVKLA AALEHHHHH*
MK141 SZ4-L-KR1-ACP1- L₁₂-M5-TE	MQKVAELKNRVAVKLN RNEQLKNKVEELKNRNAYLKNE LATLENEVARLENDVAEGGSGGGSGDEVSALRYRIEWPTG AGEPARLDGTWL VAKYAGTADETSTAAREALESAGARVREL VVDARCGRDELAERLRSVGEAVGVL SLA VDEAEPEEAP LALASLADTLSVQAMVSAELGCPLWT VTESAVATGPFERVRNAAHGALWGVGRVIALENPAWGGLV DVPAGSVAELA RHLAAV VSGGAGEDQLALRADGVY GRRWVRAAAPATDDEWKPTGT VLTGGTGGVGGQIARWLARRGAPHLLL VSRSG PDADGAGELVAE LEALGARTTVACADVTDRESVRELLGGIGDDVPLSAV F HAAATLDDGTVDLTGERIERASRAKVLGA RNLHELTRELDLTAFVLFSSFASAFGAPGLGGYAPGNAYLDGLAQQRSSDGLPATAV AWTWAGSGMAEGPVADRFRRH GVIEMPPETACRALQN ALDRAEVCPVIDV RWDRFLLAYTAQRPTRLFDEIDDAR RAAPQAAEPRVGAHMLASLPAPERE KALFELVRSHAAAVLGHASAERVPADQAF AELGVDSLSALELRNRLGAATGVRLTTVFDHPVRTLA AHLAAELGGAT GAEQAAPATTAPVDEPIAIVGMACRPGDVDSPE SFWEVSGGGD AIAEAPADRGWEPDPDARLGGMLAAAGDFDAGFFGI SPREALAMPQQRIMLEISWEALERAGHDPVSLRG SATGVFTGVGTVDYGP RPDEAPDEVLGYVGTGTASSVASGRVAYCL GLEGPA MVTDTACSSGLTALH LAMESLRRDECGLALAGGVTVMS PGAFTEFRSQGGLAADGRCKPFSKAADGFGLAEGA GVLVLQRLSAARREGRPVLA VLRGSAVNQDGASNGLTAPSGPAQQRVIRRALENAGV RAGDV DYVEAHGTGTRLGDPIEV HALLSTYGAERDPDDPLWIGSVKSNIGHTQAAAGVAGVMKA VLALRHGEMPR TLHFDEPSPQIEWDLGAVSVSQARSWP AGERP RRAVGSSFGISGTNAHVIVEAPEADEPEPAPD SGPVPLVLSGRDEQAMRAQAGRLADHLAREPRNSLRTGFTLAT RRSAWEHRAVVVGDRDDALAGLRAVADGRIADRTATGQARTRRGVAMVFPQGAQWQMARDLLRESQVFADSIRDCE RALAPHVDWSLTLDLSGARPLDRV DVVQPALFAVMVSLAALWRSHGVEPAAVVGH SQGEIAAAHVAGALTLEDAKLVA VRSRV LRLGGQGGMASFGLCTEQAAERIGRFAGALSIA SVNGPRSVV VAGESGPLDELIAECEAEGITARRIPDVYASHSPQ VESLREELLTELAGISPVSADVALYSTTGQPIDTATMDTAYWYANLREQVRFQDATRQLAEAGFDAFVEVSPHPVLTVGIE ATLDSALPADAGACVVGTLRRDRGGLADFHTALGEAYAQGVEDWSPA FADARPVELPVYFQRQRYWLPI TGGRARDE DDDWRYQVVWREAWE SASLAGRVLLVTGP GPVSELS DAIRSGLEQSGATVLTCDVESRSTIGTALEAADT DALSTVVSLL SRDGEAVDPSLDALALVQALGAAGVEAPLWV LTRNAVQVADGELVDPAQAMVGGLGRVVGIEQPGRWGGLV DAD AASI RSLA VLA VLPD RGEQ VAI RADGI KVARL VPAPARAARTRWSPRGTVL VTGGTGGIGAHVARWLARSGAEHLVLLGRR GADAPGASELREELTALGTGVTIAACDVADRARLEAVLA AERAEGRTVSA VMHAAGVSTSTPLDDLT EA EFTIADVKVRG TVNLDELCPDLD A VFVLFSSNAGVWGSPGLASYAAANAFLDGFARRRSEGAPVTSIAWGLWAGQNMAGDEGG EYLRSQG LRAMD PDR AVEELHITLDHGQTSVSVVMDRRRFVELFTAARHRPLFDEIAGARA EARQSEEGP ALA QR LAA STAER REH LAHLIRAEVAAVLGHGDDAIDRDR AFRDLGFD SMTA VDLRNRLAA VTGVREAATVVF DHTITRLADHYLERL VSGTPA

	REASSALRDGYRQAGVSGRVRSYLDLLAGLSDFREHFDGSDGFSSDLVDMADGPGEVTVICCAGTAISGPHEFTRLAGALRGIA PVGRLAAVLEELTATLFDRETVMDDTRLTALGAYDRLTGQWRPRETGLPTLLVSAGEPMGPWPDDSWKPTWPFEHD TVAVPGDHFTMVQEHAIAHIDAWLGGGNSSSVDKLAAALEHHHHHH*
MK142 SZ4-L-KR1-ACP1-L₁₂-M6-TE	MQKVAELKNRAVKLNNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAEGGSGGGSGDEV SALRYRIEWRPTG AGEPARLDGTWLVAKYAGTADETSTAAREALESAGARVRELVDARCGRDELAERLSVGEAVGVL SLLAVDEAEPEEAP LALASLADTLSLVQAMVSAELGCPLWTVTESAVATGPFERVRNAAHGALWGVGRVIALENPAVWGLVDVPAGSVAELA RHLAAVVSGGAGEDQLALRADGVYGRWRVRAAAPATDDEWKPTGTVLVTGGTGGVGGQIARWLARRGAPHLLLVSRS PDADGAGELVAELEALGARTTVACDVTDRS VRELLGGIGDDVPLSAVFHAAATLDDGTVDLTGERIERASRAKVLGA RNLHELTRELDLTAFVLFFFASAFGAPGLGGYAPGNAYLDGLAQQRSDGLPATAVAWGTWAGSGMAEGPVADRFRRH GVIEMPPETACRALQNALDRAEVCPVIDVRWDRFLAYTAQRPTRLFDEIDDARRAAPQAAAEPRVGAHMLASLPAPERE KALFELVRSHAAAVLGHASAERVPADQAFELGVDSL SALELRNRLGAATGVRLPTTVFDHPDVRTLA AHLAAELGGAT GAEQAAPATTAPVDPDPIAVGMACRFPGGVHNPGELWEFIVGGDAVTEMPTDRGWDL DALFD PDPQRHGTSYSRHGAFL DGAADF DAFFGISPREALAMD PQQ RQV LETT WELFENAGIDPHSLRGSDTGVFLGAAYQGYGQDAV VPEDSEGYLLTG SSAVVSGRVAYVLGLEGPAVTVDTACSSLVALHSACGSL RDGDCGLAVAGGV SVMAGPEVFT EFSRQGGLAVDGRCKAF SAEADGFGFAEGVAVVLLQRLSDARRAGRQVLGVVAGSAINQDGASNGLAAPSGVAQQRVIRKA WARAGITGADV AVVE AHTGTRLGD PVEAS ALLATY GKS RGSS GPVLLGSVKS NIGH AQAAGVAGVIKVVLGLNRGLVPPMLCRGERSPLIEWSS GGVELAEAVSPWPPAADGVRRAGVSAFGVSGTNAHVIIAEPPEPEPLPEPGPVGLAAANSVPVLLSARTETALAAQARLLE SAVDDSVPLTALASALATGRAHLPRAALLAGDHEQLRQLRAVAEGVAAPGATTGTTASAGGVVFVPGQGAQWEGMAR GLLSVPVFAESIAECDAVLSEVAGFSASEVLEQRPDAPS LERVDV VQPVLFSVMVSLARLWGACGVSPSAVIGH SQGEIAAA VVAGVLSLEDG VRVVAL RAK AL RALAGKGGMVSLA PGERARALIAPWEDRISVAAVNSPSSVVSGDPEALAE LVARCE DEGVRAKTL PV DYASHSRHVEEIRETILADLDG ISARRA AIPLYSTLHGERRDADM GPR YWYDNLSQRFDEAVSAAVA DGHATF VEMSPHPVLTAAVQEIAADAVAIGSLHRDTAEELHIAELARA HVHG VAVDWRNVFP APPVALPNYPFEPQRYW LAPEVSDQLADS RYRVDWRPLATTPDLEGGFLVHGSAPESL TSAVEKAGGRVVPV ASADREALAAALREV PG EVAGVLS VHTGAATHLALHQSLGEAGVRAPLWL VTSRAVALGESEPVDPEQAMVWGLGRVMGETPERWGLV DLP AEPAPGDGE AFVACLGADGHEDQVAIRDHARYGRR LVRAPL GTRESSWE PAGTAL V TGGT GALGGHVARHLARCGVEDLVLV SRRGV APGAAELEAELVALGAKTTITACDVADREQLSKLLEELRGQGRPV RTV VHTAGV PESRPLHEIGELESVCAAKVTGARLL ELCPDAETFVL FSSGAGVWGSANLG AYSAANAYLDALAHRRRAE GRAAT SVAWGA WAGE GMAT GDLEG LTRR GLRPMA PERAIRALHQALDNGDTCV SIADVDWERFAVGFTAARPRPLL DELVTPAVGAVPAVQAAPAREMTS QELF THSHVAAIL GHSSPDAVGQDQPFTELGFDSL TAVGLRNQLQQATGLALPATLV FEHPTV RRLADHIGQQLDSGT PAREASSALRDGYRQA GVSGRVRSYLDLLAGLSDFREHFDGSDGFSSDLVDMADGPGEVTVICCAGTAISGPHEFTRLAGALRGIA PVGRLAAVLEELTATLFDRETVMDDTRLTALGAYDRLTGQWRPRETGLPTLLVSAGEPMGPWPDDSWKPTWPFEHD TVAVPGDHFTMVQEHAIAHIDAWLGGGNSSSVDKLAAALEHHHHHH*
MK147 SZ4-L-M6-TE	MQKVAELKNRAVKLNNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAEGGSGGGSGKLDPIAVGMACRPG GVHNPGELWEFIVGGDAVTEMPTDRGWDL DALFD PDPQRHG TSYSRHGAFL DGAA FD A AFFG I SPRE ALAMD PQQ RQV LETT WELFENAGIDPHSLRGSDTGVFLGAAYQGYGQDAV VPEDSEGYLLTGNSA VSGRVAYVLGLEGPAV TVDTACSS LVALHSACGSL RDGDCGLAVAGGV SVMAGPEVFT EFSRQGGLAVDGRCKAF SAEADGFGFAEGVAVVLLQRLSDARRAG RQVLGVVAGSAINQDGASNGLAAPSGVAQQRVIRKA WARAGITGADV AVVEAHGTGTRLGD PVEAS ALLATY GKS RGSS GPVLLGSVKS NIGH AQAAGVAGVIKVVLGLNRGLVPPMLCRGERSPLIEWSSGGV LAEAVSPWPPAADGVRRAGVSAF

	GVSGTNAHIIIAEPPEPEPLPEPGPVGVLAANSVPVLLSARTETALAAQARLLESAVDDSVPLTALASALATGRAHLPRRA ALLAGDHEQLRGQLRAVAEGVAAPGATTGTASAGGVVFVPGQGAQWEGMARGLISVPFAESIAECDAVLSEVAGFSAS EVLEQRPDAPSLERVVVQPVLFSVMVSLARLGACGVSPSAVIGHSQGEIAAAVVAAGVLSLEDGVRVALRAKALRALA GKGGMVSLAAPGERARALIAPWEDRISVAAVNSPSSVVSGDPEALAEVARCEDEGVRAKTLVDYASHSRHVEEIRETIL ADLDGISARRAIAPILYSTLHGERRDGADMGRPYWYDNLRSQRFDEAVSAAVADGHATFVEMSPHPVLTAAVQEIAADAV AIGSLHRDTAEEHLIAELARAHVGVADWRNVFPAPPALPNYPFEPQRYWLAPEVSDQLADSRYRVDWRPLATPPVD LEGGFLVHGSAPESLTSAVEKAGR VVPVASADREALAAALREVPGEVAGVLSVHTGAATHLALHQSLGEAGVRAPLWL TSRAVALGESEPVDPEQAMVWGLGRVMLET PERWGGLVDPAPGDGEAFVACLGADGHEDQVAIRDHARYGRRLV RAPLGTRESSWEPA GTALVTGGTGALGGHVARHLARC GVEDLVLVSRRGDAPGAAELEAEVALGAKTTITACDVADRE QLSKLLELRGQGRPVRTV VHTAGVPESRPLHEIGELESVCAAKVTGARLLDECPDAETFVLFSSGAGVWGSANLGAYSA ANAYLDALAHRRRAEGRATSAWGA WAGEGEMATGDLEG LTRRGLRP MAPERAIRALHQALDNGDTCVSIADVDWERF AVGFTAARPRPLLDELVTPAVGAVPAVQAAPAREMTSQUELLEFTSHVAAILGHSSPD AVGQDQPFTELGFDSL TAVGLRN QLQQATGLALPATLVFEHPTV RRLADHIGQQLDSGPAREASSALRDGYRQAGVSGRVRSYLDLLAGLSDFREHFDGSDGF SLDLVDMADGPGEVTVIC CAGTA A ISGPHEFTRLAGALRGIA PVRAV P QPGYEEGEPLPSSMAA VAAVQADAVIRTQGDKP FVVAGHSAGALMAYALATELLDRGHPPRGVVLIDVYPPGHQDAMNAWLEELTATLFDRETVMDDTRLTALGAYDRLTG QWRPRETGLPTLLVSAGEPMGPWPDDSWKPTWPFEHDTVAVPGDHFTMVQEHA DAIARHIDA WLGGGNSKLAALEHHH HHH*
MK148 SZ4-L-M2-TE	MQKVAELKNRAVKLN RNEQLKNKVEELKNRNAYLKNE LATLENEVARLENDVAEGGSGGGSKLEPIAIVGMACRLPG EVDSPERLWELITSGRDSAEEVPDDR GWPVDELMASDAAGTRRAHGNFMAGAGDFDAAFFGISPREALAMD PQQRQALET TWEALESAGIPPETL RGSDTGVFGMSHQGYATGRPRPEDGVDG YLLTGNNTASVASGRIA YVLGLEGPALTVDTACSSLV ALHTACGSLRDGDCGLAVAGGV SVMAGPEVFTESRQGALSPDGRCKPFSDEADG FGLGEGSAFVVLQRLSDARREGRRV LGVVAGSAVNQDGASNGLSAPSGVAQQRVIRRAWARAGITGADVAVVEAHGTGTRLGDPVEASALLATYGKSRGSSGPV LLGSVKS NIGHAQAAAGVAGVIKVLLGLERGVVPPMLCRGERS GLIDWSSGEIELADGVREWSPAADGV RAGVSAFGVS GTNAHIIIAEPPEPEPVQPRRMLPATGVVPVLSARTGAALRAQAGRLADH LAHHPGIAPADVSWTMARARQHFEERA VLAADTAEAVHRLRAVADGAVVPGVVTGSASDGGSVFVPGQGAQWEGMARELLPVPVFAESIAECDAVLSEVAGFSVSE VLEPRPDAPSLERVVVQPVLFAVMVSLARLWRACGAVPSA VIGHSQGEIAAAVVA GALSLEDGM RVVARRSRAVRA GRGSMLS VRGGRSDVEKL LADDSWTGRLEVAAVNGPD AVV VAGDAQAAREFLEYCEGVGIRARAIPV DYASHTAHV EPV RDELVQALAGITPRRAEVPFSTLTGDFLDGT ELDAGY WYRNL RHPVEFHSAVQALTDQGYATFIEVSPHPV LASSVQETLD DAESDAAVLGTLERDAGDADRFLTALADAHTRGVAVDWEAVLGRAGLVDLPGYFPQGKRFWLLPDRT PRDELDGWFYR VDWTEVPRSEPAALRGRWL VVVPEGHEEDGTV E VRSALAEAGAEPEVTRGVGGLVGDCAGVVSLLAEGDGA VQTLVL VRELDAEGIDAPLWTTFGAVDAGSPVARPDQAKLWGLQV ASLERGPRWTGLV DLPMPDPELRGRL TAVLAGSEDQV AVRADAVRARRLSPAHTATSEYAVPGGTILV TGGTAGL GAEVARWLA GRGA EHLALV SRRGPDT EGVGDLTAELTRLGA RVS VHACDVSSREP VRELVHGLIEQGDVVRGVVHAAGLPQQAINDMDEAAFDEVVAAKAGGA VHL DELCSDAELFLLFS SGAGVWGSARQGAYAAGNAFLDAFARHRRGRGLPATSVAWGLWAAGGMTGDEEAVSFLRERGV RAMPVPRAL AALDR VLA SGETA VVVT DWDWPAFAESYTAARPRPLLDRIVTTAPSERAGEPETESLRDRLAGL PRAERTAELVRL VRTSTATVLGH DDPKAVRATTPFKELGFDSLAVR L RNLLNAATGLRLPSTLVFDHPN ASAVAGFLTSELGS GTPAREASSALRDGYRQAGV SGRVRSYLDLLAGLSDFREHFDGSDGFSLDL VDMADGPGEVTVIC CAGTA A ISGPHEFTRLAGALRGIA PVRAV P QPGYEEG EPLPSSMAA VAAVQADAVIRTQGDKP FV VVAGHSAGALMAYALATELLDRGHPPRGVVLIDVYPPGHQDAMNAWLEELTA TLFDRETVMDDTRLTALGAYDRLTGQWRPRETGLPTLLVSAGEPMGPWPDDSWKPTWPFEHDTVAVPGDHFTMVQEHA DAIARHIDA WLGGGNSKLAALEHHHHH*

MK149 SZ4-L-M3-TE	MQKVAELKNRVAVKLNRNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAEGGSGGGSGKLDPIAIVSMACRLPG GVNTPQRLWELLREGGETLSGFPTDRGWDLARLHHPDPDNPGBTSYVDKGGLDDAAGFDAEFFGVSPREAAAMDPQQRLLETSWELVENAGIDPHSLRGATGVFLGVAKFGYGEDTAAAEDVEGYSVTGVAPAVASGRISYTMLEGPSISVDTACSSLVALHLAVESLRKGESSMAVVGAAVMATPGVFVDFSRQRALAADGRSKAFGAGADGFGFSEGVTLVLLERSEARRNGHEVLAVVRGSAQNQDGASNGLSAPSGPAQRVRIRQALESCGLEPGDVDAVEAHGTGTALGDPIEANALLDTYGRDRDADPRLWLGSVKS NIGHT QAAAGVTGLKVVLA LRNGELPATLHVEEPTPHWDWSSGGVALLAGNQPWRRGERTRRARVSAFGISGTNAHVIVEEAPEREHRETTAHDGRPVLVVSARTTAALRAQAAQIAELLERPDADLAGVGLGLATTRARHEHRAAVVASTREEAVRGLREIAAGAATADAVVEGVTEVDGRNVVFLPGQGSQWAGMGAELLSSSPVFAGKIRACDESMAPMQDWKVSDVLRQAPGAPGLDRV DVVQPVLFAMVSLAEWR SYGVEPAAVVGH SQGEIAAAHVAGALTLEDAAKL VVGRSRLMRSLSGEGGMAAVALGEAA VRERL RPWQDRLSVAAVNGPRSVVSGEP GALR AFSEDCAA EGIRV R D IDV DYASHSPQI ERV REELLET TDIA PRPARVTFHSTVESRSM DGTEL DARYW YRNL RETV RFADAV TRLAESGYD A FIEVSPHPVVQAVEE AVEEADGAEDAVVVGSLHRDGGDLAFLRSMATAHSGVDI RDV ALPG A FPAL PTYF QRKRYWLQPAAPAAAS DELAY RVS WTPIEKPESGNLDGDWL VVTPLISPEWTEMCEAINANGRALRCEVDT SASRTEMAQAVAQAGTGFRGVLSLLSDESACRPGVPAGAVGLLTLVQALGDAGVDAPVWCLTQGAVRTPADDLARPAQT TA HGFAQVAGLELPGRWGGVVDLPESVDDAALRLLVAVL RG GGRAEDH LA VRDGR L HGRV VVRASLPQSGRSWTPHGT VLTGAASPVG DQLVRWLADRG AER LVAGACPGDDLLA AVEEAGASA VVCAQD A A ALREALGDEP VTALV HAGT L TNFG SISEV APEEFAETIAAKT ALLA VLDEV LGDRAVEREVYCSSVAGI WGGAGMA AYAAGSAYLDALAEHHRAR GR SCTSVAWTPWALPGAVDDG YL RER GLRSLSADRAMRTWERVLAAGPVSVAVADVDWPVLSEGFAATRPTA LFAELA GRGGQAEAE PD SGPTGEPAQRLAGLSPD EQQENLLEVANAVAEVLGHESA EINVRRAFSELGLDSLNAMLRKRLSASTGLRPASL VFDHPTV TALA QH L TS QL DSGT PAREASSALRDGYRQAGVSGRVRSYLDLLAGLSDFREHFDGSDGFSDLVDMADGPGEVTVICCA GTAAISGPHEFTRLAGALRGIAPVRAVPQPGYEEGEPLPSSMAAVAAVQADAVIRTQGDKPFFVAGHSAGALMAYALATELLDRGHPPRGVVLIDVYPPGHQDAMNAWLEELTATLFDRET VRMDDTRLTALGAYDRLTGQWRPRETGLPTLLVSAGEPMGPWPDDSWKPTWPFEHDTVAVPGDHFTMVQEHA DAIRHIDAWLGGGNSKLA AALEHHHHH*
MK150 (5)M1-L-SZ3	MSGDNGMTEEKLRRYLKRTVTELDVTARLREVEHRA GEPVAVVAMCRLPGGVSTPEEFWELLSEGRDAVAGLPTDRGWLDLSLFHPDPTRSGTAHQRGGGFLTEATAFDPAFFGMSPREALAVDPQQRLMLELSWEVLERAGIPPTSLQASPTGVFVGLIPQEYCPRLAEGGEGVEGYLMTGTTSVASGRIATLGLEGPAISVDTACSSLVAVHLACQSLRRGESSLAMAGGGVTVMPTPGMLVDFSRMNSLAPDGRCKAFSAGANGFGMAEGAGMLLERLSDARRNGHPVLA VLRGTAVNSDGASNGLSAPNGRAQVRVIQQLAESGLGPADIDA VEA HGTGTRLGDPIEARALFEAYGRDREQPLHLSVKSNLGHTQAAAGVAGVIKMVLAMRAGTLPTLHASERSKEIDWSSGAISLLDEPEPWPGARPRRAGVSSFGISGTNAHAIIEAPQVVEGERVEAGDVVAPWVLSASSAEGLRAQAARLA AHLREHPGQDPDIAYSLATGRAALPHRAAFAPVDESAALRVL DGLATGNADGA AVGTSRAQQRADVFPQGWQWAGMAV DLLTSPVFAA ALRECADA LEPLDFEVIPFLRAEAARREQDAALSTERDV VVQPMFAVMVSLASMWRAHGVEPAAVIGHSQGEIAAACVAGALS LDDAARVVALRSVIATMPGNKGMASIAAPAGEVRARIGDRVEIAAVNGPRSVVAGDSDELDRLVASCTTECIRAKRLAVDYASHSSHVETIRDALHAELGEDFHPLPGFVPFSTVGRWTQPD ELDAGYWYRNLRRTVRFADAVRALAEQGYRTFLEVAHPILTA AIEEIGDGS GADLSAIHSLR RGDGSLADFGEALSRAFAAGVAVDWESVHLGTGARRVPLPTYFQ RERVWLEPKVARRSTEVDEVSALRYRIEWRPTGAGEPARLDGTWL VAKYAGTADETSTAAREALESAGARVRELVVDARCGRDELAERLRSVGEVAGVLSLLAVDEAEPEEAPLALASLADTLSVQAMVSAELGCP LWTVTESAVATGPFERVRNAAHGALWG VGRVIALENPAVWGGLV DVPAGSVAELRH LA AVVSGGAGEDQLALRADGVYGR RWVRAA APATDDEWKPTGTVLVTGGTGGVGGQIARWLARRGAPHLLL VSRSGPDADGAGELVAELEALGARTVACDVTDR E S VRELLGGIGDDVPLSAVFHAAATLDDGTVD TLTGERIERASRAKVLGARNLHELTRELDLTAFVLFSSFASAFGAPGLGGYAPGNAYLDGLAQQR RSDGLPATAV AWTWAGSGMAEGPVADRFR RHGVIEMPPETACRALQNALDRAEV

	PIVIDVRWDRFLAYTAQRPTRLFDEIDDARRAAPQAAAEPRVGALASLPAPEREKALFELVRSHAAVLGHASAERVPAD QAFAELGVDSLSALELRNRLGAATGVRLPTTVFDHPVRTLAAHAAELGGSGGGSG NEVTLENDAAFIENENAYLEK EIARLRKEKAALRNRLAHHK LEHHHHHH*
MK152 SZ4-L-KR1-ACP1-L₁₂-M3-TE	MQKVAELKNRAVAKLNRNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAEGGSGGGSGDEVSALRYRIEWRPTG AGEPARLDGTWLVAKYAGTADETSTAAREALESAGARVRELVDARCGRDELAERLRSVGEAVGVLSSLAVDEAEPEEAP LALASLADTLSLVQAMVSAELGCPLWTVTESAVATGPFERVRNAAHGALWGVGRVIALENPAWGGLVDVPAGSVAELA RHLAAVVSGGAGEDQLALRADGVYGRWRVRAAAPATDDEWKPTGTVLVTGGTGGVGGQIARWLARRGAPHLLVSRSG PDADGAGELVAELEALGARTTVACDVTDRESVRELLGGIGDDVPLSAVFHAAATLDDGTVDTLTGERIERASRAKVLGA RNLHELTRELDLTAFVLFSSFASAFGAPGLGGYAPGNAYLDGLAQQRSDGLPATAVAWTWAGSGMAEGPVADRFRRH GVIEMPPETACRALQNADRAEVCPVIDVRWDRFLAYTAQRPTRLFDEIDDARRAAPQAAAEPVGAHMLASLPAPERE KALFELVRSHAAVLGHASAERVPADQAFELGVDSLSELRNRLGAATGVRLPTTVFDHPVRTLAAHAAELGGAT GAEQAAPATTAPVDPIAIVSMACRLPGGVNTPQRLWELLREGGETLSGFPTDRGWDLARLHHPDPDNPGBTSYVDKGFLD DAAGFDAEFFGVSPREAAAMDQPQRLLLETSWELVENAGIDPHSLRGATGVFLGVAKFGYGEDTAAEDVEGYSVTGVA PAVASGRISYTMLEGPSISVDTACSSLVALHLAVESLRKGESSMAVVGGAAMATPGVFVDFSRQRALAADGRSKAFGA GADGFFGSEGVTLVLLERLSEARRNGHEVLAVERGSALNQDGASNGLSAPSGPAQRVRVIRQALESCGLEPGDVDAVEAHGT GTALGDPIEANALLDTYGRDRDADRPWLGSVKSIGHTQAAAGVTGLLKVVLAIRNGELPATLHVEEPTPHVDWSSGGV ALLAGNQPWRGERTRRARVSAFGISGTNAHVIVEEAPEREHRETTAHDGRPVLVVSARTTAALRAQAAQIAELLERPDA DLAGVGLGLATTRARHEHRAAVVASTREEAVRGLREIAAGAATADAVVEGVTEDGRNVVFLPGQGSQWAGMGAELLS SSPVFAGKIRACDESMAPMQDWKVSDVLRQAPGAPGLRVDVVPVLFAVMVSLAEWRSYGVPAVVGHSQGEIAAA HVAGALTLEDAKLVVGRSRLMRSLSGEGGMAAVALGEAAVERLRPWQDRLSVAAVNGPRSVVVSGEPGALRAFSEDC AAEGIRVRDIDVDYASHSPQIERVREELLETTGDIAPRPARVTFHSTVESRSMDGTELARYWYRNLRETVRFADAVTRLAE SGYDAFIEVSPHPVVQAVEEAVEADGAEDADEVVVGSLHRDGGDLSAFLRSMATAHSGVDIRWDVALPGAAPFALPTYP FQRKRYWLQAPAAPAAASDELAYRVSWTPIEKPESGNLDGDWLVVTPLISPEWTEMCEAINANGRALCEVDTASRTE MAQAVAQAGTGRGVLSLLSSDESACRPGVPAGAVGLLTVQALGDAGVDAPVWCLTQGAVRTPADDDLARPAQTTAHC FAQVAGLELPGRWGGVVLDLPESVDDAALRLLVAVLRGGRGAEDHLAVRDGRHLHGRVVRSALPQSGRSWPHGTVLVT GAASPVGDQLVRWLADRGAEVLVLAGACPGDDLLAAVEEAGASAVVCAQDAALREALGDEPVTAHVAGTLTNFGSIS EVAPEEFAETIAAKTALLAVLDEVLGDRAREVENEVYCSSVAGIWGGAGMAAYAAGSAYLDALAEHHRARGRSCTSVAWTP WALPGGAVDDGYLRERGLRSLSADRAMRTWERVLAAGPVSVAVADVDWPVLSEGFAATRPTALFAELAGRGGQAEAEPM DSGPTGEPAQRLAGLSPDEQQENLLEVANAVERLGHESAAEINVRRAFSELGLDSLNAMALRKRLSASTGLRLPASLVFD HPTVTALAQHLSQLDSGTPAREASSALRDGYRQAGVSGRVRSYLDLLAGLSDFREHFDGSDGFSLDLVMDADPGEVTVI CCAGTAAISGPHEFTRLAGALRGIAVPRAVPQPGYEEGEPLPSSMAAAVAVQADAVIRTQGDKPFVVAAGHSAGALMAYAL ATELLDRGHPPRGVVLIDVYPPGHQDAMNAWLEELTATLFDRETVRMDDTRLTALGAYDRLTGQWRPRETGLPTLLVSAG EPMGPWPDDSWKPTWPFEHDTVAVPGDHFTMVQEHAADAIHIDAWLGGGNSSSVDKLAAALEHHHHHH*
MK168 SZ4-L-KR5-ACP5(2)	MQKVAELKNRAVAKLNRNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAEGGSGGGSGPIPTGGRARDEDWW RYQVVWREAWESESAGRVLLTGPVGVPSELSDAIRSGLEQSGATVLTCDVESRSTIGTATEAADTDALSTVVSLLSRDGE AVDPSLDALALVQALGAAGVEAPLWVLTNAVQVADGELVDPAQAMVGGLGRVVGIEQPGRWGGLVLDVADAASIRS LAAVLADPRGEEQVAIRADGIKVARLVPAPARAARTRWSPRGTVLVTGGTGGIGAHVARWLARSGAEHLVLLGRRGADAP GASELREELTALGTGVTIAACDVADRARLEAVLAERAEGRTVSAMHAAVGSTTPLDDLTEAEFTIADVKVRGTVNLD ELCPDLDASFVLFSSNAGVWGSPGLASYAAANAFLDGFARRRSEGAUTSIAWGLWAGQNMAGDEGGEYLRSQGLRAMD PDRAVEELHITLDHGQTSVSVVDMDRRFVELFTAARHRPLFDEIAGARAEARQSEEGPALAQRALSTAERREHLAHLIR

	AEVAAVLGHGDDAIDRDRFLGFDSMTAVDLRNRLAAVTGVREAATVVFDHPTITRLADHYLERLV GTEVRGEAPSA LAGLDALEALPEVPATEREELVQRLERMLAALRPVAQAADASGTGANPSGDDLGEAGVDELLEALGRELDGDPNSSVD KLAAALEHHHHHH*
MK169 SZ4-L-RIFS(KR1-ACP)(2)	MQKVAELKNRVAVKLNNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAEGSGGGSGEPAEPASAGDPLLGTV VSTPGSDRLTAVAQWSRRAQPWAVDGLVPNAALVEAIRLGDLAGTPVVGELVVDAPVVLPRRGSRREVQLIVGEPGEQRR RPIEVFSREADEPWTRAHGTLAPAAAAPPEAAAGDATDVTAGLRDADRYGIHPALLDAAVRTVVGDDLLPSVWTGVS LLASGATAVTVTPTATGLRLTDPAQPVLTVESVRGTPFVAEQGTTDALFRWDWPEIPLPTAETADFLPYEATSAEATLSAL QAWLADPAETRЛАVVTGDCTEPGAAAIWGLVRSAQSEHPGRIVLAADDDPAVLPAVVASGEPVQRVRNGVASVPLTRVT PRQDARPLDPEGTVLITGGTGLALTARHLVTAGVRHLVLVSRGEAPELQEELTALGASVIAACDVADRAQLEAVL AIPAEHPLTAVIHTAGVLDGVVTELTPDRLATVRRPKVDAARLLDELTREADLAAFVLFSSAAGVLGNPGQAGYAAANAE LDALARQRNSLDPAVSIAWGYWATVSGMTEHLDGADLRRNQRIGMSGLPADEGMALLDAAIATGGTLVAAKFDVAALR ATAKAGGPVPLRGLAPLPRRAAKTASLTERLAGLAETEQAAALLLVRRHAAEVLGHSGAESVHSRFTKDAGFDSL AVELRNRLAAATGTLSPAMIFDYPKPPALADHLRAKL GTEVRGEAPSALAGLDALEALPEVPATEREELVQRLERMLA ALRPVAQAADASGTGANPSGDDLGEAGVDELLEALGRELDGDPN SSSVDKLAALEHHHHHH*
MK170 SZ4-L-M5-TE	MQKVAELKNRVAVKLNNEQLKNKVEELKNRNAYLKNELATLENEVARLENDVAEGSGGGSGKLEPIAIVGMACRFPG DVDSPEFWEFVSGGGDAIAEAPADRGWEPDPDARLGGMLAAAGDFDAGFFGISPREALAMDQQRIMLEISWEALERAG HDPVSLRGSATGVFTGVGTVDYGPRPDEAPDEVLYGVTGTAASSVASGRVAYCLGLEGPAMTVDTACSSGLTALHAMES LRRDECGLALAGGVTVMSSPGAFTEFRSQGGLAADGRCKPFSKAADGFGLAEGAGVVLQRLSAARREGRPVLAVRGSA VNQDGASNGLTAPSGPAQQRVIRRALENAGVRAGDVYVEAHGTGTRLDPIEVHALLSTYGAERDPDDPLWIGSVKSNI HTQAAAGVAGVMKAVLARHGEMPRTLHFDEPSPQIEWDLGAVSVSQARSWPAGERPRRAVGSSFGISGTNAHVIVEEA PEADEPEPAPDGPVPLVLSRDEQAMRAQAGRLADHLLAREPRNSLRDTGFTLATRrsaWEHRAVVVGDRDDALAGLRAV ADGRIADRTATGQARTRRGVAMVPGQGAQWQGMARDLLRESQVFADSIRDCCRALAPHWDWSLTLSGARPLDRV VQPALFAVMVSLAALWRSHGVEPAAVVGHSQGEIAAAHVAGALTLEDAAKLVAVRSRVLRLGGQGMASFGLGTEQA AERIGRFAGALSIASVNGPRSVVVAGESGPLDELIAECEAEGITARRIPDVYASHSPQVESLREELLTELAGISPVSADVALYST TTGQPIDTATMDTAYWYANLREQVRFQDATRQLAEAGFDAFVEVSPHPVLTGIEATLDSALPADAGACVVGTLRRDRGG LADFHTALGEAYAQGVEVDWSPAFAADARPVELPVYPFQRQRYWLPIPTGGRARDEDWWRYQVVWREAEWESASLAGR VLLVTGPGVPSELSDAIRSGLEQSGATVLTCDVSRSTIGTALEAADTDALSTVVSLLSRDGEAVDPSLDALALVQALGAAG VEAPLWVLTRNAVQVADGELVDPAQAMVGLGRVVGIEQPGRWGLVLDVADAASIRSLAAVLADPRGEEQVAIRADG IKVARLVPAPARAARTWSPRGTVLVTGGTGGIGAHVARWLARSGAEHLVLLGRRGADAPGASELREELTALGTGTVIAA CDVADRARLEAVLAAERAEGRTVSAMHAAGVSTPLDDLTEAEFTIADVKGRTVNLDELCPDLDASFVLSNAGVW GSPGLASYAAANAFLDGFARRRSEGAPVTSIAWGLWAGQNMAGDEGGYLRSQGLRAMDPDRAVEELHITLDHGQT SVVDMDRRFVELFTAARHRPLFDEIAGARAEARQSEEGPALAQRALSTAERREHLAHLIRAEVAAVLGHGDDAIDRD RAFRDLGFDSMTAVDLRNRLAAVTGVREAATVVFDHPTITRLADHYLERLVSGBTAREASSALRDGYRQAGVSGRVR SYLDLLAGLSDFREHFDGSDGFSLDLVDMADGPGEVTVICCACTAAISGPHEFTRLAGALRGIAPVRAVPQPGYEEGEPLSSMA AVAAVQADAVIRTQGDKPFVVAGHSAGALMAYALATELLRGHPPRGVVLDVYPPGHQDAMNAWLEELTATLFDRET RMDDTRLTALGAYDRLTGQWRPRETGLPTLLVSAGEPMGPWPDDSWKPTWPFEHDTVAVPGDHFTMVQEHA DAIARHIDAWLGGGNSKLA ALEHHHHHH*

Table S3: Yields of proteins used in this study. If no indication is given the PKS domain/modules are derived from DEBS. Typical yields are presented.

Construct	Protein	Yield /mg/L of culture
MK96	KR1-ACP1(2)	8.6
MK111	(5)KS1-AT1-SZ3	1.3
MK112	SZ4-KR1-ACP1(2)	7.9
MK113	(5)KS1-AT1	1.1
MK136	SZ4-KR1-ACP1-M2-TE	0.9
MK137	SZ4-KR5-ACP5-M6-TE	2.1
MK138	SZ4-KR1-ACP1-M2-TE	3.4
MK141	SZ4-KR1-ACP1-M5-TE	1.6
MK142	SZ4-KR1-ACP1-M6-TE	1.4
MK147	SZ4-M6-TE	7.9
MK148	SZ4-M2-TE	1.5
MK149	SZ4-M3-TE	3.8
MK150	(5)M1-SZ3	15.3
MK152	SZ4-KR1-ACP1-M3-TE	3.6
MK168	SZ4-KR5-ACP5(2)	2.3
MK169	SZ4-RIFS(KR1-ACP1)(2)	9.8
MK170	SZ4-M5-TE	3.3
BL12	LDD(4)	6.7
BL13	(5)M1(2)	2.4
BL16	(3)M2-TE	1.6
BL17	(3)M5-TE	0.8
BL18	(3)M6-TE	4.9
RSG34	(3)M3-TE	9.4

References:

- (1) Tang, Y., Kim, C., Mathews, I. I., Cane, D. E., and Khosla, C. (2006) The 2.7-Å crystal structure of a 194-kDa homodimeric fragment of the 6-deoxyerythronolide B synthase. *Proc. Natl. Acad. Sci.* **103**, 11124–11129.
- (2) Keatinge-Clay, A. T., and Stroud, R. M. (2006) The Structure of a Ketoreductase Determines the Organization of the β-Carbon Processing Enzymes of Modular Polyketide Synthases. *Structure* **14**, 737–748.
- (3) Reinke, A. W., Grant, R. A., and Keating, A. E. (2010) A synthetic coiled-coil interactome provides heterospecific modules for molecular engineering. *J. Am. Chem. Soc.* **132**, 6025–6031.
- (4) Thompson, K. E., Bashor, C. J., Lim, W. A., and Keating, A. E. (2012) SYNZIP Protein Interaction Toolbox: *in Vitro* and *in Vivo* Specifications of Heterospecific Coiled-Coil Interaction Domains. *ACS Synth. Biol.* **1**, 118–129.
- (5) Viegas, M. F., Neves, R. P. P., Ramos, M. J., and Fernandes, P. A. (2018) Modeling of Human Fatty Acid Synthase and in Silico Docking of Acyl Carrier Protein Domain and Its Partner Catalytic Domains. *J. Phys. Chem. B* **122**, 77–85.
- (6) Kapur, S., Lowry, B., Yuzawa, S., Kenthirapalan, S., Chen, A. Y., Cane, D. E., and Khosla, C. (2012)

- Reprogramming a module of the 6-deoxyerythronolide B synthase for iterative chain elongation. *Proc. Natl. Acad. Sci.* **109**, 4110–4115.
- (7) Kao, C. M., Katz, L., and Khosla, C. (1994) Engineered biosynthesis of a complete macrolactone in a heterologous host. *Science* **265**, 509–512.
- (8) Pfeifer, B. A., Admiraal, S. J., Gramajo, H., Cane, D. E., and Khosla, C. (2001) Biosynthesis of complex polyketides in a metabolically engineered strain of *E. coli*. *Science (80-.)* **291**, 1790–1792.
- (9) Chen, A. Y., Cane, D. E., and Khosla, C. (2007) Structure-Based Dissociation of a Type I Polyketide Synthase Module. *Chem. Biol.* **14**, 784–792.
- (10) Klaus, M., Ostrowski, M. P., Austerjost, J., Robbins, T., Lowry, B., Cane, D. E., and Khosla, C. (2016) Protein Protein Interactions, not Substrate Recognition, Dominates the Turnover of Chimeric Assembly Line Polyketide Synthases. *J. Biol. Chem.* **291**, 16404–16415.
- (11) Lowry, B., Robbins, T., Weng, C., Brien, R. V. O., Cane, D. E., and Khosla, C. (2013) In Vitro Reconstitution and Analysis of the 6-Deoxyerythronolide B Synthase. *J. Am. Chem. Soc.* **135**, 16809–16812.
- (12) Gokhale, R. S., Tsuji, S. Y., Cane, D. E., and Khosla, C. (1999) Dissecting and exploiting intermodular communication in polyketide synthases. *Science* **284**, 482–485.