

Phylogenomic analysis of type I polyketide synthase genes in pathogenic and saprobic ascomycetes

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Fungal type I polyketides (PKs) are synthesized by PK synthases (PKSs) and include well known secondary metabolites such as the anticholesterol drug lovastatin and the potent natural carcinogen aflatoxin. Other type I PKs are known to be virulence factors for some plant pathogens and pigments such as melanin. In this study, a phylogenomic approach was used to investigate the origin and diversity of fungal genes encoding putative PKSs that are predicted to synthesize type I PKs. The resulting genealogy, constructed by using the highly conserved PKS ketosynthase (KS) domain, indicated that: (i) Species within subphylum Pezizomycotina (phylum Ascomycota) but not early diverging ascomycetes, like *Saccharomyces cerevisiae* (Saccharomycotina) or *Schizosaccharomyces pombe* (Taphrinomycotina), had large numbers (7–25) of PKS genes. (ii) Bacteria and fungi had separate groups of PKS genes; the few exceptions are the likely result of horizontal gene transfer from bacteria to various sublineages of fungi. (iii) The bulk of genes encoding fungal PKSs fell into eight groups. Four groups were predicted to synthesize variously reduced PKs, and four groups were predicted to make unreduced PKs. (iv) Species within different classes of Pezizomycotina shared the same groups of PKS genes. (v) Different fungal genomes shared few putative orthologous PKS genes, even between closely related genomes in the same class or genus. (vi) The discontinuous distributions of orthologous PKSs among fungal species can be explained by gene duplication, divergence, and gene loss; horizontal gene transfer among fungi does not need to be invoked.

Type I fungal polyketide (PK) synthases (PKSs) are closely related to fatty acid synthetases (FASs) (1). Both are multifunctional enzymes with the same ancestral enzymatic domain structure, namely ketoacyl synthase (KS), acyl transferase (AT), ketoreductase (KR), dehydratase (DH), enoyl reductase (ER), and acyl carrier protein [also known as a phosphopantetheine attachment site (PP) domains]. The KS, AT, and PP domains are essential for both FASs and PKSs, whereas the KR, DH, and ER domains are present in all FASs, but some or all are absent in various PKSs. KR, DH, and ER domains catalyze, in a stepwise fashion, reduction of a keto to a hydroxyl group, dehydration of the hydroxyl to an enoyl group, and reduction of the enoyl to an alkyl group, respectively. PKSs that lack some or all of these domains produce partially reduced or fully oxidized PKs.

Fungal type I PKSs, like FASs, are monomolecular enzymes; most are iterative and use their active sites repeatedly to synthesize a PK, adding a two-carbon molecule (i.e., a CoA ester) to the growing chain with each condensation. The noniterative fungal PKSs perform only one condensation cycle and make a diketide. Each one that has been characterized so far is encoded by a gene that resides in a gene cluster (genes adjacent along one stretch of a chromosome), along with a PKS gene encoding an iterative PKS (2, 3). The products of an iterative and a noniterative PKS are joined to make a branched PK. The diversity of PKs is generated, in part, through the use of the three optional PKS reducing domains.

Access to genomes of saprobic and pathogenic members of the Ascomycota allowed us to address the following questions regarding distribution and evolution of Type I PKS genes: (i)

How many PKS genes are there in these genomes? (ii) Do type I PKS genes from fungi and bacteria fall into the same or different groups? (iii) How many subgroups of type I PKS genes are found in the Ascomycota? (iv) Do species in the different classes of Ascomycota share the same or have different subgroups of type I PKS genes? (v) What is the frequency of putative orthologs among distantly related fungi, including those that evolved independently as plant pathogens and among closely related species? (vi) Which evolutionary process can account for the phylogenetic distribution of PKS genes: gene duplication, divergence, gene loss, horizontal gene transfer (HGT), or all of these?

To address these questions, we performed phylogenetic analyses on the amino acid sequences of KS domains encoded by all previously characterized fungal PKS genes and by all putative PKS genes extracted from genomic sequences of five taxonomically diverse fungal species in the Ascomycota, subphylum Pezizomycotina (4) [the saprobe *Neurospora crassa* (5); the maize pathogens *Cochliobolus heterostrophus* (*Bipolaris maydis*) (6, 7) and *Gibberella moniliformis* (*Fusarium verticillioides*) (8); the cereal pathogen *Gibberella zeae* (*Fusarium graminearum*) (8); and the cosmopolitan dicot pathogen *Botryotinia fuckeliana* (*Botrytis cinerea*) (8)]. We also searched for PKS genes in the genomes of three earlier diverging ascomycetes [*Saccharomyces cerevisiae* (9), the plant pest *Eremothecium* (*Ashbya*) *gossypii* (10), (both in the Saccharomycotina), and the yeast saprobe *Schizosaccharomyces pombe* (11) (Taphrinomycotina)].

Materials and Methods

Detailed treatments of all section below are provided as *Supporting Materials and Methods*, which is published as supporting information on the PNAS web site.

DNA and Protein Genomic Databases. Four genomes were sequenced by Celera Genomics for the Torrey Mesa Research Institute (TMRI)/Syngenta: *B. fuckeliana* strain B05.10 (to $\approx 5\times$ coverage), *C. heterostrophus* strain C4 (ATCC 48331) (to $\approx 5\times$ coverage), *G. moniliformis* strain ATCC 38932 (FGSC 7600) (to $\approx 4\times$ coverage), and *G. zeae* lineage 7 strain GZ3639 (12) (to $\approx 2\times$ coverage). The shotgun sequence assemblies of these genomes were used for these studies, along with the genome

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Abbreviations: PK, polyketide; PKS, PK synthase; KS, ketoacyl synthase; AT, acyl transferase; DH, dehydratase; NPS, nonribosomal peptide synthetase; ME, methyl transferase; ER, enoyl reductase; KR, ketoreductase; PP, phosphopantetheine attachment site (acyl carrier protein of PKSs and thiolation of NPSs); AMP, adenylation; CON, condensation; HGT, horizontal gene transfer; TMRI, Torrey Mesa Research Institute; FAS, fatty acid synthase.

Data deposition: The sequences reported in this paper have been deposited in the GenBank database [accession nos. AY495643–AY495666 (*C. heterostrophus* PKS2–25), AY495606–AY495625 (*B. fuckeliana* PKS1–20), AY495591–AY495605 (*G. moniliformis* PKS1–15), AY495626–AY495641 (*G. zeae* PKS1–16), and AY495642 (PKS1 from *Didymella maydis*)].

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sequences of *N. crassa* (5), *S. cerevisiae* (9), *E. gossypii* (10) (sequenced by Syngenta), and *S. pombe* (11). The protein databases for *C. heterostrophus*, *B. fockeliana*, and *G. moniliformis* were compiled by using four gene models that were found to generate the largest numbers of predicted proteins: GENEMARK.HMM (13) (*Arabidopsis* and *Caenorhabditis* models), FGENESH (14) (dicot model), and GENSCAN (15) (*Arabidopsis* model). *N. crassa*, *S. cerevisiae*, and *S. pombe* protein databases are available at the National Center for Biotechnology Information.

PKS Gene Identification. To retrieve PKS genes or predicted proteins from each genome, we used the consensus sequence of the KS domain, the most highly conserved domain in type I PKSs and FASs (1), as a query in TBLASTN searches of the TMRI genome assemblies; in PSI-BLAST searches of the TMRI protein databases of *C. heterostrophus*, *B. fockeliana*, and *G. moniliformis*; and in PSI-BLAST searches of the eukaryote protein database (www.ncbi.nlm.nih.gov/BLAST) for *N. crassa*, *S. cerevisiae*, and *S. pombe*. The PSI-BLAST searches were also used to retrieve all previously published fungal PKS protein sequences from National Center for Biotechnology Information.

PKS Gene Annotation. The protein sequences of the KS domains obtained from TBLASTN and PSI-BLAST searches were used as queries to retrieve the complete PKS protein sequences from the protein databases. The *C. heterostrophus*, *B. fockeliana*, and *G. moniliformis* PKS-predicted sequences from each of the four protein databases were retrieved and aligned by CLUSTALW by using the amino acid substitution matrix BLOSUM62, implemented in MEGALIGN (DNASTAR, Madison, WI), and the protein predictions and corresponding DNA sequences were evaluated for their predicted introns. The predicted PKS proteins from *N. crassa* were also aligned to verify intron predictions. The predicted proteins were submitted to Hidden Markov Model searches, implemented in TIMELOGIC (TimeLogic, Carlsbad, CA), to determine their multidomain PKS protein structures.

KS-Domain Genealogy Construction and Evaluation. The predicted KS domains of all newly sequenced fungal genomes plus those from all previously published PKS protein sequences from ascomycete fungi, the homologous FASs from animals (1), and representative type I PKSs from bacteria (Table 1, which is published as supporting information on the PNAS web site) were aligned with CLUSTALW (Table 2, which is published as supporting information on the PNAS web site) and phylogenetically analyzed with neighbor joining and maximum parsimony in PAUP4.0b8 (Sinauer, Sunderland, MA), by using the settings given in *Supporting Materials and Methods*. The resulting KS-domain genealogy was evaluated to rank major clades and subclades of PKS enzymes. The criteria used to categorize proteins in major clades and subclades were moderate bootstrap support, similar domain structure, and similar chemical characteristics of the resulting PK products for those PKSs that have been previously characterized. KS-domain genealogy was also used to predict putative phylogenetic orthologs that are expected to encode enzymes of identical or nearly identical biochemical function (16, 17). The criteria used to predict proteins as orthologs were bootstrap values >80%, identical predicted domain structure (or nearly so), and chemical characteristics similar to those of the resulting PK product for those proteins whose product has been characterized previously. Putative orthologs were further supported by a KS-domain genealogy that was consistent with the currently understood organismal phylogeny (4), applicable when three or more orthologs are present. Alternative phylogenetic hypotheses were tested for significance with a Kishino-Hasegawa test (18). The hypothesis underlying the constraint tree is rejected if the difference in logarithm likelihood com-

pared to the unconstrained tree divided by the SD (Ln L/SD) is >1.96 ($P \leq 0.05$) (19).

Results and Discussion

Species Within the Pezizomycotina Have Many PKS Genes. The four newly sequenced genomes have large numbers of putative PKS genes: 15 in *G. moniliformis*, 16 in *G. zaeae*, 20 in *B. fockeliana*, and 25 in *C. heterostrophus* (Table 3, which is published as supporting information on the PNAS web site). The PKS genes in *G. zaeae* were represented as fragments, due to the small contig size and many gaps in the 2× coverage; however, a search of the publicly available *G. zaeae* genome {albeit a different strain [PH-1 (NRRL 31084, but the same lineage, lineage7)]} did not reveal additional PKS genes beyond those we identified in our 2× coverage. These numbers are greater than those of the saprobe *N. crassa*, which is thought to produce only one PK, namely PK-derived melanin, yet contains seven putative PKS genes, only one of which is a melanin-type PKS gene (5). By contrast, the three earlier-diverging ascomycetes (*S. pombe*, *S. cerevisiae*, and *E. gossypii*) lacked identifiable PKS genes, including those involved in pigment formation.

KS-Domain Genealogy and Prediction of PK Structure. KS-domain genealogy (Fig. 1) was used to infer the genealogy of type I PKSs. The predicted fungal and bacterial PKSs grouped in three main clades (20–22). Together, these three major clades grouped into a larger clade of type I PKSs, which was sister to the clade of FASs found in animals. All of the type I PKSs and FASs grouped into a larger clade, which was sister to the type II PKSs found in bacteria and to the bacterial and mitochondrial acyl-ACP synthetases (1). KS-domain genealogy suggested that the ancestral domain structure of type I PKSs was KS-AT-DH-ME-ER-KR-PP, because these domains were present, and in this order, in the PKS protein sequences in members of two of the three main clades of PKSs, one of which (the bacterial PKS clade) was in a basal position.

Two of these three main clades of microbial type I PKSs were identified as exclusively fungal and correlated with the two largest structural classes of fungal PKs: reduced and unreduced (1). Each main fungal PKS clade was further divided into four groups, each with a typical domain structure, depending on the loss of ancestral domains, or the gain of novel domains (mapped onto Fig. 1 and listed in Table 3). The third major clade included all bacterial PKSs, and within this clade were nested two small, additional fungal clades, the first of which comprised the 6-methylsalicylic acid synthetases, and the second of which was a nonribosomal peptide synthetase (NPS)/PKS hybrid gene found in *C. heterostrophus*.

Fungal PKSs Producing Reduced PKs. The first main fungal clade included the PKSs that synthesize variously reduced, and usually linear, PKs (Fig. 1). The characterized PKs serve as precursors to toxins that are active in animals [e.g., lovastatin (2), citrinin (3), and fumonisin (23)] and to toxins that are active in plants [e.g., T-toxin (6), and PM-toxin (24)]. These reduced PKs frequently are synthesized from CoA thioesterified carboxylic acids other than acetyl and malonyl CoA, and the extent of keto-group processing varies from one condensation cycle to the next (1). In addition, one or more sites may be methylated by a ME domain (3, 25). However, many of the predicted PKSs in the PKS clade that produces reduced PKs have highly divergent and presumably nonfunctional ME domains.

The clade of PKSs that synthesize reduced PKs was subdivided into four subclades (I–IV), each of which had a typical domain structure (Fig. 1). Reducing PKS subclade I included *Aspergillus terreus* LovF and its ortholog *Penicillium citrinum* MlcB, which synthesize the diketide portion of lovastatin (2) and citrinin (3), respectively, and *C. heterostrophus* PKS1 and its ortholog *Didy-*

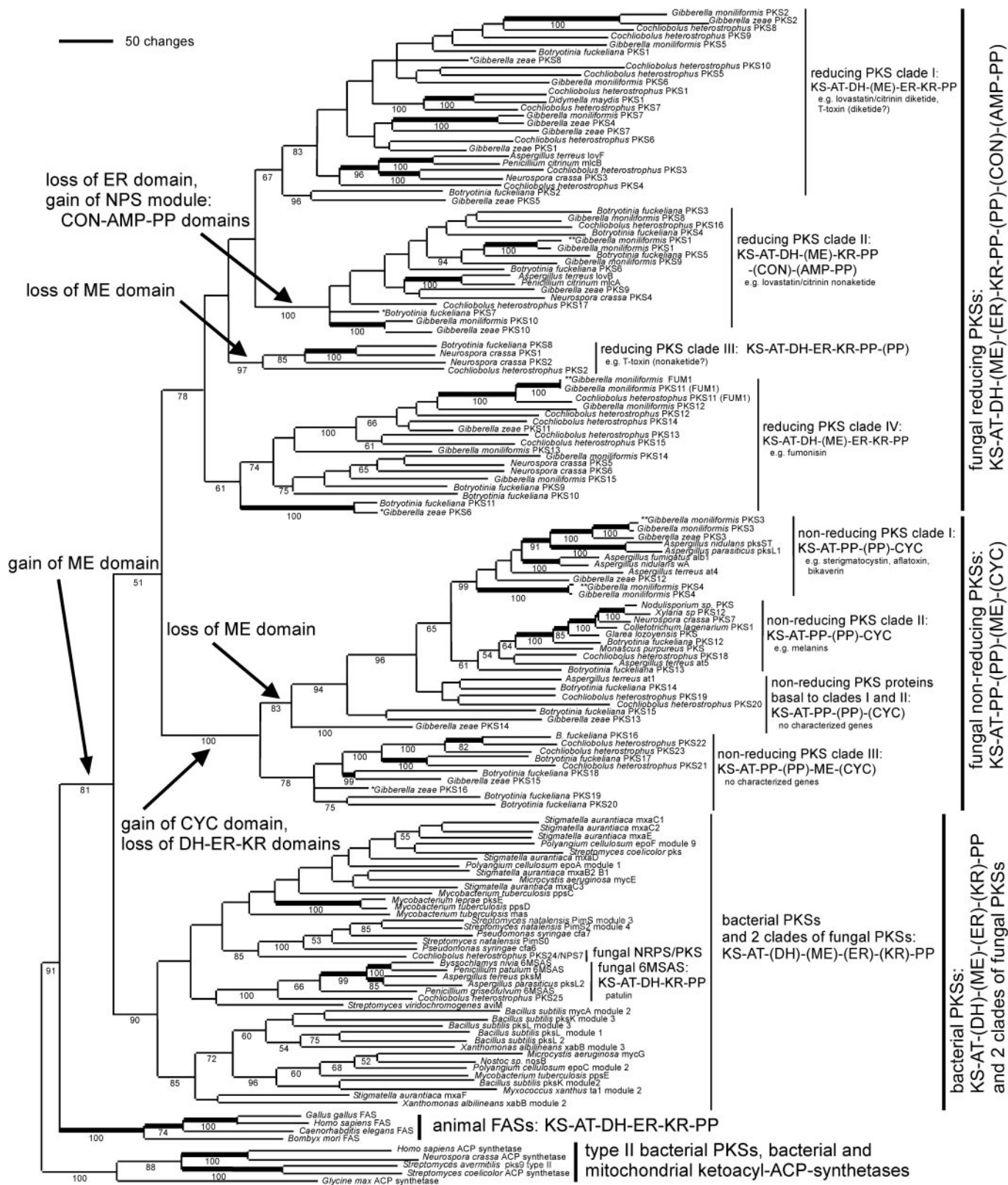


Fig. 1. Genealogy of type I PKSs, inferred by maximum parsimony analysis of the KS domain. Major clades and subclades are indicated by vertical bars, each of which shares a common organization of domains (those in parentheses are variable in their presence/absence within that clade). Branch length indicates number of inferred amino acid changes. Numbers below branches indicate percentage bootstrap support for each clade. All branches present in a strict consensus of the maximum parsimony trees received bootstrap support. Bold branches indicate putative orthologs, which were inferred as described in *Materials and Methods*. Monophyletic gains and losses of domains are noted by arrows. Three *G. moniliformis* PKSs were previously represented by GenBank submissions; these are noted with double asterisks. The accession numbers for all sequences obtained from GenBank are given in Table 1. Three *G. zeae* and one *B. fockeliana* PKSs represented by partial C-terminal fragments of the KS domain were mapped onto the tree that was based on an alignment that included only N-terminal

mella (*Mycosphaerella*) *maydis* PKS1, which are necessary for the synthesis of structurally similar linear PKs, T-toxin (6), and PM-toxin (26), respectively.

Reducing PKS subclade II was characterized by enzymes that have lost the ER domain (Fig. 1); PKs synthesized by PKSs in this clade are predicted to either lack reduced alkyl groups or to contain alkyl groups whose reduction is completed by the product of an external ER domain-containing gene [e.g., *A. terreus* *lovC* (25), *P. citrinum* *mlcG* (3)]. The PKSs of this clade were also found to have either a condensation (CON) domain typical of NPSs (25) or an entire NPS module consisting of a CON domain, an adenylation (AMP) domain, and a PP domain (Table 3 and Fig. 1). Genealogies of the AMP and CON domains, along with those from characterized fungal and bacterial NPS proteins, grouped the AMP domains with those that adenylate amino acids that are subsequently *N*-methylated and grouped the CON domains with those that condense *N*-methylated amino acids (analysis not shown). These data suggest that the PKSs in this subclade gained an *N*-methylated amino acid-type NPS module (described in refs. 27–30). However, none of these PKSs had an *N*-methylation domain that is found between the AMP and CON domains of *N*-methylated amino acid type NPS modules. Many of these PKSs also lacked AMP and PP domains; these PKSs were scattered throughout this subclade, suggesting that the AMP and PP domains have been lost repeatedly, including those from the only two characterized proteins, *A. terreus* *LovB* and its ortholog *P. citrinum* *MlcA*, which synthesize the cyclic nonaketide portion of lovastatin and citrinin, respectively (3, 25). Each cyclic PK is derived from the action of the CON domain on a methylated keto group. This mode of action suggests that the loss of the AMP and PP domains was involved in a change of function of the CON domain from acting on methylated amino acids to acting on methylated keto groups. Uncharacterized PKSs with the same domain structure are predicted to make cyclic PKs. PKSs that have lost the entire C-terminal NPS module, consisting of the CON, AMP, and PP domains, are also scattered throughout this clade. The inferred polyphyletic losses of the CON, AMP, and PP domains were tested by constraint topologies forcing subsets of sequences into clades, which were subjected to parsimony analyses, and the likelihoods of the resulting genealogies were compared to the likelihood of the maximum parsimony tree found without these topological constraints. The Kishino–Hasegawa tests (18) rejected the alternative hypothesis of a monophyletic origin of the four PKSs that lack the entire NPS-like module via a sequential loss of first the AMP-PP domains and then the CON domain [$P < 0.0001$] (19) or single loss of all three domains ($P = 0.0302$).

PKSs in reducing subclade III all lacked a ME domain, and two of the four PKSs had an additional PP domain. The only characterized protein is *C. heterostrophus* PKS2, which, along with PKS1, is required for synthesis of T-toxin (S. E. Baker, personal communication).

Reducing PKS subclade IV, like subclades I and II, included PKS enzymes that may or may not have a conserved ME domain. The only characterized PKS in this subclade is *G. moniliformis* FUM1, which makes the linear PK precursor of the toxin fumonisin (31, 32).

The PKSs that lack conserved and putatively functional ME domains are scattered throughout reducing PKS clades I, III, and IV and comprise reducing PKS clade II. The alternative hypothesis of a monophyletic loss of ME domains in the reducing PKS clade was rejected by the Kishino–Hasegawa test ($P < 0.0001$).

Fungal PKSs Producing Unreduced PKs. The second main fungal clade included fungal PKSs that synthesize unreduced, and usually cyclic, i.e., aromatic, PKs that are precursors to toxins [e.g., sterigmatocystin (33) and aflatoxin (34)] and pigments [e.g., melanin (35), bikaverin (36), and green spore pigment (37)]. All PKSs within this clade lacked ER, DH, and KR domains, which is interpreted as a single loss of these reducing domains, compared to the inferred ancestral domain structure of type I PKSs (Fig. 1, arrows). The nonreducing fungal PKSs are predicted to synthesize PKs in which the keto groups are either not reduced or may be reduced by enzymes other than a PKS. Unreduced PKs are typically synthesized from acetyl- and malonyl-CoA. PKS proteins with an additional PP domain were scattered throughout the nonreducing PKS clade, as was the case for reducing PKS subclade III. The functional significance of these duplicated PP domains is not known.

A Claisen-type cyclase (38) domain was present at the C terminus of all PKS proteins in sister subclades I (toxins and nonmelanin pigments) and II (melanin). The PKSs that make melanin precursors are encoded by the most widely distributed fungal PKS genes and have been characterized in *Colletotrichum lagenarium* (PKS1) (35), *Glarea* sp., *Nodulisporium* sp. (39), and *C. heterostrophus* (PKS18) (B. Robbertse and S. Baker, personal communication). *N. crassa* PKS7 maps to the same region as the *per* locus; *per* mutants have nonmelanized perithecia and ascospores (40). The melanin-type PKS gene is not ubiquitous, because the two *Gibberella* genomes lacked a gene within unreduced PKS clade I, which correlates with a lack of melanized structures in their life cycles.

All PKSs basal to nonreducing PKS subclades I and II are uncharacterized. These basal PKSs occur both with and without the Claisen-type cyclase (CYC domain); the PKSs without CYC are predicted to make unreduced noncyclic PKs. We propose calling this clade the nonreducing PKs, because this character is invariable, whereas not all PKSs in this clade are likely to make aromatic (cyclic) PKs. Several of these basal PKSs formed a subclade (nonreducing PKS subclade III) characterized by a ME domain located after the PP domain [KS-AT-PP-(PP)-ME-(CYC)], the apparent result of a domain rearrangement. The other PKSs basal to nonreducing subclades I and II do not have ME domains; we hypothesize that the genes encoding these PKSs diverged after the loss of the ME domain that was retained in nonreducing PKS subclade III (Fig. 1, arrow).

Other Fungal and Bacterial PKSs. The third main fungal PKS clade comprised the 6-methylsalicylic acid-type (i.e., simple aromatic) of PKSs that make 6-methylsalicylic acid, a precursor to toxins [e.g., patulin (41)]. These toxins have been characterized from *Aspergillus* spp. (27, 42), *Byssosclamyces nivea*, and *Penicillium* spp. (27). The *C. heterostrophus* genome had a PKS in this clade, but the other Pezizomycotina genomes did not. This fungal clade was nested within the large clade that comprised all bacterial type I PKSs. A second fungal clade was also nested within this bacterial clade and was represented by a single member, *C. heterostrophus* PKS24. PKS24 is a hybrid NPS/PKS gene, predicted to encode an enzyme that synthesizes a partly reduced PK decorated with a single amino acid.

Origin of PKS Genes in the Ascomycota. Fungal genome sequences available were limited to the three subphyla of Ascomycota, i.e., Taphrinomycotina, Saccharomycotina, and Pezizomycotina, as

fragments; these are marked with a single asterisk. Alignment was based on 4,862 amino acids from the KS domain; 462 characters were informative, 20 were uninformative, and 4 were constant. Parsimony was performed with 100 random additions. Shown is one of 18 most-parsimonious trees of 17,859 steps, consistency index (CI) = 0.2763, rescaled CI = 0.1491. The maximum parsimony trees generated by coding gaps as either 21st amino acid or as missing were not significantly different from each other or from the tree generated by neighbor joining.

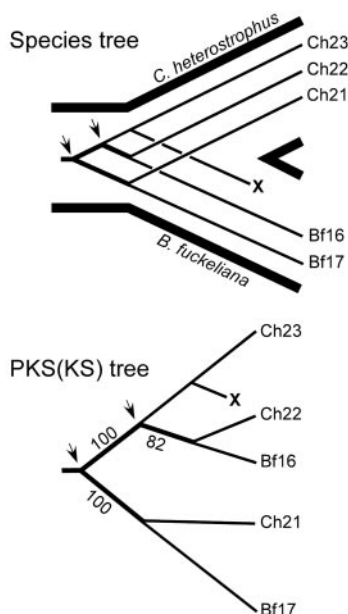


Fig. 2. Species tree (*Upper*) depicts the inferred gene duplication and gene losses necessary to account for the distribution of these PKS proteins present in fungal nonreducing PKS subclade III (*Lower*). Arrows indicate the two gene duplication events, and X indicates the extinction of the ortholog of *C. heterostrophus* PKS23 in the lineage leading to *B. fuckeliana*.

well as to one of the three subphyla of Basidiomycota, Hymenomycetes (*Phanerochaete chrysosporium*, www.jgi.doe.gov/programs/whiterot.htm). PKS genes were found only in the genomes of Pezizomycotina. It is possible that these genes have been lost in the non-Pezizomycotina taxa or that PKS genes came to the Pezizomycotina via HGT, perhaps from the bacterial PKS genes that form the sister clade to the main fungal clade. However, until fungal genomes from other Basidiomycota and from Zygomycota and Chytridiomycota are sequenced, this question will remain unresolved.

Diversification of PKS Genes Among Species in the Pezizomycotina.

The diversity of PKS genes and PKs found in Pezizomycotina led to speculation, including our own (6, 43), that HGT is involved in generating and maintaining this diversity (ref. 44 and references therein). However, other discontinuous genome events such as gene duplication and differential gene loss are alternative explanations to HGT for the distributions of PKS genes (45).

We have data for genomes of species within three classes (Sordariomycetes, Dothideomycetes, and Leotiomycetes) of the Pezizomycotina (Table 3). For the fungal-reducing PKSs, all three classes are represented in each of the four subclades. For the fungal-nonreducing PKSs, all three classes are represented, but not in each of the four subclades. We also have data from a fourth class, Eurotiomycetes, in the form of many PKS proteins in GenBank from *Penicillium* and *Aspergillus* species (Table 1); these PKSs were found in two of the four fungal-reducing PKS subclades and three of the four fungal-nonreducing PKS subclades. This phylogenomic distribution (Fig. 1) is consistent with the hypothesis that all eight types of PKS gene were present in the common ancestor of the four classes of the Pezizomycotina, before the radiation of Pezizomycotina 300–700 million years ago (46).

Evidence for gene duplication and gene loss was observed; for example, in nonreducing PKS subclade III, a clade with many strongly supported internal branches (Fig. 2). Using only the genomes of *C. heterostrophus* and *B. fuckeliana*, two gene dupli-

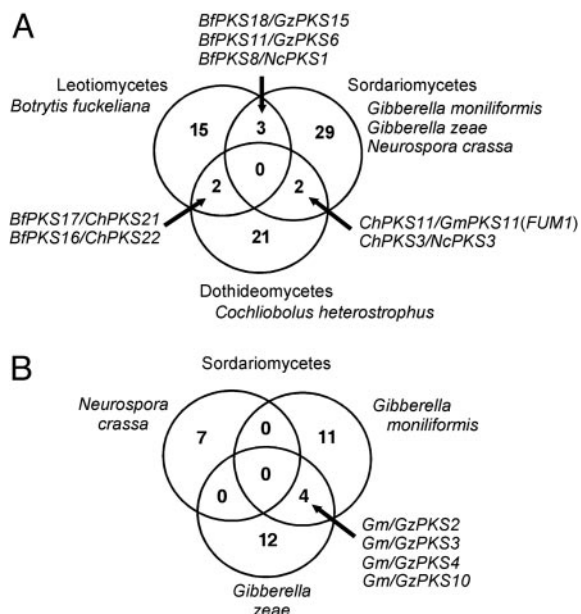


Fig. 3. Venn diagrams showing predicted PKS gene orthologs shared between and among taxa, inferred as described in *Materials and Methods*. (A) Among the classes Leotiomycetes (represented by *B. fuckeliana*), Dothideomycetes (*C. heterostrophus*), and Sordariomycetes (*N. crassa*, *G. moniliformis*, and *G. zeae*). (B) Among the Sordariomycetes: *N. crassa*, *G. moniliformis*, and *G. zeae*.

cations in an ancestor of these two fungi would account for the present distribution of three pairs of paralogs, *ChPKS21/BfPKS17*, *ChPKS22/BfPKS16*, and *ChPKS23*, provided that the counterpart of *C. heterostrophus* PKS23 was lost in the lineage leading to *B. fuckeliana*. HGT does not need to be invoked.

It would seem possible to distinguish between gene duplication and loss and HGT by comparing the time of divergence of PKS genes to that of genes taken to represent the organismal divergence. Similar divergence times for the two types of gene would favor gene duplication. More recent divergence for the PKS genes would favor HGT. Although rates of nucleotide substitution have been estimated for fungal tubulin genes (47), substitution rates for PKS genes are not known and will require data sets with large numbers of orthologous PKS genes.

The *N. crassa* genome is remarkable for its lack of duplicated genes (5), due presumably to repeat-induced point mutation that alters nucleotides in both copies of duplicated genes and renders them nonfunctional. If HGT were the main method of generating PKS gene diversity, *N. crassa* might be expected to have as many PKS genes as the other fungi. However, *N. crassa* has by far the fewest PKS genes, suggesting that gene duplication may be more important than HGT for generating PKS gene diversity.

Origins of Fungal PKS Genes in the Bacterial PKS Gene Clade.

We found two PKS genes in *C. heterostrophus*, each sister to different PKS genes from bacteria. Several branches uniting the fungal and bacterial PKS genes in the bacterial PKS clade were so well supported that it seems certain that these fungal sequences belong in the bacterial clade. Kishino–Hasegawa tests rejected the alternative hypothesis that these sequences are fungal in origin [*C. heterostrophus* PKS24 ($P = 0.0001$) and *C. heterostrophus* PKS25 and the other members of the MSAS clade ($P < 0.0001$)]. These fungi may have acquired these PKS genes by one or more HGT events from bacteria.

Orthologs. Gene duplication and loss have left few putative orthologous matches among fungal PKS genes, and in no case

were orthologous genes found for members of all of the three classes (Sordariomycetes, Dothideomycetes, and Leotiomyces) for which we have genomic sequence (Fig. 3A). Few orthologs were observed among members of the Sordariomycetes, even within a genus (Fig. 3B); the *N. crassa* PKS genes have no orthologs in *G. zeae* or *G. moniliformis*, and only four PKS genes of 25 are orthologous between *G. zeae* and *G. moniliformis*.

Orthologous PKS genes found among closely or distantly related species reflect examples of nearly identical PKs being produced by different fungi. For example, citrinin and lovastatin are nearly identical compounds, made by closely related *P. citrinum* (3, 24) and *A. terreus* (25), respectively. Both citrinin and lovastatin require two PKs for their synthesis; *A. terreus* *LovF* and *P. citrinum* *MlcB* genes are orthologs, and *LovB* and *MlcA* genes are orthologs. We note that both pairs of proteins receive strong bootstrap support (Fig. 1) and have identical domain structures, which is why we used these criteria to predict orthologs among the uncharacterized proteins. Another example of orthologs that fit these criteria, *C. heterostrophus* PKS1 (6) and *D. maydis* PKS1 (26), synthesize nearly identical compounds (T- and PM-toxins, respectively), and the biological activities of these toxins on the fungal host, maize, are identical (24). In a third example, *Dothistroma pini* makes dothistromin with products of genes that are orthologous to those of the aflatoxin biosynthetic pathway found in *Aspergillus* spp. (48).

PKS Diversity in Ascomycota. Given that a fungal species has as many as 25 PKS genes, our small sampling of fungal genomes predicts that an astonishingly large number of PKs are produced by species of Pezizomycotina. The unknown PKs may be toxins

and pigments, or they may be involved in developmental processes that require small molecule signaling, such as growth and hyphal fusion, aerial hyphae formation, conidiation, and sexual reproduction. For example, in *Dictyostelium discoideum*, a type II PKS produces the 12-carbon PK precursor for a diffusible signaling molecule (DIF-1, a chlorinated alkyl phenone), which induces the differentiation of prestalk-O cells (49).

This inventory and categorization of all type I PKS genes in the genomes of five species within the Pezizomycotina is a starting point in our understanding of the diversity of PKs generated by these multidomain enzymes. A better understanding of the function of these compounds requires the characterization of PKs, including their modification and decoration by enzymes that are part of the biochemical pathway [some or all of which may be found in gene clusters (50, 51)], and the analysis of their roles in the biology of these fungi (52).

We thank Steven Briggs, President and Chief Executive Officer of TMRI, for his helmsmanship. Intermediate genome assembly of raw shotgun sequences was performed with in-house software (Don Hutchinson, TMRI). The gene predictions were created by Darrell Ricke (TMRI). Preliminary analyses of PKSs in *C. heterostrophus* were conducted by Barbara Robbertse and Scott E. Baker (TMRI). S.K. was supported by a collaborative agreement between University of California, Berkeley, and TMRI/Syngenta. The *C. heterostrophus*, *B. fuckeliana*, *G. moniliformis*, and *G. zeae* genome sequences were provided by Celera Genomics (Rockville, MD) for TMRI/Syngenta. The genome of *E. gossypii* was sequenced by SBI Technology/Syngenta (Research Triangle Park, NC). The genome of *P. chrysosporium* was made available by the U.S. Department of Energy (<http://genome.jgi-psf.org/whiterot1/whiterot1.home.html>).

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Supporting Materials and Methods

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DNA and Protein Genomic Databases. Four genomes were sequenced by Celera Genomics for Torrey Mesa Research Institute (TMRI)/Syngenta: *Botryotinia fuckeliana* strain B05.10 (to $\approx 5\times$ coverage), *Cochliobolus heterostrophus* strain C4 [American Type Culture Collection (ATCC) 48331] (to $\approx 5\times$ coverage), *Gibberella moniliformis* strain (ATCC 38932) (FGSC 7600) (to $\approx 4\times$ coverage), and *Gibberella zeae* lineage 7 strain GZ3639 (1) (to $\approx 2\times$ coverage). The shotgun sequence assemblies of these genomes were used for these studies, along with the genome sequence of *Neurospora crassa* (2), *Saccharomyces cerevisiae* (3), *Eremothecium gossypii* (4) (sequenced by Syngenta), and *Schizosaccharomyces pombe* (5). The more complete coverage ($\approx 10\times$) of *G. zeae* [lineage 7 strain PH-1 (NRRL 31084), available from the Whitehead Institute (www-genome.wi.mit.edu/annotation/fungi/Fusarium/index.html)], was used to check whether our $2\times$ genome coverage of *G. zeae* lacked any putative *polyketide (PK) synthase (PKS)* genes.

The protein databases for *Cochliobolus heterostrophus*, *B. fuckeliana*, and *G. moniliformis* were compiled by using four gene models that were found to generate the largest numbers of predicted proteins: GENEMARK.HMM (6) (*Arabidopsis* and *Caenorhabditis* models), FGENESH (7) (dicot model), and GENSCAN (8) (*Arabidopsis* model). *N. crassa*, *S. cerevisiae*, and *S. pombe* protein databases are available at NCBI (www.ncbi.nlm.nih.gov).

PKS Gene Identification. Initially, we searched our $2\times$ coverage of fungal genomes available at TMRI for *PKS* genes. Because $2\times$ coverage yielded average contig lengths shorter than the expected length of most *PKS* genes (5,000-7,500 bp), most *PKS* genes were represented as fragments. When greater coverage was available, the process was repeated, but no additional genes were found. We used the ketoacyl synthase (KS) domain, the most highly conserved domain in type I *PKS*s and in fatty acid synthases (FASs) (9), as a query. The KS domain is composed of a KS_n Protein Families database Pfam00109.6) and a KS_c (Pfam2801.6) domain (www.sanger.ac.uk/Software/Pfam). The consensus sequences of the two Pfams were concatenated and used in TBLASTN to retrieve *PKS* genes from each genome. In addition, to find *PKS* genes that may have been missed in a TBLASTN search, the concatenated Pfams were used in PSI-BLAST as a query against the predicted protein databases from *C. heterostrophus*, *B. fuckeliana*, and *G. moniliformis* and against the protein databases from *N. crassa*, *S. cerevisiae*, and *S. pombe* [available in the eukaryote protein database (www.ncbi.nlm.nih.gov/BLAST)]. The PSI-BLAST searches were also used to retrieve all previously published fungal *PKS* genes from NCBI. After the initial phylogenetic analyses were performed (described below), representative KS domains from each clade were used as queries in TBLASTN searches against all databases described above to identify any previously missed *PKS* genes.

PKS Gene Annotation. The protein sequences of the KS domains obtained from TBLASTN and PSI-BLAST searches were used as queries to retrieve the complete *PKS* protein sequences from the protein databases. The *C. heterostrophus*, *B. fuckeliana*, and *G. moniliformis* *PKS*-predicted sequences from each of the four protein databases were retrieved and aligned by CLUSTALW by using the amino acid substitution matrix BLOSUM62, implemented in MEGALIGN (DNASTAR, Madison, WI). This was done to evaluate, manually, the predicted introns removed from each sequence by each algorithm. *PKS* genes frequently have large numbers of introns, particularly so in the KS domain (which can have up to nine introns each). None of the algorithms reliably removed all of the introns, nor were all of the 5' and 3' ends

correctly predicted. The majority of introns in the *PKS* genes were therefore predicted manually by using progressive alignments with previously annotated *PKS* proteins as a guide. The predicted *PKS* proteins from *N. crassa* were also aligned to verify intron predictions. The predicted proteins were submitted to Hidden Markov Model searches, implemented in TIMELOGIC (TimeLogic, Carlsbad, CA), to determine their multidomain structures.

KS Domain Genealogy Construction and Evaluation. The predicted KS domains of all newly sequenced filamentous fungal genomes plus those from all previously published *PKS* protein sequences from ascomycete fungi, the homologous FASs from animals (9), and representative type I *PKS*s from bacteria, including those that occur as nonribosomal peptide synthetase (NPS)/*PKS* hybrids (Table 1) were aligned with CLUSTALW (NEXUS file provided as Table 2) and phylogenetically analyzed with neighbor joining and maximum parsimony (MP) in PAUP4.0b8 (10). MP was performed with the following settings: gaps treated alternately as "21st amino acid" and as missing, 100 repetitions performed with random sequence addition, branch swapping by tree bisection-reconnection (TBR), and heuristic search performed with multiple trees saved and steepest descent invoked. Only the MP genealogies were retained.

The resulting KS-domain genealogy (randomly selected from the MP trees) was evaluated to rank major clades and subclades of *PKS* enzymes. The criteria used to categorize proteins in major clades and subclades were moderate bootstrap support (performed with MP, with 100 repetitions with 10 random additions per repetition, MaxTrees set to 1,000), similar domain structure, and similar chemical characteristics of the resulting PK products for those *PKS*s that have been previously characterized (e.g., *PKS*s producing lovastatin and citrinin). KS-domain genealogy was also used to distinguish orthologous genes that are expected to encode functionally homologous enzymes (11, 12). The criteria used to categorize proteins as orthologous were bootstrap values >80%, identical predicted domain structure (or nearly so), and similar chemical characteristics of the resulting PK product (e.g., ascomycete-type melanin). Putative orthologs were further evaluated by comparing their topology in the KS domain genealogy to that of the currently understood organismal phylogeny (13). The latter is applicable when three or more orthologs are present.

Alternative phylogenetic hypotheses were tested for significance with a Kishino--Hasegawa (K-H) test (14). The hypothesis was stated as a tree with a topology constraining the taxon in question as a monophyletic group. The constrained tree was imported as a PAUP file and analyzed to find the best tree given the constraint, by using parsimony and the settings described above to find the MP tree without the constraint. The likelihood of the resulting tree was compared to that of the most likely tree, given no constraint. The hypothesis underlying the constraint tree is rejected if the difference in logarithm likelihood divided by the standard deviation ($\text{Ln } L/\text{SD}$) is > 1.96 ($P \leq 0.05$) (15).

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Table 1. GenBank accession nos. for previously published sequences included in Fig. 1

Species	Protein (module)	Accession nos.
<i>Aspergillus fumigatus</i>	alb1	AAC39471
<i>Aspergillus nidulans</i>	pksST	AAA81586
<i>A. nidulans</i>	wA	CAA46695
<i>Aspergillus parasiticus</i>	pksL2	AAC23536
<i>A. parasiticus</i>	pksL1	AAC41675
<i>Aspergillus terreus</i>	at1	BAB88688
<i>A. terreus</i>	at4	BAB88689
<i>A. terreus</i>	at5	BAB88752
<i>A. terreus</i>	lovB	AAD39830
<i>A. terreus</i>	pksM	AAC49814
<i>A. terreus</i>	lovF	AAD34559
<i>Bacillus subtilis</i>	pksK module 2	NP_389599
<i>B. subtilis</i>	pksK module 3	NP_389599
<i>B. subtilis</i>	pksL module 1	CAB13602
<i>B. subtilis</i>	pksL module 2	CAB13602
<i>B. subtilis</i>	pksL module 3	CAB13602
<i>B. subtilis</i>	mycA pks1 module 2	AAF08795
<i>Bombyx mori</i>	FAS p270	AAB53258
<i>Byssochlamys nivea</i>	6MSAS	AAK48943

<i>Caenorhabditis elegans</i>	FAS	NP_492417	
<i>Colletotrichum lagenarium</i>	pks1	BAA18956	
<i>Gallus gallus</i>	FAS	P12276	
<i>Glarea lozoyensis</i>	PKS1	AAN59953	
<i>Glycine max</i>	beta-ketoacyl synthetase	AAF61730	
<i>Gibberella moniliformis</i>	FUM1	AAD43562	
<i>G. moniliformis</i>	PKS1	CAC44633	
<i>G. moniliformis</i>	PKS3	CAC88775	
<i>G. moniliformis</i>	PKS4	CAB92399	
<i>Homo sapiens</i>	FAS	AAC50259	
<i>H. sapiens</i>	3-oxoacyl-ACP-synthetase	NP_060367	
<i>Microcystis aeruginosa</i>	mycE	AAF00958	
<i>M. aeruginosa</i>	mycG	AAF00957	
<i>Monascus purpureus</i>	PKS1	CAC94008	
<i>Mycobacterium leprae</i>	pksE	AAA17364	
<i>Mycobacterium tuberculosis</i>	mas	CAB06108	
<i>M. tuberculosis</i>	ppsD	CAB06093	
<i>M. tuberculosis</i>	ppsE	CAB06094	
<i>Myxococcus xanthus</i>	ta1 module 2	CAB38084	
<i>Neurospora crassa</i>	PKS1	XP_325868	
<i>N. crassa</i>	PKS2	XP_324368	
<i>N. crassa</i>	PKS3	XP_324222	
<i>N. crassa</i>	PKS4	XP_329445	
<i>N. crassa</i>	PKS5	XP_330288	
<i>N. crassa</i>	PKS6	XP_330106	

<i>N. crassa</i>	PKS7	XP_322886	
<i>N. crassa</i>	3-oxoacyl-ACP-synthetase	AAB81078	
<i>Nodulisporium</i> sp.	melanin PKS	AAD38786	
<i>Nostoc</i> sp. GSV224	nosB	AAF15891	
<i>Penicillium citrinum</i>	mlcA	BAC2056	
<i>P. citrinum</i>	mlcB	BAC20564	
<i>Penicillium griseofulvum</i>	pks2	AAB4968	
<i>Penicillium patulum</i>	6MSAS synthetase	CAA39295	
<i>Pseudomonas syringae</i>	cfa6	AAD03047	
<i>P. syringae</i>	cfa7	AF098795	
<i>Sorangium cellulosum</i>	epoA module 1	AAF26919	
<i>S. cellulosum</i>	epoC module 2	AAF2692	
<i>S. cellulosum</i>	epoF module 9	AAF62885	
<i>Stigmatella aurantiaca</i>	mxkB2, B1	AAK57186+AAK57185	
<i>S. aurantiaca</i>	mxnC1	AAK57187	
<i>S. aurantiaca</i>	mxnC2	AAK57187	
<i>S. aurantiaca</i>	mxnC3	AAK57187	
<i>S. aurantiaca</i>	mxnD	AAK57188	
<i>S. aurantiaca</i>	mxnE	AAK57189	
<i>S. aurantiaca</i>	mxnF	AAK57190	
<i>Streptomyces natalensis</i>	PimS0	CAC20930	
<i>S. natalensis</i>	PimS2 module 3	CAC20921	
<i>S. natalensis</i>	PimS2 module 4	CAC20921	
<i>Streptomyces viridochromogenes</i>	aviM	AAK83194	
<i>Streptomyces coelicolor</i>	act typeII	NP_823552	

<i>Streptomyces avermitilis</i>	pks9 typeII	NP_823551	
<i>Xanthomonas albilineans</i>	xabB module 2	AAK15074	
<i>X. albilineans</i>	xabB module 3	AAK15074	
<i>Xylaria</i> sp.	PKS12	AAM9354	

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Table 2. KS domain alignment used for phylogenetic analyses in NEXUS format

#NEXUS

```
begin data;
  dimensions ntax=144 nchar=486;
  format datatype=Protein interleave gap=- missing='.';
  matrix
[
  1
  60]
Bf_PKS7      AIAIVGNSCRLL--PGGSHCPSKLWDL-LKSPIDLVAEIRQNRFN-----AGFYHPRAEHS
Aspergillus_terreus_at1 AI--VGMSCRFL--PGGADDPERFWEL-LEEGRDVHKKVPADRFVDLHCDPTDKRK----
Aspergillus_terreus_at4 AI--VGYSGRF--PSA-ASNEAFWEL-LRAGRVDHREIPDRDFDWEITYDPTGKRR----
Aspergillus_terreus_at5 AI--VGMAGRF--PNS-DDLESFWST-LRQGLDLHRRVPPDRFDIDAHYDPTGKKL----
Gm_PKS5      AVIGSVAISRMLMRAKTHEINHSRTI-LTTAR-VIHEVRPNRMKN----KSIMSMLLSLI
Ch_PKS9      AV--VGFQFKF--PQDVTNAETLWKL-LIERRSTMTETPKNRWNI----DGFYKENGHRP
Ch_PKS8      AI--IGMNMKF--PGDAVSAQSFWKL-IVSARNVSKVEVPADRFKI----DSFYHDPNRL
Gz_PKS2      AIV--GYSFKL--PQGVEDDDAFWDV-LENRRNLMTDWPESRVKT----DSFAHPDRSS
Gm_PKS2      AIV--GYSFKL--PQGAEDDDVFDV-LENRRNLMTDWPESRVKT----DSFANNKH---
Ch_PKS10     AVI--GLDAKL--PCGDGDSVQQFFDF-LIAGRSARKVPVPSDRYNA----DAFWHPD-HHR
Ch_PKS5      AI--VGLATRF--PQEAYNTEQLWQF-LLAKRCAHTPIPEGRMGPF----GHYHPDPEHG
Ch_PKS7      AI--VGMSCRF--PGSANNIENLFKM-ICEKRESRTAIPKSRWNN----KAFYHPNQRH
Ch_PKS1      AV--VGMFRG--PGDATNVEKLLNM-ISEGRESRAEVQAKKWDPF----EGFYHPDSSRH
Didymella_maydis_PKS1 AI--VGLSFRG--PGDATSAENLLRM-VAESRESRSPGPSQKRNA----SGFYHPDPSRQ
Gz_PKS4      AI--VGLACRF--PGDATSPSKFWDL-LKSGKDAYSE-TTDRYNA----QAFYHPNSKRQ
Gz_PKS7      AV--VGMSCRF--PGDASSPEELWKV-LERGESQWSEFPKDRFNI----EGFYHPDGRQ
Gm_PKS7      AI--VGIGLRF--PGDASSPEELWKV-LERGESQWSEFPKDRLNI----DGYHPSGDRQ
Ch_PKS6      AI--IGIAFRG--PGDARDPEAFYRM-LIEGRSARTEIPKDRYNV----DAFYHPDPERL
Gz_PKS1      AV--IGIGCRF--PGNSSNPEALWKL-LSDARSMYSKVPVPPDRYNV----DGFHPSNKLN
Ch_PKS4      AI--VGMSCRF--SGMADTPGAFWQM-LSKGMTSWSRDARDFRKL----ESFVHPDNDLS
Gm_PKS6      AV--IGVSGRF--PGDATSPRHLWDL-LKEGRNALS DVPESTRFNI----DGFYHPDGGRA
Nc_PKS1      AI--VGMACRM--PGNVSTPAEFWEL-CTRARS GWSETPKQRFNS----ARFHHPNQGGK
Bf_PKS8      AI--VGISCRF--PGNVSTPDEFWEL-CSTRARS GWSETPKERFDT----ASFYHPNPGKA
Nc_PKS2      AI--VGMSCRF--PGGVETPGDFYRM-MCRGRSGWSKVPKDRFNQ----EGFYHPDPDHK
Ch_PKS2      AI--IGMSCRFL--PGKVASLEDFWDM-LSNSKHGYSRQFPRERFNW----EAFYHPNQSRK
Aspergillus_terreus_lovF AM--VGMGCRF--GGGATDPQKLWKL-LEEGGSAWSKIPPSRFNV----GGVYHPNGQRV
Penicillium_citrinum_mlcB AI--VGMGCRF--AGDATSPQKLWEM-VERGGSAWSKVPSSRFNV----RGVYHPNGERV
Ch_PKS3      AI--IGMSCRFL--AGGATDPEKLWQL-CAEGRTGWSEIPEDRFKI----DGHYHPDNDL
Nc_PKS3      AI--IGMSCRFL--AGDVDSPSKLWDL-LAQGKSAWSEIPKDRFNI----DGFHHPNFEKL
Bf_PKS2      AI--VGISCRF--PAGANSPEELWSL-ISQKSAWS DVPEDRFNW----KSFHLPSSDVV
Gz_PKS5      AI--IGVGCRL--PGGANLNDLWKL-LSES RNGQTEIPKDRWNA----DAWFDAYPDAK
Gm_PKS1_CAC44633 AI--IGSACRF--PGDSSSPSKLWDL-LKSPRDL LTKVPPNRYNA----DAFYHADSKHH
Gm_PKS1      AI--IGSACRF--PGDSSSPSKLWDL-LKAPRDL LTVPSNRYNA----DAFYHADSKHH
Bf_PKS5      AI--IGMACRF--PGGSNSPSKLWDL-IKSPRNLSKRVPDERFNS----EAFHHTNGSYH
Gm_PKS9      AV--VGTACRF--PGGNTTPSKLWDL-VFSSRDV LQKVPARFNV----DSFYHSDPTH
Aspergillus_terreus_lovB VV--VGSACRF--PGGNTTPSKLWEL-LQHPRDVQSRIPKERFDV----DAFYHADSKHH
Penicillium_citrinum_mlcA VV--VGSACRF--PGGNTTPSKLWEL-LKEPRDVQTKIPKERFDV----DTFYSPDGTHP
Gz_PKS9      .....SKLWDL-LKRPKDVSKETIPEDRFNL----DRFYHKDS SHH
Ch_PKS16     AI--VGSACRF--AGGANSPSKLWDL-LRNPKDIRSNITDKRFNA----NGFYHPDGSHH
Bf_PKS6      AV--IGTGCRF--PGGASSPSKLWNL-LHHPYDLTQKVPSSRFNI----KAFYHPNGEHH
Bf_PKS3      AI--IGSSCRF--TGSANT-SKLWEL-LKEPRDVLSKIPERFLA----EGFYHQDQHH
Ch_PKS17     AV--VGVGCRF--PGGNTTPSKLWDL-ICEQRDIQSKIPAERYNS----DAFYNEDGSKP
Gz_PKS10     AI--IGTFCRF--HGGANTPSKLWDL-LIEKRDVQSSIPPVRFNV----DAFIIVM-DKN
Gm_PKS10     AI--IGTSCRF--PGGANTPSKLWDL-LCEKRDVQSRIPNDRFNV----DAFYSTNGDKN
Nc_PKS4      AI--VGMSCRF--TGEATSPAKLWEL-LKDPKDLTREVPKDRFNV----EGFYHPDGEYH
Bf_PKS4      AI--IGRGCRF--AGGCQSPSELWNF-LQTPYDV SQEFPDR-----NSASRH
Gz_PKS11     AICGMSV----RLPGGLHSPQDLWEL-LVSKGDARGVPVPSRYNA----SSYWSEKM-KP
Ch_PKS14     ICGMGL----RLPGGNTTPQEFWEL-LINKGDGRVVRVPMSTRYNA----SAYHETTK-RP
Ch_PKS12     AICGMAI----RLPGALKTPAQFWTF-LTEKDKARAPIPFARFDA----EAFYSASG-KS
Gm_PKS12     AICGTAL----RLPGGISNCQDYWDL-LYHGLDARRPIPSRFNI----NGFNDSL G-K
Ch_PKS13_pseudogene TIVGMAM----RLLGVRHNVDELWEM-LIAKKSGLVDILKKRYNA----EGFQSDTP-MV
Ch_PKS11_FUM1 AIVGLGM----RLPGAHTAEQLWKT-IVQKRSTRCEIPASRFVS----DGFHSPSA-KP
Gm_FUM1_AAD43562 AIVGMGM----RLPGGIHTPDELWGM-LVEKRSTRCEIPPTRFVS----DGFHSPSS-KP
Gm_FUM1      AIVGMGM----RLPGGIHTPDELWGM-LVEKRSTRCEIPPTRFVS----DGFHSPSS-KP
Ch_PKS15     AIVGMGM----RLPGNCHNSEAFWDL-LINQRDGMIDIPSSRWNS----EGFYDPHG-RP
Nc_PKS5      AI--VGMACRW--PGGVHDPSQFWEF-LRNKVNWGKEFDDPRFSS----AGFHHPNSDRP
Gm_PKS15     AI--VGMSCRF--PGGVNPKEFWEF-LVNGESGYSDFPEERLNI----DSWYHPESVRP
Nc_PKS6      AI--IGMGCRF--PGGSHSSKLWEL-LKAGRTAQSFRPSSRFNI----DGFYHPNSDRP
Gm_PKS14     CI--VGMACRF--PGGIRSPDDLWDF-MIQKESAYGVPVADRYNI----DGFYHHQNLS
Bf_PKS9      .....CRL--PGGLTTPDELWDF-LLAKKDARCRVPHSRDYI----DSYY-SDTKKP
Gm_PKS13     AI--VSMACRF--PGGIDKPLDLWDH-VRAGRSSATAIPKDRFNA----ENFLSMDPNQK
Bf_PKS10     AI--IGFACRF--PGGNTTPKLLWEL-LERGDIASNKVPSRFNI----EGHWD-GSQKP
```


Stigmatella_aurantiaca_mxaC2
Stigmatella_aurantiaca_mxaC3
Stigmatella_aurantiaca_mxaD
Stigmatella_aurantiaca_mxaE
Stigmatella_aurantiaca_mxaF
Streptomyces_natalensis_PimS0
Streptomyces_natalensis_PimS2_mod3
Streptomyces_natalensis_PimS2_mod4
Streptomyces_viridochromogenes_aviM
Xanthomonas_abilineans_xabB_mod2
Xanthomonas_abilineans_xabB_mod3

AI--VGMGCRF--PGGANPEAYWRL--LSDGVDAVREVPADR---WQSSSTA-AEHKGT-
AI--VGIGCRV--PQAS-GPEAFWKL--LEGGVDAIREVPASR---WKVEDYY-DPQPGVP
AI--VGTSCRV--PGGARTPEAFWRL--LHGGVDAITEVPRDR---WDAEALF-RDDEPNKQ
AI--IGMGRF--PGGNDPEYWNL--LCNGVDAVTEVPSSR---WTRREME-DMPEAL
AI--VGMACRY--PQ--ASDPEAFWRL--LAGGNHAIETIPPER---WDINRLY-DRDPAAA
AI--VGMACRL--PGGADSPDALWEL--LADGTDAMSPFPTDRG---WDLDRLF-DEDADRP
AI--VGMSCRV--PGGVRSPRLWDL--VSEGVDAISDFPVNRG---WNT-GLF-HPDPDNP
AI--IGMSCRY--PGGVSPEDLWRL--VTEGGDATGEFPADR---WDAEGLY-DPDPDRA
AV--VGIGCRF--PGGVNSPGEFWDL--LTAGRNTVGEPPDR---WEYRDF-GPRFDAA
AI--VAMHCEV--PGAGENTALWSF--LRSDVNAIRPIESTRFDLWAMRAY----PGLA
AI--IGLAGRF--PGADTLEE--FWNN--LRNGQSSMGEVPGER---WD-HQHYFDSERQAP

[
Bf_PKS7
Aspergillus_terreus_at1
Aspergillus_terreus_at4
Aspergillus_terreus_at5
Gm_PKS5
Ch_PKS9
Ch_PKS8
Gz_PKS2
Gm_PKS2
Ch_PKS10
Ch_PKS5
Ch_PKS7
Ch_PKS1
Didymella_maydis_PKS1
Gz_PKS4
Gz_PKS7
Gm_PKS7
Ch_PKS6
Gz_PKS1
Ch_PKS4
Gm_PKS6
Nc_PKS1
Bf_PKS8
Nc_PKS2
Ch_PKS2
Aspergillus_terreus_lovF
Penicillium_citrinum_mlcB
Ch_PKS3
Nc_PKS3
Bf_PKS2
Gz_PKS5
Gm_PKS1_CAC44633
Gm_PKS1
Bf_PKS5
Gm_PKS9
Aspergillus_terreus_lovB
Penicillium_citrinum_mlcA
Gz_PKS9
Ch_PKS16
Bf_PKS6
Bf_PKS3
Ch_PKS17
Gz_PKS10
Gm_PKS10
Nc_PKS4
Bf_PKS4
Gz_PKS11
Ch_PKS14
Ch_PKS12
Gm_PKS12
Ch_PKS13_pseudogene
Ch_PKS11_FUM1
Gm_FUM1_AAD43562
Gm_FUM1
Ch_PKS15
Nc_PKS5
Gm_PKS15
Nc_PKS6

61 120]
GTSNVTKAYLLEG--DPWTFDNEFFKISAREAESMDPQQR.....
NTSLTPFGCFIDQ---PGLFDAGFFDMSPREAANTDPMHRLALVTAYEALESQSGF-----
NTRSVYKGFIDQ---PGLFDTKFFNMSPRKPENTDPAQRALITTYEAMEMMAGM-----
NSTHTPYGCFIEK---PGLFDPRFFNMSQREAYQTDPMGRLLALVTAYEALEMESGF-----
KTFALR-AHFLKG--ETAADFAPFFSITPAEAGGMDPQQRGLL----ENTYRA-LENAGC
GTVKNRGGHFLSD--DPAFADAPFFSIPAEAECEMDPQQRLLL----ETSYHA-LENAGI
DSIRIRNAHFMTD--DPAFADAPFFFNMSHAEASVLDPPQQRGLL----EGAYRT-FENAGI
HSRRKKLRRWIRC--.....EGVLTA-CLVACL
QKWNGKGGHFISD--DAAAFDAPFFSLTAKEASAMDPQRWTL----EATYHA-FENAGL
DAIASKQGHFMNS--SVKAFDAPFFGITPAEAAAALDPQQRLLL----ESSYLA-LENAGY
GTHAV--AHFLAE--DPYFDSFFGITKGEAMSLDPQQRVVL----ENVYLA-LENAGS
GSHNVEYGHFFHD--DISKFDAPFFNMTKEEAAAALDPAQRLLL----ESTYEA-LENGGI
GTHNVEYGHWFQ--DVYNFDAPFFNVPAAEAAAALDPQQRMLL----ECSYEA-FENSGT
GSHIVEYGHWFQ--DVYEFDAPFFNLSAVESAALDPQQRMLL----ECTYEA-FENSGM
NVLPVTVGGHFLQ--DPHVFDAAFFNITAAEAI SLDPKQRIAL----EATYEA-FENAGK
GSI SFRGAHFIKS--DFAAFDASFFFAVAEDAKAIDPQQRILL----EASYEA-LENAGI
GSI SFRGAHFIKG--NFASFDAFFSISAEDAQAIDPQQRILL----EASYEA-LENAGI
GSIQQRyahflq--DFKAFDAPFFSITPKEAKAMPDTHRILL----EAAYEG-FENAGL
TTIA-EGAHFLSE--NIAAFDAAFFNIAPIDAKSMDPQQR.....GI
GSFNASGLHLLRQ--NPAAFDNDFFSISGLEAKAMPDQQRMLL----ELAYET-FENAGL
GTLNTKQGYFLKS--DVKDFDAGFFSITPEEARGMDPTQRILL----ELAYEG-LENAGL
GTLNVPVGGNFLNV--DLAAFDAFFFLTEKEAISMDPQQRLLL----ECTFEA-LESAGI
GCFNAAGNFLE--DVLFDAPFFSLTAQEATSMDPQQRILL----ECTFEA-LESAGI
GTFNSQGGYFIKQ--DLSGFDAAFFDVTRREAEAMDPQQRLLM----ECTYEA-LESAGI
DCIDVNCGYFLDG--DIAEFDAQFFKMGNTDA--DPQGRMILL----ECVYEALENAGGS
GSMHVRGGHFLDE--DPALFDASFFNMSTEVASCMDPQYRLLI----EVVYEA-LEAAGI
GSTHVKGGHFLDE--DPALFDAAFFNMTTEVASCMDPQYRLLI----EVVYEA-LESAGI
NTTNVKGACFLDE--DVGNFDAFFNLPAAETA AV-----SIIII----CSSLPC-ILTKGN
NGTNVIGGHFMKE--DVLFDAAFFNLSAETAALDPQFRLLQ----ESTYEA-LESAGI
GTINSRGGHFIQ--DIRTFDAGFFGIPPAEANAMPDQHRLLQ----ETAYEA-LENAGI
QSMVTKYGYFLQ--DISQDAKFFGISSAEANSMDPQQRLLI----ETVYEA-LEDAGI
GTTNVRHSYFLNE--DPAFDDNFFNIQPGAEAEIDPQQRLLM----EVVYQG-LCASGQ
GTTNVRHSYFLSE--DPASFDDNFFNIQPGAEAEIDPQQRLLM----EVVYQG-LCASGQ
GATDSREAYFLEE--DVALFDNAFFNIQPGAEAEAVDPQQRLLM----ETVYDS-LCAGGQ
GTTNVTQSYFLDE--DVTKFDNGFFGIQPMEEAEAMDPQQRLLI----ETVYDS-VVDAGL
GRTNAPYAYVLQD--DLGAFDAAFFNIQAGEAEAMDPQHRLLL----ETVYEA-VTNAGM
GRTNAPFAYLLQE--DLRGFDASFFNIQAGEAETIDPQQRLLL----ETVYEA-VSNAGL
GTANVRRSYLLDE--DVRLFDQFFGISPGEAQAMPDQHRVLL----EVVYEA-IESAGK
GHMNVKQSYLLEE--DTRLFDAAFFGNIQVAKAMPDQQRLLI----EVVYEA-IESAGL
GTHNATKSYFLNE--DPTTFDAPFFNINPREAEALDPQQRLLI----ETVYEA-LEAAGL
GTSNVLHSYLLDE--NPLAFDASAFFHIHNREAEICIDPQQRLLI----ENVYET-IESACY
GCTNVNHGYFLDE--DIRAFDAAFFRMPAEVEAVDPQQRMLL----ETVFEA-LESAGT
GCTDVKKAYLWAE--DTRLFDASFFKINPREAEAMDPQQRLLI----EAVYEA-TEAAGL
GCTDVKKAYLLSE--DIRVFDASFFKINPREAEAMDPQQRLLI----EAVYEA-TEAAGL
GTTNSAKAYFLEQ--DHRFLDASFFNITPKEAEAEIDPQQRMLL----EVVYEA-LESAGY
STS--TRGYFLSQ--DVREFDASFFSLSPLEAQAMPDQHRLLL----ETVYEA-LEEAGI
GTVTKYGYFLDESIDIASVDTSFFTMRKDEVGKADPQQRQML----EVAREC-LEDAGE
TTVATEYGYFLDEDVKGAMDTSRFMSRADVEFSDPQQRHLL----EVVMEA-LEDAGE
GYFNKNGYLLDSDAEVGLDTSLFRMSQSESERMDPQAKILL----ELAREC-FESAGE
DSIKTRHGYFIED--DLSRLDTSFFSLTKNELERVDPQQRLLL----EVTHEC-LEDAGE
GTMQTKHGYFLND--IDLTFDALFFSFSYKEAERIDLAQRQLL----EIIYEC-LENVGE
GSMVRRHGHLDFDLHRLDTSLSFSMGITVNDIDPQQRMLL----EVVYEA-LESAGQ
GSIAMRHGHFLDDKDLHRLDTSFFSMGMTEVSDIDPQQRMLL----EVAYEC-MQSSGQ
GSIAMRHGHFLDDKDLHRLDTSFFSMGMTEVSDIDPQQRMLL----EVAYEC-MQSSGQ
GTTKVNKGNFLG--SIDPADFDGSSFFMSAAQIAKVDPQHRVLL----ETTYEA-LENAGE
GSMKLEGAFLAQ--DARLFDHTFFGMTGLEVETMDPSQRKLL----EVAYEA-IESAGE
GSVVTKGGYFLKH--DMNEFDNEFFGITAAEARLMDPAQKQML----EVVFES-AEAGV
GSLNMEGGYFIED--DIRGFENSFFGINNLEATYMDPQQRKLL----EVVFET-FENAGF

Sorangium_cellulosum_epoC_mod2
Sorangium_cellulosum_epoF_mod9
Stigmatella_aurantiaca_mxaB2_B1
Stigmatella_aurantiaca_mxaC1
Stigmatella_aurantiaca_mxaC2
Stigmatella_aurantiaca_mxaC3
Stigmatella_aurantiaca_mxaD
Stigmatella_aurantiaca_mxaE
Stigmatella_aurantiaca_mxaF
Streptomyces_natalensis_PimS0
Streptomyces_natalensis_PimS2_mod3
Streptomyces_natalensis_PimS2_mod4
S_viridochromogenes_aviM
Xanthomonas_albilineans_xabB_mod2
Xanthomonas_albilineans_xabB_mod3

AGSVLE-----DVDRFDAAFFGISPREAELMDPQHRIFMCEAWALEENAGYDPTAY
----RWAGLLTE--AVDGFDAAFFGTSPREARSLDPQORLLLEVTWEGLEDAIAPQSL
GKMYGTRGGFLDD---VEHFDAEFFGISPREAASLDPQORLLEVAWEALEENAGLAPDQL
----RWGGFL-D--SVDSFDPFFGGISPRAVAVRMDPQORLLEVAWEALEENAGLDPVKL
----RWGGFL-D--RVDGFDPEFFGIAPRAVAVMDPQORLLEVAWEALEENAGLPKSRL
GKTYTRWGGFLDG---ADQFDAQFFGIAPRAANMDPQORLLEVAWEALEENAGIAPARL
GRTYARWGGFIDG---VDRFDAAFFGVSPREASRMDPQORLLEVAWEALEERAGQPPDQL
EKLGARWGAFFI-N--KVDQFDADFFGISPREAHRMDPQORLLEVAWEALEERAGQDMTQL
GKVTSRWGAFLD---QVDGFDPLFFGISPKALHMDPQORLLEVAWEALEEDAGIAAEBGL
GTSYAREGGFLHD---AGDFDAGFFGLSDQEATATDPQORLLEAAWETFERAGIDPQSL
GTTYSTQGGFLHD---AGEFDASFFGISPREALSMDPQORLLETTWEAFEHAGIDPTTV
GHTYSTRGGFLHD---AADFDASFFGISPREALAMDPQORLLETSWEAMERAGIDPATL
LRTAIRSGSFLDD--DIAGFDAEFFGISPREAELMDPQORLLEVAWAQALEHAGIPPHL
GEQLPRYAGFLDD---VDAFDAAFFGISRRAECMDPQORLVLEMVWKLIEQAGHDP LSW
GKTYSRWGAFLRD---IDGFDAAFFWFSDVALES DPQARIFLEQAYAGIEDAGYTPGSL

[
Bf_PKS7
Aspergillus_terreus_at1
Aspergillus_terreus_at4
Aspergillus_terreus_at5
Gm_PKS5
Ch_PKS9
Ch_PKS8
Gz_PKS2
Gm_PKS2
Ch_PKS10
Ch_PKS5
Ch_PKS7
Ch_PKS1
Didymella_maydis_PKS1
Gz_PKS4
Gz_PKS7
Gm_PKS7
Ch_PKS6
Gz_PKS1
Ch_PKS4
Gm_PKS6
Nc_PKS1
Bf_PKS8
Nc_PKS2
Ch_PKS2
Aspergillus_terreus_lovF
Penicillium_citrinum_mlcB
Ch_PKS3
Nc_PKS3
Bf_PKS2
Gz_PKS5
Gm_PKS1_CAC44633
Gm_PKS1
Bf_PKS5
Gm_PKS9
Aspergillus_terreus_lovB
Penicillium_citrinum_mlcA
Gz_PKS9
Ch_PKS16
Bf_PKS6
Bf_PKS3
Ch_PKS17
Gz_PKS10
Gm_PKS10
Nc_PKS4
Bf_PKS4
Gz_PKS11
Ch_PKS14
Ch_PKS12
Gm_PKS12
Ch_PKS13_pseudogene
Ch_PKS11_FUM1
Gm_FUM1_AAD43562
Gm_FUM1

121
.....
---VLNRTPATHCKRVGTFYQGASDDYREASSGQ-N-----VDTYIITGGCRAFATG
---VRNRPSTQDRVGVFFGTTSDDWRGVNSGQ-D-----VGTYFIPGNRAFPVPG
---VFNRTSSMLDRIGTFYQTSDDWRTLNAE-K-----IDMYIIPGTIRAFATS
P--MNKVLGTNTGVFIGCFT----REYEAIMFKET----EQQRYFA-TGTGTT--MLAN
T--MQDAVGTRTSVHVGCLL----QEYSQISQRDA----QMPGDYRI-VGSS-GLAMLSN
P--MESLRSRTPTSVYCASFS----RDGETITGRDF----ASQRYHA-TANG-S-SMLSN
P--VDSLKGSRTAVFSASML----EDYCRVTAVDP----DNLERTAV-TGSIVS-CIIPN
P--VESLKGSRTAVFSASML----EDYSRMTIVDP----DNLERTAV-TGSTVS-CIIPN
T--LQELGNSNTGVYTGFSV----YDYRDVMIKDT----DVAL-T-Y-SGTGTVPSTLAG
T--LEQVAGSNTSVYVSGFS----HDHLAILNSDP----ETTLRHRV-T--GLTNSMHSN
P--LEKIVGKTSTVVFVGSFA----TDYDILLTRDP----ESVPMYQC-TNSGQSRAMISN
P--MSKIVGTDTSVVFVSSFA----TDYDMLWRDP----ESVPMYQC-TNSGFSRSNLAN
P--LNKLVGTDTSVFTAVFC----TDYDMLWRDP----EMVPMYQC-TNSGATRANMAN
P--LKQVAGTTTACFVGSFS----SDYRDAVVRDF----AHNPKYHV-LGTCEE--MIAN
R--KEDVDGTDAAVYVGSFV----KDYEQVCLRDP----DWQPQYAA-TGNIA--IMAN
R--KEDIDGSDAAVYVGSFV----KDYEQVCLRDP----DWQPQYAA-TGNIA--IMAN
T--LDQVSGTQTSYIGTFT----ADFPNLQARDN----EGPSIYHA-TGMS-S-SLANS
R--MEDLTGSTSCYVGTFT----RDWSDMLMRDP----ETAPKYSG-ATG-KFFLVSN
T--IKALEKSKTGVYCASTY----QDYDQILGRDP----ELSAKYRF-TGTGAS--MLAN
K--IDEVANQHMSCYIGACQ----HDYWDLQAYDM----DSAPKYTA-TGTG-P-ALLSN
PK--HTIAGKDVGVFVGGSF----PEYESHLFRDP----DTIPMHQ--ATGCAYAMQSN
PK--HELVGKDVGVFVGGSF----SEYESASFVDT----DSVPMYQC-ATG-KFFLVSN
PR--EHIAGTRTGVFIGGNY----GEHRFHIRDL----DTIPSPFD--ATGNQPAFLSG
Q--ESLVGSKVGVFSTNT----SDYTLSLKDDI----YSMPALV--GVLATACMLSN
P--LEQVSGSKTGVFAGTMY----HDYQGSFQRQP----EALPRYFI-TGNAGT--MLAN
T--IDMGAGSNTSVFVGVVY----HDYQDSLNRDP----ETVPRYFI-TGNSGT--MLSN
SAWIPSSGSCSRAPTRPSRM----QDYHDAGLRDV----TTLPRFFL-VGVGSA--MASN
T--LQDVAGSNTSVYAGSFF----RDYHESLIRDP----DTLPRFLL-MGTGAA--MASN
S--LEKVRGSSTSVYVAIFN----RDYDRMMFKDT----NDIAKYHL-LGSGEA--IASN
P--VEVTRGNSNTGVYASIFE----RSYDRMGHKDL----STIGTYM--ATGKFFLVSN
T--IEGLRGSPTAVYVGVVY----DDWSGIIITRDL----EVFPQYG--ATGMARSIMSN
T--IEGLRGSSTSVYVGVVY----DDWNGILTRDL----EVFPQYG--ATGMARSIMSN
T--IEGLRGSNTGIYVGMVY----DDWAQAINRDW----ESTMTYA--ATGQSRAIVSN
R--PEGLRQSTSVFVGVVY----DDWAQMLAKDW----DIIPTYM--ATGVSRAVMSN
R--IQDLQGSTAVYVGVVY----HDYETVSTRDL----ESIPTYS--ATGVAVSVASN
R--IQDLQGSSTAVYVGVVY----HDYETIVTREL----DSIPTYS--ATGVAVSVASN
T--IHGLHNSDTAVYVGLMC----TDYVVIQAADL----NSVPTYN--ATGVANSNASS
S--IDRLGSDTAVFAGLMC----GDYBARMRLDL----GDYPTHF--ATGTSRAVMSN
S--IEEMQGSTAVYVGLMC----ADYFDVLMRDI----EDIPQYL--ATGTARSIMSN
P--METIRGSDTGVFVGLMC----ADYDVMQMRDP----ETLPQYF--STGTARSIVSN
T--IDQLQGSTAVYVGCMT----SDYVEMLLRDP----LDFPKYM--APGTARSILSN
P--MEDLKGSDTAVYVGCMT----GYYHEMLMREP----QDMPKYM--ATGTARSILSN
P--MEDLKGSDTAVYVGCMT----GDYHEMLMRDP----QDMPKYM--ATGTARSILSN
T--LQDYSGKKVAVFAGVMT----ADYDTLSQRDD----LSVSQYY--ATGNARSISN
P--AETLRGSDTAVYTGVMF----HDYLSLSQDH----MAIPKYH--ITGTAPNKASN
FDW----KGRPIGCFMGSFG----EDWVEMFAKEA----QQYGIYRV--MGYGDFFLSN
ANF----RGKIGCYFVGNMC----EDWGEEMNRDP----LWHPYKI--DGYQWMLAN
TDW----RGKIDIGTYVGSFG----NDWLEMAAKDR----LDKNMYKV--SGYSDFLLSN
ITY----RGKQVGCYVGTGFG----DDWLIMNAKEP----LQGGLYAT--TGGADLMMAN
TEW----RSKEIGYVGSFG----EDWLNENIMDA----HMLGIYRA--LRFIDFFLAN
VNW----QGSNIGCYVGVVY----EDWLDLHAKDL----FDSGYTRV--SGSHDFAISN
TNW----RGSNIGCYVGVVY----EDWLDLHAKDL----YDSGYTRV--SGGHDFAI SN
TNW----RGSNIGCYVGVVY----EDWLDLHAKDL----YDSGYTRV--SGGHDFAI SN

Ch_PKS15
Nc_PKS5
Gm_PKS15
Nc_PKS6
Gm_PKS14
Bf_PKS9
Gm_PKS13
Bf_PKS10
Bf_PKS11
Penicillium_patulum_6MSAS
Bysochlamys_nivea_6MSAS
Aspergillus_terreus_pksM
Aspergillus_parasiticus_pksL2
Penicillium_griseofulvum_pks2
Ch_PKS25
Nc_PKS7
Colletotrichum_lagenarium_pks1
Nodulisporium_sp_PKS1
Xylaria_sp_PKS12
Bf_PKS12
Glarea_lozoyensis_PKS1
Monascus_purpureus_PKS1
Bf_PKS13
Ch_PKS18
Bf_PKS14
Ch_PKS20
Aspergillus_nidulans_wA
Aspergillus_fumigatus_alb1
Gz_PK12
Gm_PKS3_CAC88775
Gm_PKS3
Gz_PKS12
Gm_PKS4_CAC92399
Gm_PKS4
Aspergillus_parasiticus_pksL1
Aspergillus_nidulans_pksST
Gz_PKS13
Bf_PKS15
Bf_PKS19
Ch_PKS22
Bf_PKS16
Ch_PKS19
Ch_PKS23
Ch_PKS21
Bf_PKS17
Bf_PKS18
Bf_PKS20
Homo_sapiens_FAS
Gallus_gallus_FAS
Caenorhabditis_elegans_FAS
Bombyx_mori_FAS_p270
Ch_9497_PKS24_3p_of_nps7
Nc_3_oxoacyl_ACP_synthetase
Homo_sapiens_3_oxoacyl_ACP_synth
Glycine_max_beta_ketoacyl_synth
Streptomyces_coelicolor_act
Streptomyces_avermitilis_pks9
Bacillus_subtilis_pksK_mod2
Bacillus_subtilis_pksK_mod3
Bacillus_subtilis_pksL_mod1
Bacillus_subtilis_pksL_mod2
Bacillus_subtilis_pksL_mod3
Bac_subtilis_mycA_pks1_mod2
Microcystis_aruginosa_mycE
Microcystis_aruginosa_mycG
Mycobacterium_leprae_pksE
Mycobacterium_tuberculosis_mas
Mycobacterium_tuberculosis_ppsC
Mycobacterium_tuberculosis_ppsD
Mycobacterium_tuberculosis_ppsE
Myxococcus_xanthus_tal_mod2

SNF----RGKKVGVYVMFA----DDYVEMQSKDS----EPHEFLSL---TGHLDLFGSN
T--WESVSGTRTGVFVGNFC-L---DHMMIQSRDW----DNPRPYAF-TGAGTS--ILAN
S--AQELRGSKTGVYIGNFG-L---DQALMALKDS----EFMSPYTS-TGISGT--ILSN
T--LDQVSDANIGCYVGNFV-T---DFITMQLKDS----EYTHRYSA-TGLGTT--ILAN
S--MEA IAGSNTGVYVANFT-V---DYQPLQLRDP----DYLHRYVT-TGSAT--IMSN
T--LSQLSGSNTGCYIGNFT-S---DYLLQHRDH----NNPKPYSL-LGSGYT--IISN
S--METISKSKIGCYCAMFV-S---DYHDMQMOP----EYLPTFIA-IGTRT--MLAN
P--LESLDGAPVGCYVGSYA-SGKSDYHDMQMRDP----EQRVSGHA-VGTGRA--ILSN
T--LEQLSGSNTGVYVQWA-S---DYQEMLLRDI----DFPIYQA-SGVGAA--ISSN
S-----GSDTAVFWGV---NSDDYSKLV--LE----D-LPNVEAWMGIGTAYCGVFN
S-----GSDTAVFWGV---NSDDYSKLL--LE----D-LPNIEAWMGIGTAYCGVFN
S-----GSETAVFMGV---NSDDYSKLL--LE----D-IPNVEAWMGIGTAYCGVFN
S-----GSDAAVFMGV---NSDDYGKLL--LE----D-LPHVEPVMGIGTAYCGVAN
A-----GSDTSVYMGV---NSDDYGKLV--LE----D-LTGVAHMGVGTAYCGIPS
A-----GSDTAVY--I---DPNDYSRML--ME----D-LQAI EAWSIGTAHHGISN
---VPNRPSTRLDRI GTFYGGTSDDWREINAAQ-E-----VDTYYITGGVRAFPGP
---VPNRPSTRLDRI GTFYGGTSDDWREINAAQ-E-----VDTYYITGGVRAFPGP
---VPNRPSTRLDRI GTFYGGTSDDWREINAAQ-E-----VDTYYITGGVRAFPGP
---VPNRPSTRLDRI GTFYGGTSDDWREINAAQ-D-----VDTYYITGGVRAFPG--
---TPNRPSTRLDRI GTFYGGTSDDWREINAAQ-E-----VDTYYITGGVRAFPGP
---SPNRPSTRLDRI GTFYGGTSDDWREVNEAQ-D-----IDTYFITGGVRAFPGP
---VRDRTPSTQAHRI GTFYGGTSDDWREINAAQ-D-----IDTYFITGGVRAFPGP
---VPNRPSTRLDRI GTFYGGTSDDWREINAAE-N-----IDTYFITGGVRAFPGP
---VHGR--GIHQRRVGTFFYGCSSDDYREVNTGQ-D-----IGTYFIPGGCRAFPGP
---SPNRPSTRLDRI GTFYGGTSDDWREINAAQ-N-----IGTYAVPSGERGFANG
---VPDSTPSTRDRVGI FYGMTSDDYREVNSGQ-D-----IDTYFIPGGNRAFTPG
---VPNSPSTRDRVGI FYGMTSDDYREVNSGQ-D-----IDTYFIPGGNRAFTPG
---VPDRPPSTRDRVGI FYVHPSDDWREVNSGQ-N-----VDTYYIPGGNRAFTPG
---VPDATPSTRDRVGI FYGVTSDNWMETNSAQ-N-----IDTYFIPGGNRAFTPG
---VPDATPSTRDRVGI FYGVTSDNWMETNSAQ-N-----IDTYFIPGGNRAFTPG
---VPDRTPSTRDRVGI FYGVTSDNWWEVNSQ-D-----VDTYYIPGANRAFTPG
---VPDRTPSTRDRVGI FYGVTSDNWWEVNSQ-D-----VDTYYIPGANRAFTPG
---VPDTPSTRDRVGI FYHGVTSNDWMETNTAQ-N-----IDTYFITGGNRAFTPG
---VPDTPSTRDRVGI FYHGVTSNDWMETNTAQ-N-----IDTYFITGGNRAFTPG
---SDGATRAVDPARIATFFGQCNDWDHVDVSHHTLG-----CDAYTLQGVQRAFAG
---ARDSSTSTHPSKIGTYFGQTVDDWKDINAQ-MG-----IDTHYLPDLRAFHPG
P-----SSTSETDDYGCYVAVMNSY- NLACHP-----PNTYATIGTGRAFLSG
RE-----NGDNIGCFIGASFNEYLD-NTSAHA-----PTAYTSTGTIRAFLCG
RE-----NNDPIGCFIGASFVEYLD-NTNSNA-----PTAYTSTGTIRAFLCG
---VANRTQSTRLQRI GTFYGGTSDDWREVNSGQ-E-----VGTYYIPGGCRAFPGP
RE-----NGDKVGCFIGASYVEYAD-NTNAYP-----PTAYSAVGTIRAFLCG
KQANADANGPDPKNIIGCYIGMCGNDYVA-NTSHHA-----PNTAYTSTGTIRAFVAG
NAD-----KHIGCFLGVGNVDYEA-NVACYP-----ATAYSATGNLKSFVAG
RQLISGDKGGNEQNDVGCFFGVCSDYND-NIASHP-----PNAFSSLSLRAFVAG
KQEAETHERTQNKDITGCFIGMIAPDYSL-NLASHP-----ASPYTGIMHRSYVAG
-----SLRGTHTGVVWGV---SGSETSEALSQDPE-----TLVGYSMVGCQRAMMAN
----ALRGTDTGVWVGA---SGSEALEALSQDPE-----ELLGYSMTGCQRAMLAN
----DLRGSKTGVFVGC---SASETSGMLTQDPD-----TVTGYTLTGCVSRMFSN
----ELRDTKTGVFVGA---CFSESEKTFWYEMK-----QVNGFGITGCSRAMLAN
R-----GSRTGVYMGV---CS---IPAHSTAL-----C-LQELGQYDATGSAGATMSG
E--DLEATGCMVSGIIGNLDDF-YNTSVTYDKEGYKRV--S-PLFVFKIXNLGAGHIAM
A--DQVATGVAIGMGMIPLEVV-SETALNFQTKGYNKV--S-FFFVFKILVNMAAGQVSI
SKIDKERAGVLVSGMGLTTFV-SDGVQALIEKGRKI--T-FFFIPYAITNMGSALLGI
--LDPARVGVSLGSVAVAATSL-EREYLLSDSGRDWEVDA-AWLSRHMFTYLVPSVMPA
--TDPHRCVSVGSVAIGTMSL-DREYNVLSDGGRKWLDDH-EYAVPHMYNYLVPTIAA
-PQTSVFMASNSY-----R---ALLPSDTTE-SLETPDGYVSWVLAQSGT--IPT
S-----GTGTGIFIGT---GNTGYK-DLFHRA---NLP IEGHAA-TGHMIPSVGPN
AKPKGRNKRQHVGVFAGV---MHKDYT---LVGA---EEASAENVF-PLSLNYAQIAN
GRKRDAELGGSVGVYVGV---MYEYQ---LYGA---QEQRVGRSL-ALTGPNSSIAN
NEKK-----CGVYLG I---MSNEYG-VMLNR-----QSRAN-ATGNSFAIAAA
S-----GSQLGVYVGY---SKGVYDYERLLSA---NYPEELHHY-IVGNLPSVLAS
A-----DNKVMGVFVGI---TSIDHALKVYGTN----Y-D-QIDFFGSGNALSAAAG
QGSIGIFAGASMTYLINNC-YPNRGLKDSNDELQPFTLDSMGQYTMVANDKDY--LTT
S-----GTRTAVMMGL---SSWDY--TIVNIE---R-RADIDAYLSTGTPHCAAVG
A-----GSS TAVFTGL---THEDY--LVLTIT---A-GGLASPYVVTGLNNSVASG
R-----GTQTSVFGV---TAYDYMLTLAGRL---R-PVDL DAYIPTGNSANFAAG
S-----GTRTGVMMLG---SSWDY--TIVNIE---R-RADIDAYLSTGTPHCAAVG
DGSIGVYGTSSPSGYLLHNL-LSHRDPNAVLAEGLNF-----DQFSLFLQNDKDF--LAT
-GPCGVMTASNSFY-----HQGSPQFPADGQP-VLRTAEYVWVLAQAGS--IPT

Nostoc_sp_nosB
Pseudomonas_syringae_cfa6
Pseudomonas_syringae_cfa7
Sorangium_cellulosum_epoA_mod1
Sorangium_cellulosum_epoC_mod2
Sorangium_cellulosum_epoF_mod9
Stigmatella_aurantiaca_mxaB2_B1
Stigmatella_aurantiaca_mxaC1
Stigmatella_aurantiaca_mxaC2
Stigmatella_aurantiaca_mxaC3
Stigmatella_aurantiaca_mxaD
Stigmatella_aurantiaca_mxaE
Stigmatella_aurantiaca_mxaF
Streptomyces_natalensis_PimS0
Streptomyces_natalensis_PimS2_mod3
Streptomyces_natalensis_PimS2_mod4
S_viridochromogenes_aviM
Xanthomonas_albilineans_xabB_mod2
Xanthomonas_albilineans_xabB_mod3

NGLIGVYAGVGMNRYLVNLL-YPHHQLLET-----VDPLQLTISNDKDF--LPT
H-----GSLTGVYAGI----MGTEYGTQIQ--H----A-SEDVVTGYGYMGATATCVAAG
K-----NSATGVFLGA-----NQNGYLADLQ--R-----R-NPAADGYRLQGGSSIIISG
V-----GTEGTGFFIGI----GPSEYEALPRAT----A-SAEIDAHGGLGTMPMSVVGAG
E-----GSIQVYAGAN----MSSYLTSNLHEHP----A-MMRWPGWFQTLIGNDKDYL
D-----GSRTGVFLGA----CSSDYSHVAQQR----R-EEQ-DAYDITGNTLSVAAG
A-----GSRKGVFMGV----MSSDYMARLLKEN----D-ATFRDGYMATGNQYSPVPG
S-----GSRSGVFIGA----CNDDY-HCMQVER----P-ETG-DAFSATGVAASVLSG
A-----GTRTGVFVGV----CGYDY-AMLQAER----D-VEG-DVYSVIGCSNSVIAG
A-----NTRTGVFFIGI----GSNEYAMLNGVGS----A-SAAGDAYIATGNDSSFAAG
A-----GSRTGVFLGI----IGSDYAQLQARLL----G-DSP-DIYHLTGTSLNAAAG
A-----GSTGTGFFIGI----YSDDY-ALLQVGN----P-SAR-DSSSVTGALNCVVPV
Q-----GSPAVCFGV----VWTDYEAMLQRMG----L-R-RISSYTSSSGSHSIVAN
R-----GSRTGVFTGA----MDRQYGTSA--A----A-PSAWESMLITGTAGSAVSG
H-----GTTTGTFIGS----TYQEY--GLG--V----E--DGSAGHLVGTGTPSVLSG
R-----GSTGTGFFIGI----SYQDY-RAFG--A----G-PDGAEGHLITGTISSVLSG
A-----GTDGTVFAGV----CTYDYG--GRLE----D-LPNIDAWTGIGAAVCAVSN
G-----GQPVGLFVGA----HTSDYGELLASQP----QLMAQCGAYIDSGSHLTMIPN
SKS-----QRVGVFVGV----MNGYY-----SSGARFWQIAN

[
Bf_PKS7
Aspergillus_terreus_at1
Aspergillus_terreus_at4
Aspergillus_terreus_at5
Gm_PKS5
Ch_PKS9
Ch_PKS8
Gz_PKS2
Gm_PKS2
Ch_PKS10
Ch_PKS5
Ch_PKS7
Ch_PKS1
Didymella_maydis_PKS1
Gz_PKS4
Gz_PKS7
Gm_PKS7
Ch_PKS6
Gz_PKS1
Ch_PKS4
Gm_PKS6
Nc_PKS1
Bf_PKS8
Nc_PKS2
Ch_PKS2
Aspergillus_terreus_lovF
Penicillium_citrinum_mlcB
Ch_PKS3
Nc_PKS3
Bf_PKS2
Gz_PKS5
Gm_PKS1_CAC44633
Gm_PKS1
Bf_PKS5
Gm_PKS9
Aspergillus_terreus_lovB
Penicillium_citrinum_mlcA
Gz_PKS9
Ch_PKS16
Bf_PKS6
Bf_PKS3
Ch_PKS17
Gz_PKS10
Gm_PKS10
Nc_PKS4
Bf_PKS4
Gz_PKS11
Ch_PKS14
Ch_PKS12
Gm_PKS12

181
.....
RINYFFKFSGPSFNCDTACSSGLASIQMACTSLHGDADTUVVAGGLNVLCDSDG-FTGL
RISYFFRFSGPSLSDTACSSSFAAIQAACSYLWRGEC-DTAAIAGGNILNPNPDN-FAGL
RINYHFKFKGPSYNVDTACSSSFAAIQLACTSLLAKEC-DTALAGGLNVMTTTPDL-FAGL
RLSYFYDLHGPSISLDTACSSSLNACHLACNSLRLGEC-DMALAAGYNLFYNPD-TIIPL
RLSWFYDFSGPSMTVDTACSSGLVALHLACQELLAGSV-NMSLVCGTNLCLLPDSTAL-L
RISHFFDLHGPSLTVDTACSSSLVALHLGQVQSLTGES-TMSLVCGANTFITPESQALAL
RVSWYFDLRGPSINVNACSSSLTAVDMACKALNSGDAYSQAVVTGTVNLLDPSI-FQVL
RVSWYFDLRGPSIHVNTACSSSLSAVDMACKALKSGDA-SSAVVTGANLLLDPSI-FQVL
RVAVFFDFRGPFAFTVDTACSSSMVALHQAVIGLKSRECNLVALACGTNVLISPEFG-QQL
RVSWFFDLKGPSVTIDTACSSSMAALHLGQSRLTGES-DMSIITGVITLNYPGD-VNGM
RLSYFFDLHGPSVTVDTACSSGLVALHLACQSLRAGEA-KSAIAAGVNVVNLNHEF-MTMM
RISYSFDLKGPSVLDVDTACSSGLTALHLACQSLLVGDV-RQALAAGSSLIILGPEM-MVTM
RVSYSFDLKGPSITVDTACSSGLTALHLACQSLVTGES-TQAVVSGSSLIILGPEM-MVTM
RISHFFDLHGPSATVHTACSSSLVALHLACQSLLSGDA-EMALAGGVMILTPDG-TMQL
RISYFFNLHGPSMTIDTGCSSGLSVVHLAAQSLRTKET-SLAIAAGAGMILTPTNT-MMPM
RISHFFNLHGPSMTIDTGCSSGLSVVHLASQSLRASET-SLAIAAGAGMILTPTNT-MMPM
RLSWFYDLRGPSLTVDTACSSSLTAFHLACQSLRGTGES-EMSVVGGANLMFGPDMISIL-L
RVSWFFDLHGPSLTVDTACSSSLVALHLACQSLINDGVS-KVAVAAGANLMLNPDN-PMWM
RISYFFDLRGPSMTIDTACSSSLVALIHEACALRMGEI-DQALIGGVNLIILDPD-KLMVQ
RISWFFNLKGPSVTIDTACSSSLTALHLAQSLIRNGES-DSALVGGGLHLLNPFVGF-M
RISHFFDLRGPSFTADTACSSSMVAIHLACQSLRGTGES-SAALVGGCHLNMLPEF-WISF
RISHFFDLRGPSFTMDTACSSSLVALHLACQSLIRNGEC-KSAITGGCHLNMLPEF-WISM
RLAYFFNLHGPTFTVDTACSSSLHALHLAVQSLIRNGEC-DAAVVGASHLITQPDV-WVSM
IVSNTFDLKGPSVSDTACSSAFYALQLASQSLRSGET-EMCIVSGCALNISF-WRWTML
RVSHFYDLRGPSVSDTACSSSTLTLALHLAIQSLRAGES-DMAIVAGANLMLNPDV-FTTM
RISHFFDLRGPSVTVDTACSSSTLTLALHLACQSLRGTGES-DTAAIVAGANLMLNPDV-FVTM
RLSHYFDLRGASMSIDTGCSTLTLALHLAQACNDLRNGES-DMSVVGANLMLNPDN-FITM
RLSHFFDLRGPSMSVDTGCSTLTLALHLAQACQSLRSGES-TMSIVGGANLMLNPDN-FLAM
RISYTFDLKGPSMTVDTGCSSGLVALHLAQACQSLRSGDT-DMALVGGTSLILSPDT-MVPM
RVSYCFDLKGPSMTIDTGCSSGLVGLHQACQSLRAGES-ELALVAGGSLVILQPDV-LYTV
RISYFFDWHGPSMTIDTACSSSLVAVHQAIQTLRSGES-EVAIAAGANLILTPGM-YIAE
RISYFFDWHGPSMTIDTACSSSLVAVHQAIQTLRSGES-QV.....
RLSYFFDWHGPSMTVDTACSSSLVAVHQVTSRLNGEC-PVVAAGVNLILGPGM-WIAE
RISYFFDWHGPSMTIDTACSSSLVAVHVAVRTLRSGES-NVAVAAGANLILGPGM-YICE
RISYFFDWHGPSMTIDTACSSSLVAVHLAVQQLRTGQS-SMAIAAGANLILGPM-TFVLE
RVSYFFDWHGPSMTIDTACSSSLAVHLAVQQLRTGQS-TMAVAAGANLILGPM-TFVME
RVSYFFDWHGPSMTIDTACSSSLVAVHVAQALRNQTS-RMAVACGTNLIILSP-LFISE
RVSYFFDWHGPSMTIDTACSSSLVAVHVAQALRSGDS-HAAVACGSNLIIFGPEM-YVIE
RISYFFDWHGPSMTIDTACSSSLVAVHNAISTLRSGQS-RTAIAAGANLILGPEM-YIGE
RVSYFFDWHGPSLTVDTACSSSLVAVHQAVSALRNGEC-RSAVAAGVNLIFGPEM-FNGE
RISYFYDWHGPSMTIDTACSSSLVAVHVAQALRSGIS-RVACAAGSNAILGPEM-FVIE
RISYFLDWHGPSMTIDTACSSSLVAVYDAVTLRNGVS-RIACAGGVNLIILGPEM-MISE
RISYFLDWHGPSMTIDTACSSSLVAVYDAVTLRNGVS-KIACAGGVNLIILGPEM-MISE
RVSYFFNFHGPSMTIDTACSSSLVALHLQAVLSLRSGEA-EMACVSGVNLILTPEQ-FVVE
RISYFFDWHGPSVTVDTACSSSLVALDHAVQQLRSGSS-TLAVAAGANLMLDGRP-FIGF
RVSYELDLGM.....
RISYFGLTGPSMTIRTACSSALTGLAEAFAAIQRGIC-EGAIVAGSNLILAPGMT-QQM
RLSYEYDLRGPSMTIRTACSSALMGLHEACLAIRNGDC-SAALVGGSNLFWSPDTM-ADM
RISYEYDFQGPSMVIKTCSSSAVALHEACRAIQRGDA-SSAIVGGANMIMTALP-ATM

Mycobacterium_tuberculosis_ppsC
Mycobacterium_tuberculosis_ppsD
Mycobacterium_tuberculosis_ppsE
Myxococcus_xanthus_tal_mod2
Nostoc_sp_nosB
Pseudomonas_syringae_cfa6
Pseudomonas_syringae_cfa7
Sorangium_cellulosum_epoA_mod1
Sorangium_cellulosum_epoC_mod2
Sorangium_cellulosum_epoF_mod9
Stigmatella_aurantiaca_mxaB2_B1
Stigmatella_aurantiaca_mxaC1
Stigmatella_aurantiaca_mxaC2
Stigmatella_aurantiaca_mxaC3
Stigmatella_aurantiaca_mxaD
Stigmatella_aurantiaca_mxAE
Stigmatella_aurantiaca_mxAf
Streptomyces_natalensis_PimS0
Streptomyces_natalensis_PimS2_mod3
Streptomyces_natalensis_PimS2_mod4
S_viridochromogenes_aviM
Xanthomonas_albilineans_xabB_mod2
Xanthomonas_albilineans_xabB_mod3

RLAYILGARGPAVVIDTACSSSLVAVHLACQSLRGRES-DMALVGGTNNLLSPGP-SIAC
RIAYLLGLRGPVAVDTACSSSLVAIHLACQSLRLRET-DVALAGGVQLTLPFFT-AIAL
RISHAFNLGRPSIAVQHTACSSSLVAVHLACLSSLGEC-DMALAGGSLCIPHRVYFTS
MVSYKLGKGLPSLFVHTNCSSSLVAVVQAQIAAGDC-QTALVGAATVFPANLGYLHQ
RVAYKLNLTGTAVNVQACSTSLVAVHLACQSLLNyec-DMALAGGVTLSPQKIGYLHQ
RVSYCLGLQGPALAITACSSSLVAIHTACEALRSNDC-QLALAGGVTVMPPTPGV-LIDF
RIAFVLGLRGPVAVDTACASASLTDIHLAVQSLRRHEC-SLALAGGVTDMDTPEV-FAEF
RISYVLGLRGPVAVDTACSSSLVAVHLACQSLRSGEC-STALAGGVSLMLSPST-LVWL
HVSYRLNLRGPSISVQACSTSLVAVHLACMSLLDREC-DMALAGGITVRIPIHRAGYVYA
RLSYTLGLQGPCLTVDTACSSSLVAIHLACRSLRARES-DLALAGGVNMLLSPET-MIML
RVSYVLGLQGPCMPVDTACSSSLVALHLASESLRAGES-NLALAGGVNLIILSPET-MICL
RLSYTLFNLQGPVAVDTACSSSLVSLHLACQSLRNREC-NMALAGGVNLIILSPQS-VLLV
RLSYLMDLRGPAMTVDTACSSSLVALHLASQSLRNREC-DAALVGGVNNLLSQRP-SSWL
RLAYLLRFQGPMTSLNTACSSSLVAVHLACQSLRAGES-NLALAGGVNMTLAPHS-TIYL
RLSYTLGLQGPCMSIDTACSSSLVALHVACQSLRNREC-DLALSAGVNMLMPDA-TIAL
RLSYLLDFQGPCLAVDTACSSSLVALHLACQSLRNQEC-SMALAGGVNLIILSPQS-SSRV
RVSYVLGLRGPVAVDTACSSSLVAVHLACESLRRGES-TMALVGGVNLTIPADS-TVGL
RIAYTYGLEGPALTVDTACSSSLVALHLACRSLRSGE-TALAGGVTVMTATPAP-FAHF
RLAYLFGLEGPVAVDTACSSSLVALHLACQSLRNRES-SLALAGGATVMTNPNP-FVAF
RLSYTYFNLQGPVAVDTACSSSLVALHLACQSLRNRES-SLALAGGVNLIILSPQS-FVGF
RVSHALDLRGPVAVDTACSSSLVALHHTAAQSLRLGEC-TVALAGGVNLLVSPGQ-TIAL
RASRWNFNTPGSEVINACSSSLVALHRAVQSLRQGES-SVALVGLVNLIILAPKV-LLAS
RVSYQDFRGPVAVDTACASASLTAIHLALESLSRSGSC-EVALAGGVNLLVDPGQ-YLNL

[
Bf_PKS7
Aspergillus_terreus_at1
Aspergillus_terreus_at4
Aspergillus_terreus_at5
Gm_PKS5
Ch_PKS9
Ch_PKS8
Gz_PKS2
Gm_PKS2
Ch_PKS10
Ch_PKS5
Ch_PKS7
Ch_PKS1
Didymella_maydis_PKS1
Gz_PKS4
Gz_PKS7
Gm_PKS7
Ch_PKS6
Gz_PKS1
Ch_PKS4
Gm_PKS6
Nc_PKS1
Bf_PKS8
Nc_PKS2
Ch_PKS2
Aspergillus_terreus_lovF
Penicillium_citrinum_mlcB
Ch_PKS3
Nc_PKS3
Bf_PKS2
Gz_PKS5
Gm_PKS1_CAC44633
Gm_PKS1
Bf_PKS5
Gm_PKS9
Aspergillus_terreus_lovB
Penicillium_citrinum_mlcA
Gz_PKS9
Ch_PKS16
Bf_PKS6
Bf_PKS3
Ch_PKS17
Gz_PKS10
Gm_PKS10
Nc_PKS4
Bf_PKS4

241
.....
SRGHFLSKT-GGCKTFD-----CNADGYCRGDGVSVMKRLDDAQRD--NDNLI
DRGHFLSTT-GNCNTFD-----DEASGYCRSDAVGSVILKRLDADAEADN--DP
SRAHFLSKT-GSCKTFD-----DGADGFCRGDVGTVLKRLEDADAEADN--DP
TALGFLSPD-GRCYSFDERA-----NR---YSRGEGFGMVLKRLSDAIRDG--DC
SSLNMMSKD-SVCYSFDERA-----S--GYARGEFGVVLKRLSEAIADG--NNIR
SNGGFLSVD-GKYSFDKA-----N--GYARGEFGFVLLKPLDAAIRDG--D
ANQGFLSPD-GVCYSFDERA-----N--GYARGEVIAVVLKPVQAAIENG--DM
ANQRFLSPD-GVCYSFDERA-----N--GYARGEVIAVVLKPVQAAIENG--DM
NGLGVLSFQ-GASKSFD-----KEANGYGRGEGISVVVLRKMSDAIRDG--DT
SHQGFLSPD-GRCYSFDHRA-----N--GFARGEAGTVLKRLEDALRNG--DT
SMMKFLSPD-GRCYAFDERG-----N--GYGRGEGVGTVFLKPLADALDGD--DP
SMMKFLSPD-GRCYAFDERA-----N--GYARGEVAVLKLKRLDADALDN--DT
SMMRFLSPD-GRCYAFDDRA-----N--GYARGEVTVLKLKRLDADALANG--DT
NNLGFNLPE-GHSRFDKDA-----G--GYGRGEGCGVILKRLDQAIQD--DN
TALNFLSPD-GKCFTFDSRA-----N--GYGRGEGIGVVVMKRLSDALRDN--DT
TALNFLSPD-GKCFTFDSRA-----N--GYGRGEGIGVVVMKRLSDAIRDN--DT
GAAKILSPE-GSKMWDANA-----D--GFARGEFGVTVLKPLDALTALRDG--DT
SNMSFLSAD-GLSKSFDASA-----D--GYGRGEGIAAVLKLKPLDQAVRDR--DP
SSMQFLSPD-GRCYSFDARA-----S--GYSRGEGVAGIMLKLPSKALKDR--DT
SSMSFLSAD-NKCHSFDASA-----N--GYARAEAGGFFVLLKRLDQALADG--DT
STCRLLSDA-GRSFSFDNRG-----T--GFGRGEGCGMIVLKLPLDQAIKDK--DP
NKYRFLSDE-GRSFSFDSRG-----T--GYGRGEGCGLIVLKLKPLDQALRNG--DN
SMLRFLSDE-GRTYAFDHRA-----RS--GYARGECAVILKPVKAFKDN--DH
SNLTMLNPD-GLSKSFDPA-----DA--GYVRGEGAASIVKPLDAAIRDN--DR
SNLGFLLSSD-GISYSFDRA-----D--GYGRGEGVAAIVLKLTPDAVRDG--DP
SNLGFLLSPD-GISYSFDPA-----N--GYGRGEGIAAVLKLKPLDQALRNG--DP
SSIALISKD-GRSFAFDSRA-----N--GYGRGEGAATVILKRLDADAIRDG--DP
SSMTLISKD-GRSWAFDSRA-----N--GYGRGEGSATVILKPLDAAALRDG--DP
DKRSVLDPS-GKSFVFDNRG-----V--GYGRGEGVATVILKRLDQALNAG--DH
SGMGMLNPD-GKSFVFDNRG-----A--GYGRGEGVATVILKRLDQAIKDN--DR
SKLSMLSPS-GSKMWDQDV-----N--GYARGEVIAAVLKLKPLSAAIRDN--DH
.....GIAAVLKLKPLSAAIRDN--DH
SKLHMLSPT-GSKMWDASA-----D--GYARGEVIAAVLKLKPLDQALRNG--DP
SKLRMISPT-GSRMWDANA-----D--GYARGEVIAAVLKLKPLDQALRNG--DP
SKLSMLSPS-GSRMWDAGA-----D--GYARGEAVCSVILKTLKPLDQALRNG--DP
SKLNMLSPN-GSRMWDAAA-----D--GYARGEVCSIVLKLKPLDQALRNG--DS
SNLSMLSPD-GSRMWDADA-----D--GYARGEVIAAVLKLKPLSAAIEDN--DV
SKLMLSPD-SLSRMWDKDA-----N--GYARGEVTAIVLKLKPLDQALRNG--NR
SNLHMLSPD-SLSRMWDSRA-----D--GYARGEVTAIVLKLKPLDQALRNG--DI
ANLKMISPT-GTKMWDASA-----D--GYARGEVCS--MLKLEDAIRDG--DR
SKLQLLSPT-GSRMWDAAA-----D--GYARGEVIAAVLKLKPLSAAIALDNG--DE
SKLHMLSPT-GSRMWDASA-----N--GYARGEVIAAVLKLKPLDQALRNG--DH
SKLHMLSPD-GSRMWDASA-----N--GYARGEVIAAVLKLKPLDQALRNG--DH
SSLHMLSPT-GRCHMWDRA-----D--GYARGEVIAAVLKLKPLSAAIALDNG--DR
NNMGMLSPD-GSKMWDTEA-----D--GYARGEVIAAVLKLKPLDQALRNG--DN

Microcystis_aruginosa_mycE
Microcystis_aruginosa_mycG
Mycobacterium_leprae_pk5E
Mycobacterium_tuberculosis_mas
Mycobacterium_tuberculosis_ppsC
Mycobacterium_tuberculosis_ppsD
Mycobacterium_tuberculosis_ppsE
Myxococcus_xanthus_tal_mod2
Nostoc_sp_nosB
Pseudomonas_syringae_cfa6
Pseudomonas_syringae_cfa7
Sorangium_cellulosum_epoA_mod1
Sorangium_cellulosum_epoC_mod2
Sorangium_cellulosum_epoF_mod9
Stigmatella_aurantiaca_mxaB2_B1
Stigmatella_aurantiaca_mxaC1
Stigmatella_aurantiaca_mxaC2
Stigmatella_aurantiaca_mxaC3
Stigmatella_aurantiaca_mxaD
Stigmatella_aurantiaca_mxaE
Stigmatella_aurantiaca_mxaF
Streptomyces_natalensis_PimS0
Streptomyces_natalensis_PimS2_mod3
Streptomyces_natalensis_PimS2_mod4
S_viridochromogenes_aviM
Xanthomonas_albilineans_xabB_mod2
Xanthomonas_albilineans_xabB_mod3

SQSGMMSPD-GRCKTFDASA-----NGYVRGEGCGVLLIKTLSEAQKNG--DHILA
EGL-IMSPD-GHCRAFDAAE-----KGTIFGSGVGVIMLKRSLDALADH--DHIYA
SKWSALSPT-GRCHSFDANA-----DGFVVRGEGCGVVVLRKLDALRDQ--DRVLA
SAQGMLSST-GRCHSFDAD-----ADGFVRSEGCAMVLLKRLPDALRDG--NRIFA
SRWGMLSPD-GRCKTFDASA-----DGVVRGEGAAVVVLRKLDLDAVRDQ--NRILA
SKWSALSPT-GRCHSFDANA-----DGFVVRGEGCGVVVLRKLDLDAVRDQ--DRVLA
PGS-MVSAV-GHCRPFDVRA-----DGTVFGSGVGLVVLKPLAAAI DAG--DRIHA
RGL-NFSSA-GRVKAFDAAA-----DGMIA GEGVAVLVVKDAAA AAVRDG--DPIYC
EGM-ILSPD-GHCRAFDACA-----QGTIASSGAGIVVLRKLDKDAIADR--DHIHA
SQQRVLAPD-GRCKAF-----SASADGVGLGEGVGMVLRERLSQAQAKG--RLILG
TRQNGLAAD-GYCKAF-----AEQADGTCFAEGAGVLLERLADAQRAG--HPVLA
SKTRALATD-GRCKAFSAEA-----DGFGRGEGCAVVVLRKLSGARADG--DRILA
EGG-IFSPD-GHCRAFDACA-----NGTIMGNGCGVVLKPLDRALSDG--DPVRA
GRIQALSPD-GHCRTFDASA-----NGFVRGEGCGMVVLRKLSDAQRHG--DRIWA
CSMQALASD-GRCKTFDASA-----DGYVRGEGCGVVLKRLSDAQRDG--DSILA
SKLQALSPD-GRVKAFDAAA-----NGFTRGEGCGIVVLRKLSDALVIAAG--DHIHA
SKLMALSPT-GRCHSFDASA-----NGFVRGEGCGVVVLRKLSDALASG--DNVVG
AQLRALSPD-GRCKTFDASA-----DGYVRSEGCIVVLRKLSDAKRDG--DDVLA
SSSRGLSPD-GRCKTFDAAA-----NGFVRAEGCVVLRKLSDALASG--DEILS
SRAQALSPD-GRVKAFDAAA-----NGFVVRGEGCGVVVLRKLSGARADG--DPIYC
SKLALSPD-GRCYTFDARA-----NGYVRGEGAGVAVLKLPLSRAIADG--DPIYC
SRLRALSPD-SRSMAY-----ADAANGSAWSEAGLLELRLERLSDARRNG--HRVLA
SRQALAGD-GRCKAF-----SEGADGMTLAEGVGLVLRERLSDAQRNG--HEILA
SRQRAMAAD-GRCKAF-----SDQADGMSLAEGVGLVLRERLSDARRNG--HQVLA
GTAGALAPD-GRSKPFAASAGRYAVAASADGYGRGEGCGVLLIKLLTDAVRDQ--DRVLA
ASAGMLSPD-GRCKTLDAAA-----DGFVVRSEGIAGVILKPLAQALADG--DRVYG
AGAAMLSAG-ASCRPFG-----EAADGFVAGEACGVVLLKPLKQARADG--DVIHA

[
Bf_PKS7
Aspergillus_terreus_at1
Aspergillus_terreus_at4
Aspergillus_terreus_at5
Gm_PKS5
Ch_PKS9
Ch_PKS8
Gz_PKS2
Gm_PKS2
Ch_PKS10
Ch_PKS5
Ch_PKS7
Ch_PKS1
Didymella_maydis_PKS1
Gz_PKS4
Gz_PKS7
Gm_PKS7
Ch_PKS6
Gz_PKS1
Ch_PKS4
Gm_PKS6
Nc_PKS1
Bf_PKS8
Nc_PKS2
Ch_PKS2
Aspergillus_terreus_lovF
Penicillium_citrinum_mlcB
Ch_PKS3
Nc_PKS3
Bf_PKS2
Gz_PKS5
Gm_PKS1_CAC44633
Gm_PKS1
Bf_PKS5
Gm_PKS9
Aspergillus_terreus_lovB
Penicillium_citrinum_mlcA
Gz_PKS9
Ch_PKS16
Bf_PKS6
Bf_PKS3
Ch_PKS17

301
.....
VILASATNHSANA-ISITHPHAQTQADLFRHILTQSGVS---PLEVDYVEMHGTGTQAG
VIVGTNTNHCQGT-DSITRPHGPDQVSVFKRIVRHSGID---PLDISYVEMHGTGTQAG
VVLGTATNHSSEA-VSITRPHGPAQEALYRKILKHTGVD---PVDVSYVEMHGTGTQAG
IVRGSSSNQDGNM-PGITQPTFQAQVDLINAAYRSAGLSKQT---RFFEAHGTGTQAG
VIRSTGCGQDQNT-PSITSPSQAQERLIRETYARAGLSLDETR---YFEAHGTGTQAG
VVRATGNQDQGT-PSITQPSQAQQLDRLRDTYRVLNVDAT---YVEAHGTGTQAG
VIRSIGSNQDQHT-PILTQPSQSQEELIRHVYKQAGLSMSDT---RYVEAHGTGTQAG
VIRSIGSNQDQHT-PILTQPSQSQEELIRHVYKQAGLSMSDT---RYVEAHGTGTQAG
VIRNSGIGHDKGK-APLPAPVRESQVNLVRRCYAQAKIDPSET---RLFEAHGTGTQAG
VVRGTGNQDQGT-PGLTLPSSDAQERLIRTYASAGLNFDDTT---YVEAHGTGTQAG
VIRGSASNQDQGT-SGITLPPNVAQEALIKDVYHVADLNPIDTS---IVEAHGTGTQAG
VIRGTGNQDQGT-PGITMPNSVQSQEALIRSVYKKAALDPLDPT---YVECHGTGTQAG
VIRGTGRNPDQGT-TGIAMPSSGLAQEALIRSVYAKTGLDLDLDTA---YIECHGTGTQAG
VIRASGVNSDQGT-QGVTMPSSQAQALIKHVYETRGLDYGTG---YVEAHGTGTQAG
VIRATKVNQDALK--GITLPSKEAQVANIHSVYESAGLDFSQTG---YVECHGTGTQAG
VIRATKVNQDQGT-TGITLPSKEAQVANIHSVYESAGLDFSQTG---YVECHGTGTQAG
VVLATATNEDGHT-PGISLPNSEAQQLIRRAYQMAGVDPAAETG---YVEAHGTGTQAG
VI.....
ILRGTSVVSDGKT-LGITMPMSMSQVEAITRAYEQAGLVLDLDTA---YIEAHGTGTQAG
VLRSTGNSQDQGT-LGITQPSASRQEELIRATYASAGLTFDKTN---FFEAHGTGTQAG
VIAATGLNQDQGT-PGITVPHGPAQEDLIRKVVYGRAGLDPNLCG---FVEAHGTGTQAG
VIRSGINQDQGT-PGITMPNGSAQESLMKWVYESAGIDPKDTG---YVEAHGTGTQAG
VISHSGVSHNGRT-VGIVAPSPDEQEQLLRDVFTAAKIDPREVG---FFEAHGTGTQAG
VLSDIGVNHNGRTN-GYTLDPARMQASLMRELQVRLDIKPDQEF---FVEAHGTGTQAG
IVRETAI NQDQGT-PAISTPSGEAQECLIQDCYQKQALDPKQTS---YVEAHGTGTQAG
VIRETALNQDQGT-PGITMPDVAQKSLIQECYDKAGLDMSLTS---YVEAHGTGTQAG
VIKETGLNQDQGT-ETITTPSQAQIDLMRRLYKKAAGLDPKDTG---YFEAHGTGTQAG
VIRDSGINQDQGT-ETITTPSQAQEALIRACYERAGLDPGQTT---YFEAHGTGTQAG
VIRNTGINQDQGT-SGIALPSQVAQALANHVFKQVGLDPCKIN---YVEAHGTGTQAG
VIANSQMNQDQGT-PGLNTPSQAQALSRVYQEAAGLNPADTS---FVEAHGTGTQAG
IIRATGVNQDQGT-PGLTMPNKAQADLIRSTYARAGLDINKPEDRQFFHAGHTGTQAG
IIRATGVNQDQGT-PGLTMPNKAQADLIRSTYARAGLDINKPEDRQFFHAGHTGTQAG
VIRGTGNQDQGT-PGLTMPNKAQADLIRSTYARAGLDINKPEDRQFFHAGHTGTQAG
IIRATGVNQDQGT-PGLTVPNSIAQTALIRDVYQVQAGLDITKPSDRPQLFHAHGTGTQAG
VIRETGNQDQGT-PGLTVPNSIAQTALIRDVYQVQAGLDITKPSDRPQLFHAHGTGTQAG
VIRETGINQDQGT-TGITMPNHSQAQALIRATYAKAGLDITNPQERCQFFEAHGTGTQAG
IIREVGNQDQGT-RGITMPNHSQAQASLIRQTYAKAGLDPATPEGRQFFEAHGTGTQAG
VIRETGMNSDQGT-PGITMPSASAQRDLIRSVYRKAGLDPESAEDRPQYIEAHGTGTQAG
IIRETGVNSDGKS-KGITMPLAASQADLIRQTYARAGLDCTKPAEDRCQFFEAHGTGTQAG
VIRETGVNSDQGT-MGITMPSASSQEKLIKETYRRAGLD-----RCQFFEAHGTGTQAG
IIRETGVNSDQGT-NGITMPSAQAQALIRKTYARAGLDPLK---DGCQFFEAHGTGTQAG

Gz_PKS10
Gm_PKS10
Nc_PKS4
Bf_PKS4
Gz_PKS11
Ch_PKS14
Ch_PKS12
Gm_PKS12
Ch_PKS13_pseudogene
Ch_PKS11_FUM1
Gm_FUM1_AAD43562
Gm_FUM1
Ch_PKS15
Nc_PKS5
Gm_PKS15
Nc_PKS6
Gm_PKS14
Bf_PKS9
Gm_PKS13
Bf_PKS10
Bf_PKS11
Penicillium_patulum_6MSAS
Byssoschlamys_nivea_6MSAS
Aspergillus_terreus_pksM
Aspergillus_parasiticus_pksL2
Penicillium_griseofulvum_pks2
Ch_PKS25
Nc_PKS7
Colletotrichum_lagenarium_pks1
Nodulisporium_sp_PKS1
Xylaria_sp_PKS12
Bf_PKS12
Glarea_lozoyensis_PKS1
Monascus_purpureus_PKS1
Bf_PKS13
Ch_PKS18
Bf_PKS14
Ch_PKS20
Aspergillus_nidulans_wA
Aspergillus_fumigatus_alb1
Gz_PKS12
Gm_PKS3_CAC88775
Gm_PKS3
Gz_PKS12
Gm_PKS4_CAC92399
Gm_PKS4
Aspergillus_parasiticus_pksL1
Aspergillus_nidulans_pksST
Gz_PKS13
Bf_PKS15
Bf_PKS19
Ch_PKS22
Bf_PKS16
Ch_PKS19
Ch_PKS23
Ch_PKS21
Bf_PKS17
Bf_PKS18
Bf_PKS20
Homo_sapiens_FAS
Gallus_gallus_FAS
Caenorhabditis_elegans_FAS
Bombyx_mori_FAS_p270
Ch_9497_PKS24_3p_of_nps7
Nc_3_oxoacyl_ACP_synthetase
Homo_sapiens_3_oxoacyl_ACP_synth
Glycine_max_beta_ketoacyl_synth
Streptomyces_coelicolor_act
Streptomyces_avermitilis_pks9
Bacillus_subtilis_pksK_mod2
Bacillus_subtilis_pksK_mod3

IIREIGVNSDGR-T-NGITLPSDAQKVLIRHTYRNAGLDVFK--DRQCFEAEHGTGTGAG
VIREIGVNSDGR-T-NGITLPSPEAQKFLIRQTYKKAGLDVFK--DRQCFEAEHGTGTGAG
IIRETGVNSDGRS-KGITMPNWEAQSAIQDITYRRSGLNARDPIDRCQFFAEHGTGTAAG
VIRETGVNHNGRT-SGITLPSASAQTSILIRDVYHRAGLDPTNPSDRQYIEAHGTGTQAG
.....
VIRAAVANS DGR-T-QGITQPNGYAHEAMIRLAYKQAGITV-SKYPQTAYFECHGTGTSVG
IIRGSSLNADGKT-PGMSMPSSSSQAALIRRAYQNAGIPE-SEIIRTAFAVECHGTGTATG
IVKGTAVNCDGKC-VSLVTPNGAAHEALMRKAYCDNGLDP-KD---TAFVECHGTGTPTG
VIWASATNYDGRK-IRMLNLTLVQEALICKTYAKASIT---NYRETFGFFKCYGIGMLVG
VVRATSSNSDGR-T-PGMSMPSSSESHMTLIQHAYHEAGLNP-KD---TVFVEAHGTGTVPV
LVRATSSNSDGR-T-PGMSMPSSSESHALIRRAYGEVFLDP-KD---TCFVEAHGTGTGTSVG
LVRATSSNSDGR-T-PGMSMPSSSESHALIRRAYGEVFLDP-KD---TCFVEAHGTGTGTSVG
IIRSTACNDDGKT-PGITQPNTVAHEALIRRAYRTAGISE-EDLNKTFGFFCHGTGTAVG
MIRGTAINSNGRT-GGITRPSANGQETVIREAYRNAGLPPFRDTS---YFVECHGTGTYPV
VIRGTAIGANGRG-DGMTKPDSEGGQARTIRMAYKNAGITDLAQGTG---YFVECHGTGTVPV
VIRGTAINSNGRT-NGITLPSADGQEAIVIRKAYAQAG-LGFNETD---YFVECHGTGTAVG
VIRGSALNANGRT-PGISLPDSDFQELAMRKAYSAN-LDVSQTD---YVECHGTGTAVG
VIRGTAATNANGRT-SGITQPSAKGHETVMRTAYEFAG-LDPRDTS---YFETHGTGTQV
VIRGTSNSSSGAT-PAITEPSGRAQADTILQAYAQAGINDFSETG---YFVECHGTGTVPV
VIRGSATNSDGR-T-PGMSMPSSSESHALIRRAYANAGITNLDNTA---YFVECHGTGT---G
VVQNSVANQDGR-T-NGITVPSAEQAIAIEKAYSQVGLVPYAD----YVEAHGTGTAVG
VIKGSAVCQDGR-T-GIMAPNSVAQQLAANNAL-S---AANIDPHTVRYVEAHATSTPLG
VIKGSAVAQDGR-T-GIMAPNAKQELVARTAL-K---VAKIDPRTIRYVEAHATSTPLG
IIRKGSAVAQDGR-T-GIMAPNAKQELVAVANAL-R---TAGVDRFTVGYVEAHATSTPLG
TLKGTAVAQDGR-T-GIMAPNKAQELVARKAL-D---VARVDASTIDYVEAHATSTPLG
VLKGSAVASDGR-T-GIMAPNAQAQILVAQKAL-K---EARVTPDSISYIEAHATSTSLG
MLKSSASALDGR-T-GIMAPNGAAQVDVARQALCR---AGNIDPCTVDYIEAHATSTPLG
VVLGTAATNSADA-ISITHPHGPTQSILSRAILDDAGVD---PLDVDYVEMHGTGTQAG
VILGTATNSADA-ISITHPHGPTQSILSSILDEAGVD---PLDVDYVEMHGTGTQAG
VILGAQTNSADA-ISITHPHGPTQSVLSTSLDEAGVD---PHDVDYVEMHGTGTQAG
VILGSATNSADA-VSITHPHGGTQEILYKRI LNNAGVD---PNDIDYVEMHGTGTQAG
VILGSATNSADA-VSITHPHGGTQEILYRSILRKAGVD---PLDIDYVEMHGTGTQAG
VILSTATNSADA-ISITHPHGKTQEALYKVKVLDQSGVD---PEETCYVEMHGTGTQAG
VIAGIATNSADA-VSITHPHAGAQLFLFQKVMDEARVD---IRDVKYVEMHGTGTQAG
CILGAATNSADA-VSITHPHAGAQYLYNKVLSNAGVD---AHEIDYVEMHGTGTQAG
VVLAAAGTNSADA-VSITHPHDLAQIHLFNQIVKRSQID---PLSVGYVEFHGTGTGAG
TILAAATNSADA-ISITHPHAPKQAQNYQVRMSQAGFS---PLDVSYVELHGTGTQAG
VINGAYTNSADA-VSITRPHVGAQAFIFKLLNEANVD---PKNISYIEMHGTGTQAG
VINAAYTNSADA-VSITRPHVGAQAFIFNKLLNDTNTN---PHEIDYVEMHGTGTQAG
VILGAYTNSADA-VSITRPHAGAQYIFSKLLRESGTD---PYNVSYIEMHGTGTQAG
VVLGAHTNSADA-ESITRPHVGAQRVIFNKILNEAAVD---PYSVSYVEMHGTGTQAG
VVLGAYTNSADA-ESITRPHVGAQRVIFNKILNEAAVD---PYSVSYVEMHGTGTQAG
VVLGAYTNSADA-ESITRPHAGAQAIFNKILNQADVD---PYTVGMC.....
VILGAHTNSADA-ESITRPLADAQYELFKLLNETGIH---PHDVSYVEMHGTGTQAG
VILGAHTNSADA-ESITRPLADAQYELFKLLNETGIH---PHDVSYVEMHGTGTQAG
VILDAKTNSADA-ESMTRPHVGAQIDNMTAALNTTGLH---PNDFSYIEMHGTGTQAG
TILDIKTNSADA-DSMTRPHVGAQIDNMSALLSTAGIS---PLDLSYIEMHGTGTQAG
VVAASGRNHSNGS-SSITSDAKAQEKLYRKMHNARVS---PNDISYVEMHGTGTQAG
VIKGTERNNSAGA-ASITYPGQEAQKLFGRLLRKASVD---PHDVGFEVEMHGTGTQAG
VIVGSAANQNHNH-SHITVPHADSQANLYNKVMKMGVNR---PETVTVYVEAHGTGTGTVG
VIPGIATNSADA-ASITVPHSPAQMTLYRRILKQAAMK---PEQVSYVECHGTGTQAG
VIPGAATNQGLS-SSLTIPSSPAQVQLYQNILNKAKMK---ADQVSYVEAHGTGTQAG
VILAAATNSADA-VSITHPHAGHSYLSRQIRQAGID---PLDVSYVELHGTGTQAG
VISGSSATNQGLS-ENLTLTHPPAQVDLYKSVLKGAKMS---PRHVSYIEAHGTGTQAG
VISGTAQNSND-TPIVVPNAPSLSLTLFRSVLETSNLN---AQDVSVEAHGTGTGTVG
VISSTGVYQNGND-TAITVPI SDSLSDLFHLVHLKAKLE---PKDISVVEAHGTGTGTVG
VIAGSAVNQSMNS-TSITVNPVLESQSVLFEKVAALAGSS---PSEFTYVEAHGTGTGTVG
VLLATGNQNMNH-TSITNVPVLESQSVLFEKVAALAGSS---PSEFTYVEAHGTGTGTVG
TILNKGTNTDGFKEQGVTFPQ-DIQEQPIRSLYQSAGVA---PESFEYIEAHGTGTGTVG
TIVNAGSNTDGFKEQGVTFPQ-DIQEQPIRSLYQSAGVA---PESFEYIEAHGTGTGTVG
TVVHAKSNTDGHKEHGITFPSEGERQAQLLQEVYSEAGID---PNSVYVEAHGTGTGTVG
QLLHAKTNCQDGYEQGITYPAGHIQKLLREFYECSIP---PSELEFVEAHGTGTGTVG
LLRGTAVNHGGRSAPGLTVPSGTAQQLRVLHAAL-A---SANLTPGDIDYVEAHGTGTGTVG
ELTGYGCSDAHMTGLR-DDGSGAFSLMNXALKNAGCQ---PQDVSVEAHGTGTGTVG
EVLGYGLSGDAGHITAPD-PEGEGALRCMAAALKDAGVQ---PEEISYINAHATSTPLG
EYLGGAVNCDAYHMTDPR-SDGLGVSTCIQSLEDAGVS---PEEVNYINAHATSTPLG
EISGYATRCNAYHMTGLK-ADGREMAETIRVALDESRTD---ATDIDYINAHGSGTRQN
EIGGWDVNRANAYHMTGLR-PDGPPELAKAIEVALRMAGRV---PDGVDYVNAHAGSGTRQN
LLRGIGVNNNDGADKGVFYAPSVKQADVVQVMMNQTKVQ---PESICYVEAHGTGTGTVG
VIRGTAENHGGAN-TLTSNPKAQADLLVRAY-R---QADIDPSTVTVYIEAHGTGTGTVG

Bacillus_subtilis_pksL_mod1
Bacillus_subtilis_pksL_mod2
Bacillus_subtilis_pksL_mod3
Bac_subtilis_mycA_pksL_mod2
Microcystis_aruginosa_mycE
Microcystis_aruginosa_mycG
Mycobacterium_leprae_pksE
Mycobacterium_tuberculosis_mas
Mycobacterium_tuberculosis_ppsC
Mycobacterium_tuberculosis_ppsD
Mycobacterium_tuberculosis_ppsE
Mycococcus_xanthus_tal_mod2
Nostoc_sp_nosB
Pseudomonas_syringae_cfa6
Pseudomonas_syringae_cfa7
Sorangium_cellulosum_epoA_mod1
Sorangium_cellulosum_epoC_mod2
Sorangium_cellulosum_epoF_mod9
Stigmatella_aurantiaca_mxaB2_B1
Stigmatella_aurantiaca_mxaC1
Stigmatella_aurantiaca_mxaC2
Stigmatella_aurantiaca_mxaC3
Stigmatella_aurantiaca_mxaD
Stigmatella_aurantiaca_mxaE
Stigmatella_aurantiaca_mxaF
Streptomyces_natalensis_PimS0
Streptomyces_natalensis_PimS2_mod3
Streptomyces_natalensis_PimS2_mod4
S_viridochromogenes_aviM
Xanthomonas_albilineans_xabB_mod2
Xanthomonas_albilineans_xabB_mod3

VIKGSVAVNHVGTVS-GISVSPVPSQADLIETCL-E---KTGIDPRITISYVEAHGTGTSLG
IIKGTAINHGKTN-GYSVVPNPAQADVICKAF-V---EAKVDPRTVSYIEAHGTGTSLG
IIIGSGINQDGKTN-GITAPSAKSQMDLERDIY-E---TYGHIHPSISYVEMHGTGKQG
VIKGSAINQDGGTV-GITAPSPAQTEVIEMAW-K---DAGIAPETLSFIEAHGTGTKLG
LIRGSVAVNHGAAA-GLTVPSPGPAQQELLRQAL-A---DARIVPEVDSYIEAHGTGTSLG
VIKGSAINNDGQKLGFTAPGGEGQIAAATEALAFAGVD----ANTISFVEAHGTGTPLG
VVRGSATNQDGRSN-GMTAPNALAQRDVIITLAL-R---LADITPDSVNYVETHGTGTVLG
VVRGTATNQDGRTE-TLTMPSEDAQVAVYRAAL-A---AAGVQPEVTVGVVEAHGTGTPIG
VVRGSVAVNQDQASS-GVTVPNGPAQQALLAKAL-T---SSKLTAAIDYVEAHGTGTPFG
VVRGSATNSDGRSN-GMTAPNALAQRDVIITLAL-K---LADVTPDSVNYVETHGTGTVLG
VIRGSAINNDGSAKMGYAAPNPAQADVIAEAHAVSGID----SSTVSYVECHGTGTPFG
LVRKVGINNDGQDKVGLTAPNSATGQAEVIRLFDRTGID----PASIGYVEAHGTGTLG
IIKGSAINNDGAMKVGFTAPSVSGQAAVIAEAQAIAGVD----AETISYIEAHGTATPLG
VIRGSVAVNQDQASN-GLTAPNGTAQQQVIRHAL-A---NAGLQPEVDVADEAHGTGTRLG
VIRGTAINQDQASN-GLTAPSGPAQEQVIQLAL-Q---NARLRLSIDVVEAHGTGTALG
VIRGSAINQDQASS-GLTVPNGSSQEVIRLKRAL-A---DAGCAASSVSYVEAHGTGTLG
VILGSATNNDGARKIGFTAPSEVGGQAQAIMAALALAGVE---ARSIQYIETHGTGTLG
LIRGSAMNQDGRST-GLMAPNVLAQEALLRQAL-Q---SARVDAGAI DYVETHGTGTSLG
LIRGSVAVNHQDQASS-GLTVPNGPSQQAVVQRAL-D---NARIAPALVGYIEAHGTGTPFG
TIRGSAINNDGKSA-GLTTPNVLSQQALIRQAL-E---SSGLKAEQSYVEAHGTGTPFG
VIRASAVNQDGGST-GLTAPNVLSQQALIRDAL-K---SANLSPKEIGYIEAHGTGTPFG
VIRGSVAVNHQDQASS-GLTVPNGTAQQQVLRAL-A---NAGVAPADVDYIEAHGTGTSLG
LIAGSAVNQDQASS-GLMVPNGPAQERVIEQAL-A---SGGLKPSQISFVEAHGTGTSLG
LVRGSVAVNQDQASN-GLTAPNVLAQQALIRQAL-Q---SANLSPKEIGYIEAHGTGTSLG
VIRGSAINNDGASN-GLTAPNTQAQAEVIRQAC-A---RAGVAPTQVYVEAHGTGTPFG
LVRGSVAVNQDQASN-GLTAPSGPAQQRVIRQAL-A---DAGLTPQVDVADEGHGTGTPFG
VVRGSAINQDQASN-GLSAPNGPSQQRVIRQAL-A---NSGLTPADIDAVEAHGTGTALG
VIRGSVAVNQDQASN-GLTAPNGPSQQRVIRQAL-A---NAEVDPDGIDVIGHGTGTALG
VLRGSVAVNQDGRTN-GIMAPCGQAQEHVMRRAL-T---AAGVAADTVDFVEAHGTGTRLG
LVRGVAVNHGGRSNS-LRAPNVNAQRQLLIRTY-Q---EAGVEPASVGYVELHGTGTSLG
VIRGSMINAGGHTS-AFSSPNPAAQAEVVRQAL-Q---RAGVAPDSISYIEAHGTGTVLG

[
Bf_PKS7
Aspergillus_terreus_at1
Aspergillus_terreus_at4
Aspergillus_terreus_at5
Gm_PKS5
Ch_PKS9
Ch_PKS8
Gz_PKS2
Gm_PKS2
Ch_PKS10
Ch_PKS5
Ch_PKS7
Ch_PKS1
Didymella_maydis_PKS1
Gz_PKS4
Gz_PKS7
Gm_PKS7
Ch_PKS6
Gz_PKS1
Ch_PKS4
Gm_PKS6
Nc_PKS1
Bf_PKS8
Nc_PKS2
Ch_PKS2
Aspergillus_terreus_lovF
Penicillium_citrinum_mlcB
Ch_PKS3
Nc_PKS3
Bf_PKS2
Gz_PKS5
Gm_PKS1_CAC44633
Gm_PKS1
Bf_PKS5
Gm_PKS9
Aspergillus_terreus_lovB
Penicillium_citrinum_mlcA
Gz_PKS9

361
.....
DAVEMESVTSVSPG-----RTMR-----PHPLHIGSVKGNVGH-GEAAAGVTALIKTL
DATEMNSVLSVFPYK---RMQMP-----ARPLYLGSAKANIGH-AESASGVSSLIKVM
DGTEMKSITNVFAPRD---KRRQP-----DQLIHLGALKANIGH-GEASAGVASLIKT
DPIEASAI SGAFLEYCSEQE-----PMVVGAVKTNIGHLEGSAG.....
DPCEAAAINSVFSA-RTPED-----PIFVGALKSNMGHPEGASG-IAGVIKTL
DPIEASAIQAFFRDGRSLDR-----PLYIGSVKASIGHLEGSAG-LAGVIKAV
DPIEVKAI GRCFQEHRSHE-----PLYVGSVKANIGHLE-GASALASLVKCI
DPIEVEAI GRCFQAHRSPE-----PLYVGSVKANIGHLE-GASALASLVKCV
REIQLRLVPLVTSSAIGQKKS-----PIYIGALKSNIGH-TEGNSGVSAFIKAV
DAIEATGIAQAF---KSRKDK-----PLYLGSVKASVGHLEGAAG-VAGI KSV
DPLELRALTNFCGANRPEDR-----PLIIGSVKSNIGHLEGSAG-IAAVKAV
DTTEASALS KVFSPGRR---L-----PLLIGSVKTNIGHLEGSAG-LAGVVKSI
DTTEARAI SNVFGPGRQ---V-----PLAIGSVKTNIGHLEAASG-LAGVLKCI
DPVETGAIHRTIGQGASKNRK-----LWVGSVKPNIGHLEGAAG-VASVIKGV
DWRELKAI SESLGRVGDIDN-----PIVVGSLKPNIGHLEGAAG-VAGLIKGV
DWRELKAI SESLCTVRDIDN-----PMVVGSIKPNIGHLEGAAG-VAGLIKGV
DPLEAKAILKTVGSVEGRKS-S-----LYVGSVKTNIGHLEGAAG-VAGVIKAA
.....
DKAEALAFTTTTGKDRREK-----IIVGSVKSNLGHLENASG-LASVIKTV
DPIECSVIGNVF--GKTRER-----PVYVGSVKSNIGHLEGSAG-LAGLVKTI
DPIEATAIHNVLGQERSPRN-----PLWIGSVKSNIGHLEGSAG-IAGVIKAA
DPIEATALYNVFGEGRNARN-----PLFIGSVKSNIGHLEGAASGGIISVIKTA
DPIEATAIYKAVGRYLSADD-----PLYIGSSKPNVGHLECASG-LVSVIKGV
DPIEISALQEVFSTASRTLED-----PLLIGSVKANVGHLE-BSSSGFPPLIKAA
DPLELAVISAAFFPGQ-----IQVGSVKANIGHTEAVSG-LASLIKVA
DPLEISAISAAPFKGH-----PLHLGSVKANIGHTEAASG-LASIIKVA
DPLEVGAIAAIVFKDSRPTTT-----PLPIGSIKPNVGHLETCASG-LASIVKVV
DPIEVKAIARVFKDSRKGNGEDAL-----LRIGSVKTNIGHTETASG-VAIIKVA
DLAETKSIANVFCTE--RKKT-----LYIGSVKSNIGHLESASG-AAGLLKAI
DREEIGSISKVFCEDSGRTDD-----LYIGSVKTNIGHLEATSG-IAGLLKSI
DPREAEASTVRSI--PDV-----KDDKLYVGSIKTVLG-HTEGTAGLASLIGTA
DPREAEAIYRAFY--SDV-----KDDKLYVGSIKTVLG-HTEGTAGLASLIGTA
DPQESQAIDS AFFSGDPK-----KLETMAVGSIKTIIGYHTEGTAGLASLIGSV
DPKEAEAI FRAFFGDDQP-----HDEKLNVISIKTVLG-HTEGTAGLASLIGTM
DPQEAETAIATAFFGHEQVARSDG-----NERAPLFVGSIAKTVV-HTEGTAGLMLKAS
DPQEAETAIATAFFGHKGDITSD-----GEKDELVGSIKTVLG-HTEGTAGLMLKAS
DPQEAETAIATAFFPNETDSVTNGTNGLLSEADNLLVGSIKTVIG-HTEGTAGLMLKAC

Streptomyces_coelicolor_act
Streptomyces_avermitilis_pks9
Bacillus_subtilis_pksK_mod2
Bacillus_subtilis_pksK_mod3
Bacillus_subtilis_pksL_mod1
Bacillus_subtilis_pksL_mod2
Bacillus_subtilis_pksL_mod3
Bac_subtilis_mycA_pksL_mod2
Microcystis_aruginosa_mycE
Microcystis_aruginosa_mycG
Mycobacterium_leprae_pksE
Mycobacterium_tuberculosis_mas
Mycobacterium_tuberculosis_ppsC
Mycobacterium_tuberculosis_ppsD
Mycobacterium_tuberculosis_ppsE
Mycococcus_xanthus_tal_mod2
Nostoc_sp_nosB
Pseudomonas_syringae_cfa6
Pseudomonas_syringae_cfa7
Sorangium_cellulosum_epoA_mod1
Sorangium_cellulosum_epoC_mod2
Sorangium_cellulosum_epoF_mod9
Stigmatella_aurantiaca_mxaB2_B1
Stigmatella_aurantiaca_mxaC1
Stigmatella_aurantiaca_mxaC2
Stigmatella_aurantiaca_mxaC3
Stigmatella_aurantiaca_mxaD
Stigmatella_aurantiaca_mxaE
Stigmatella_aurantiaca_mxaF
Streptomyces_natalensis_PimS0
Streptomyces_natalensis_PimS2_mod3
Streptomyces_natalensis_PimS2_mod4
S_viridochromogenes_aviM
Xanthomonas_albilineans_xabB_mod2
Xanthomonas_albilineans_xabB_mod3

DRHETAAYKRAL-----GEHARTPVSSIKSMVGHSLGAIGSLEIAA-CV
DAHETDAFKRTL-----GDHAYRTPVSSIKSMIGHSLGAIGSIEIAA-CA
DPIELAAALNTVYRQYTNKTQF-----CGIGSVKTNIGHL-DTAAGLAGCIKVV
DPIEINGLKAAPKELSNMRGESQPD---VPDHRCGIGSVKSNIGHL-ELAAGISGLIKVL
DPIEIQGLVKAQFRQYTDQR-----QFCSIGSVKSNIGHA-ESAAGISGLSKVA
DPIEITGLSKVFTQETDDK-----QFCAIGSAKSNIGHC-ESAAGIAGVTKVL
DPIELEALSTVFQEKTDKK-----QFCAIGSVKSNIGHT-SAAAGVAGVQKVL
DPVEFNGLCKAFKVFTEKK-----QFCAIGSVKANIGHL-FEAAGIVGLIKSA
DPIELNAIASVY-----GKRSD-----PLYVASVKTNIGHL-EAAAGMAGIIKTI
DPIEVDALAKVYQ--GANEGE-----CALGSVKTNIGHM-QIASGIAGLIKAT
DPIEFESLAATY-----GAG-----TPCALGSAKSNMGS-TASAGITGLIKAI
DPIELDLSLKV-----SDRAGS---DQ-L--VIGSVKTNLGH-AAAAGVAGLMKAV
DPIEFESLAATY-----GLGKGQ---GE-SPCALGSVKTNIGHL-EAAAGVAGFIKAV
DPIEIQGLRAAFVVSQTSRSA-----PCVLGSKSNIGHL-EVAAGIAGLIKTI
DPVEALSEAFRTFTDRRGY-----CRLGSKSNLGH-ETVAGITGLIKTA
DPIEIAALTAQAFSQTDDKGF-----CAIGSLKTNLGH-DTAAGVAGLIKTV
DPIEANALLATY-----GQ-RPA---ER--PLLLGSIKSNIGHT-QAAAGVASLMKMV
DPIEAGALINTY-----GRGRPP---ER--PLWLGLSKSNIGHT-QLAAGVASVIKMV
DPIEIQALNAVY-----GLGRD---VA-TPLLLGSKSNIGHT-DYVAGITGLIKVV
DAIETAALRRVFRDASTRRS-----CAIGSVKTNIGHL-ESAAGIAGLIKTV
DPIEVDALRAVM-----GPARA---DG-SRCVLGAVKTNLGH-EGAAGVAGLIKAA
DPIELRALGAVL-----GKGRP---AD-RPFFIGSVKTNIGHL-EPAAGIAGVIKTI
DPIEAALRETY-----GVRP---DG-SVCGIGSVKTNIGHL-ESAAGIAGIKVV
DPIEVEALRETY-----GVRD---DG-FPCFLGSAKTNIGHL-EAAAGMAGLIKVL
DPIEVQALANVL-----GQGR---SE-QRLLGSKSNIGHT-EAAAGIAGLIKVV
DPIELQALARVL-----GTGRS---AE-TPLFVGSVKTNVGHL-EATAGLTGLVTKTA
DPIEAQALLATY-----GSRG---AK-QALVVGAAKTNIGHT-ESAAGIAGIKMV
DPMEARALGEAL-----GAGRPS---DK--PLHIGSCKTNIGHL-EAAAGITGLIKVA
DPIEAQALLATY-----GQGRPV---ER--PLWLGSVKSNGFHT-QAAAGVVGVIKTV
DPIEAQALLATY-----GPRDP---DS--PLLLGSVKSNGHT-QSAAGVAGVIKMV
DPIEAQALLATY-----GQGRDP---EH--PLLLGSVKSNGHT-QMASGVASVIKMV
DPMEIGAAVY-----GRDRS---GQ-EPCAVGSVKSNGHL-EGAAGVAGVIKAI
DPIEIQALKEAF-----IALGAQ---AAPSNCIGSVKSALGH-EAAAGLTGLIKVL
DAVELGALNKVFDKRAAP-----CPIGSLKANIGHA-ESAAGIAGLAKLV

[
Bf_PKS7
Aspergillus_terreus_at1
Aspergillus_terreus_at4
Aspergillus_terreus_at5
Gm_PKS5
Ch_PKS9
Ch_PKS8
Gz_PKS2
Gm_PKS2
Ch_PKS10
Ch_PKS5
Ch_PKS7
Ch_PKS1
Didymella_maydis_PKS1
Gz_PKS4
Gz_PKS7
Gm_PKS7
Ch_PKS6
Gz_PKS1
Ch_PKS4
Gm_PKS6
Nc_PKS1
Bf_PKS8
Nc_PKS2
Ch_PKS2
Aspergillus_terreus_lovF
Penicillium_citrinum_mlcB
Ch_PKS3
Nc_PKS3
Bf_PKS2
Gz_PKS5
Gm_PKS1_CAC44633
Gm_PKS1
Bf_PKS5

421
.....
LAFQKQAIKPHVGIK--TMLNP--KFP-GLDSLNIHIALEQ--VPWPRN--DARTRYALVN
LMMKHGIBIPPHCGIK--NRINH--NYPLYLEQRGVRIALNV-TAWKRDALPSGKRSVFLN
IMMQKNAIPHHVGIK--TTMKN--TFPHDLNERGVRIAFKE-TPWVRPDGGK-RR-AYLN
.....
LVLEKGIIPPNVYPERINPAVTAAGPNLRF--PLEPVTWP-T-SG-V-----RRASVN
LSLEKAVIPPIAMFEKVNPAIDAADLQLAF--PKEAIPWP-A-SE-V-----RRASVS
CNLEKGVILPNALLQKMPAMNADTY--SIEVPIHNQWQPV-----Q--GLRRVSLN
LILEKGVIPPNALLQKNTALKADSY--NITVPTKSIWPT-----E--GLRRVSLN
LTLESGIIPANAHFKEVNHAI---PKKWHFKFPTVATPYPKTPSG-----VRRVSN
LALENGVIPPQANFEKANPKIPFQKWNLS--IADKAVPWP-T-QG-L-----RRVSN
LMLEQEVILPNQNFEPNPRIPFVDWKL--RVATNVQPWD-S-SG-P-----LRVSN
LMLEQGVILPNRNFEEFKHENPAGKMELAYKVPTTLECWN-N-VK-T-----RRVSN
LMLENEIILPNRNFKHANLNIPLEEWKL--RVPTTVEPWN-S-MT-T-----RRASVN
LAMENSLIPPNIHFAFPNPEIPLDEWNM--AVPTKLTWPWAA--R-T-----KRMSVS
LLEHAKIIPPNINLDPNPDIDFKNKV--KVPTKLLDWPPL--G-L-----RRVSN
LLEHAKIIPPNINFEKPNPDIDFKKWKV---PTELLDWPVS--G-L-----RRVSN
LAVERGLIPQNLWFEKLNPEINLPE-NVEI--PLKLTWPW-S-DG-P-----RRASIN
.....
LMIENGVIPPVPTFEKSSEQLPVDQ--MGITIPREAILWPAGTH-----KRASIN
YSLESGVISPTYGLENVNPKIKLDEWKINL--PTEKIKWP-A--G-L-----RRASIN
LMLERGFILPNHDFKQPNRIPWKEWNLE--VPVTPQRPWA---RG-K-----KYISVN
MMLERGFILPNHDFKTPNENIPFSKWLQ--VPISQRPWP---RA-K-----KFASVN
LSLYGFILPNADDFKENPGIPFKEWNMT--VAKQKQWP---AH-K-----KYACVN
MMLKGLVVPNANFENESMNSHLKEKNMR--VPISTQPW---KG-K-----TYIAIN
LAVEKGVIPPNARFLQPSKLL-LKDTHI--QIPLCSQSWIPT-DG-V-----RRASIN
LALKGLIIPPNARFLQKSNKMLDQKNI--KIPMSAQDWPVK-DG-T-----RRASVN
KAIKGLIIPPNMHFDVNLNPKIK--PFYDNLEVPTKAIWPEPTHKGGP-----RRASIN
LALERGQIPPSINFETPNAKLSLDEWKL--KVPTLEEWVGK-DG-I-----RRASIN
LVLEKGLIIPPNVNLNSHKGLDLEQSN--VVPKIEKLSVDDEG-K-----SRIAIN
LILKHGQIPATLNFIFKPKPSLKYEKKI--KIPSELTKLPTPQNG-P-----PRVSLN
LAIQNKTIIPPNMHFDVNLNPKIK--PFYDNLEVPTKAIWPEPTHKGGP-----RRASIN
LAIQNKTIIPPNMHFDVNLNPKIK--PFYDNLEVPTKAIWPEPTHKGGP-----RRASIN
MAIKHGVFPNPNLHFQNLSPKVA--PFYNHLHIPTTATPWPPELPSGVP-----RRISCN

Gm_PKS9 MAMKNATVPPNMHFETLNPDIIE--PFYSNLEVPVTSKAWPNVH-GSV-----RRASIN
Aspergillus_terreus_lovB FAVRHGVIIPPNNLLFDKISPRVA--PFYKNLRIPTEATQWPALPPGQP-----RRASVN
Penicillium_citrinum_mlcA FAVRNGVIPPNNLLFEKISPRVA--PFYTHLKATEATEWPIVAGQP-----RRVSVN
Gz_PKS9 MALKHAV-PPNLLFNRLNPALE--PFTK.....
Ch_PKS16 QAMQNSVTPPNLWFQQINPKLK--PFYGNLQIPHTHALIWPKPFGRRP-----KRASIN
Bf_PKS6 LALKEKTIIPPNNLHFQSLNSAIE--PFYGNLNVPTAPLPWP--AVEGP-----LRASVN
Bf_PKS3 LALQHSFIPPNNMHFQELNPEIE--PFYQNLKVVTSLQWPWPLIPDNTP-----RRASIN
Ch_PKS17 EAVRRVAVPPNMLLQKLNPAIE--PFCVNLKVPTQSLPWPPLPLGGP-----RRASVN
Gz_PKS10 EAVKTGIIPPNNLQFETLNPAVK--PYTSNLKLPVESQWPWPKLLEGFP-----RRASVN
Gm_PKS10 EAVKRGVIPPNNLQFENLNPAIK--PYVSNLRLPTESKFPWPKLAPGSA-----RRASVN
Nc_PKS4 HAMINGTVPPNNLHFNRLQPAVA--KYYSHLVVPTKALAWPSVAAGQP-----RRATVN
Bf_PKS4 LAIQHGIIIPPNLGFQQLNKKVA--PYCTNMDVVTSVQWPWAMASNTP-----RRVSVN
Gz_PKS11
Ch_PKS14 MSLENRTIPPNIKFTTPNPKIPFKE--AKLTVVPEPTPFPE-----D---RCERVSVN
Ch_PKS12 LALENRVIPPNIKFTTPNPKIPFLE--KDLKVPTDCLGWPK-----D---KLERVSVN
Gm_PKS12 LALEHGMIIPPNIKLVNPNPK-PT--MPSSVPLSPTQFPS-----D---RQNRVSVN
Ch_PKS13_pseudogene LTLEHKTIPPNIYFNIPNPRSTFKS--ARLAVPTEPLPWPT-----D---HTKRISVN
Ch_PKS11_FUM1 LALENRKIPPNIINFSIPNPKIPFNE--ANMVVPCVLEVPWE-----N---QPLRASVN
Gm_FUM1_AAD43562 LALENRTIPPNIINFSIPNPKIPFSE--MNMAVPVDAIPWPR-----D---RPLRVSVN
Gm_FUM1 LALENRTIPPNIINFSIPNPKIPFSE--MNMAVPVDAIPWPR-----D---RPLRVSVN
Ch_PKS15 LSMEHQIIPPNNVFETPNPKIPWKE--ANLRVPTCEMKWPS-----D---RLERASVN
Nc_PKS5 LALEHGAIPIIDYLRNPNIDFEGA--KVQPVTEVTPWPKD-----R---L-QRASIN
Gm_PKS15 LAIENELIPTTIGIENFNPNKIFQDW--KLHVVEATAWPSNIP-----IRRASVN
Nc_PKS6 MALEKGIIPPNTYGVKNINPKIKTDEW--NVQIVTETTPWPKLPHNAGR--LFRRAGVN
Gm_PKS14 KSFEKGVPPTRGIVNLPKLVLAER--NLMIAQDAHDWPRDL-----RRASIN
Bf_PKS9 LAMEKRTIPATIGIKKLNXPVDFKGG--RIEIVVQKMTWPWPKGFS-----VCRASIN
Gm_PKS13 LALEKGEIIPPNTNYKTTPNPKIDFDGW--RVRVPTAPQWPWS-----K---SIRRASVN
Bf_PKS10 MAIEKAIIPPNTLNLKRLNSRIDWDQ--AGVKVVEALEWPGNED--DVR--RAGVC
Bf_PKS11 LMLEHNMIIPPQVNFEEKANPEIDLAL--NLRIPITLESRP-----LKRISVN
Penicillium_patulum_6MSAS LAIQKGVLPQANLTKLNSRIDWKT--AGVKVVEATPWPE--D-----PIR--RAGVC
Byssoschlamys_nivea_6MSAS LALKKGIILPPQANLQKLNKVDWDH--AGVKVVEATKWPES--D-----EVR--RAGVC
Aspergillus_terreus_pksM MAVEKAIIPPQNLKRLNSRIDWDQ--AGVKVVEALEWPGNED--DVR--RAGVC
Aspergillus_parasiticus_pksL2 MSVQKGIILPPQANLTKLNRVNWNE--GVRVVDIEDWPSSGY-----P-R--RAGVC
Penicillium_griseofulvum_pks2 LVLRHGQVPPQANLQTLNSKIAWKK--SLCPCARELVTLPH--GT-----PSRPLRAAVA
Ch_PKS25 LSVQRGIIPPQRLLENFNTNIDWDS--SGLKVLREGKIWPENGW-----WRAAVC
Nc_PKS7 LMLHKNAIIPPHVGIKKSIIINK--TFPKDLSDRNVNIAFHL--TPFKR--RDGKPRR--IFVN
Colletotrichum_lagenarium_pks1 MMRKNAIIPPHVGIKKE--INK--TFPKDLSENVNIAFHL--TPFKR--RDGKPRR--IFVN
Nodulisporium_sp_PKS1 MMLKKNAIIPPHVGIKKE--INK--TFPKDLGERGVNIAFHK--TPFTR--KDGKPRK--VFN
Xylaria_sp_PKS12 MMLQRNAIIPPHVGIKKE--INK--TFPKDLGQGVNIAFHK--TPFLR--KDGKPRR--LNV
Bf_PKS12 MMLQKNAIIPPHVGIK--TINQ--GFPKDLAERNVHIAFKN--TPLPRKKSAGPRR--IFVN
Glarea_lozoyensis_PKS1 MMLQKNAIIPPHVGIK--TINV--GFPKDLAERNVHIAFHK--TPL--RARAGMPRR--VFN
Monascus_purpureus_PKS1 LMLRKNAIIPPHVGIK--DINK--GFP--DLAARNIKIAFDL--TPLPR--EAGKLRK--VFN
Bf_PKS13 MMLKKSMIIPPHVGIK--TKINQ--TFPKDLDRNVLNMAFKP--TPFPRPTNGK--RY--VFMN
Ch_PKS18 MMLKKNAIIPPHVGIK--VMNK--TFPKDLAQRNVHIEETQ--VAWR--KGAKRK--IFLN
Bf_PKS14 LVFQKSTIPPHVGIK--TGLNP--ALPKDLQRGVIVPYTA--TSWEKS--SSQKRLAMVN
Ch_PKS20 LMYQKSAIIPQIGVKK--LNP--TLPDLEERTIGLNWEY--VEWPRPKTGS--RLAIVN
Aspergillus_nidulans_wA LMMKENMIIPPHVGIK--TKINH--NFPTDLAQRNVHIAL--QPTAWNRFSGKRQI--FLN
Aspergillus_fumigatus_alb1 LMLKKNMIIPPHVGIK--TKINH--NFPTDLAQRNVHIAF--EPTPWRINLTPRR--IFVN
Gz_PK12 LMMEKNMIIPPHVGIK--TKINH--HFPTDLTQRNVHIAK--VPTSWTRSGQANPRI--AFVN
Gm_PKS3_CAC88775 QMLKKDTIVPHVGIK--NKINH--RFPTDLQQRNVRIAM--GPTQWKKGTEINPRR--VFN
Gm_PKS3 QMLKKDTIVPHVGIK--NKINH--RFPTDLQQRNVRIAL--EPTGWKKGTEINPRR--VFN
Gz_PKS12 QMMQKNTIIPPHVGIK--TKINH--RFPTDLAQRNVHIAF--EPTPWRINLTPRR--IFVN
Gm_PKS4_CAC92399 LMMQKNTIIPPHVGIK--TKINQ--GFPKDLDRHGVRIALKDGVDWTRPEGGK--RR--VLVN
Gm_PKS4 LMMQKNTIIPPHVGIK--TKINQ--GFPKDLDRHGVRIALKDGVDWTRPEGGK--RR--VLVN
Aspergillus_parasiticus_pksL1 MMMQHDITIPPHVGIKPGSKINR--NFP--DLGARNVHIAF--EPKWPRTHTPRR--VLIN
Aspergillus_nidulans_pksST LMMKNDITIPPHVGIKPGSRINR--NYP--DLPARNVHIAF--EPKWPRTDTPRR--VLIN
Gz_PKS13 MMFQKNIIPPHVGIK--ALNP--NFP--PLSEINIEIP--SEPSTFESVPSQP--RRILLN
Bf_PKS15 LILREGSIPTQPDWPF--KLNH--KFP--DLEASNIMIA--DGKGRVLRPNNGDKRRIVVN
Bf_PKS19 LMMKHKRIIPRQASYK--ELNP--QIPS--LDQDRMAIPTSV--MPW-----NPPVRLACVN
Ch_PKS22 AMLQKQRIIPPLASFK--SLNP--KIPA--LAPDKMAVAKQVE--AW-----DAPLRAACVN
Bf_PKS16 AMINQGVIPPLASHN--NLNP--KIPA--LEPDRMAITTKAI--PW-----TAKFRAACVN
Ch_PKS19 LMLQKNAIIPPHVGIK--TEVNP--RLPKDLQQRNVAIFFSM--TEWKS--GKKKRIAAVN
Ch_PKS23 AMLQKKSIPPHASFN--TLNP--KIPS--LGPDKITIARNLE--PW-----NIPFRALVN
Ch_PKS21 CMLHEATIPQASHT--VMNP--AIGA--TKSDGIQITLQP--KPW-----NADFKAALIN
Bf_PKS17 LMVQEGEIPQASFT--SVNP--NLGL--TPDDKIEISAQL--KPW-----NADFKAALIN
Bf_PKS18 LMMKHSKIPKQANHL--SLNP--KIRA--LEPKIKIPLQT--TPW-----EADLRIASIN
Bf_PKS20 LMMKHGKIPIQIQAF--TLNP--NIPA--LETDKMAISTSLQKQW--SDDLRLALVN
Homo_sapiens_FAS LSLEHGLWAPNLHFHSPNPEIP--ALLDGRQLVVDQPLPVRGG--VGIN
Gallus_gallus_FAS LSLEHGLWAPNLHFNDPNPDIP--ALHDGSLKVVCKPTPVKGG--VGIN
Caenorhabditis_elegans_FAS LSIERQLIIPPNLHYNTPNQYIP--GLTDGRQLVVTEPTALPGL--IGIN
Bombyx_mori_FAS_p270 IAYTTGYIPPNNLNSIPREGVS--ALVDGRMNVITDKQAWNRGM-----SGIN

Ch_9497_PKS24_3p_of_nps7 LAMQHNKIPRTLHAEKPTSVAVDWQG--ANMALVQEPQPWLQPAD-----R--PRRAGVS
Nc_3_oxoacyl_ACP_synthetase LAIAEGXIPPTLNLTSPNVGVGFNFVVPVQ-----AQQ-KEVK--VAMSN
Homo_sapiens_3_oxoacyl_ACP_synth LACYYQKLPPTLNLDCEPEFDLNYVPLK-----AQEWKTEKRFIGLTN
Glycine_max_beta_ketoacyl_synth KAITTGWLHPTINQFNPEPAVDFDTVANV-----KQQHEIN---VAISN
Streptomyces_coelicolor_act LALEHGVPPTANLRTSDPECDLDYVPLE-----ARERKLR---SVLTV
Streptomyces_avermitilis_pks9 LAIDAGLIPPTANLHEPDPELGLDYVPLS-----ARQADLR---TVLKV
Bacillus_subtilis_pksK_mod2 MSLYHQELAPSVNYKEPNPNTDLAS--SPFYVVDQKKTLS-----REIKTHRAALS
Bacillus_subtilis_pksK_mod3 LQMKHKTLVKS LHCEFTLNPLYQLTD--SPFYVQEKQEWK-SVTDTRDNE--LPRRAGIS
Bacillus_subtilis_pksL_mod1 LQLHHQKLVPSLHSEELNPNYVDFEK--SPFYVQHETETWKQPVIKENGEDVPYPRRAGIS
Bacillus_subtilis_pksL_mod2 LQMKYRHVAPSLHSNVLNPNIDFLN--SPFKVQEELEEWKRP IISVNGKDIELPRIAGVS
Bacillus_subtilis_pksL_mod3 LCMNHKTLVPTLNFTTPNEHFEFEH--SPLYVNTTELKPEW-----TADGK----PRRACVS
Bac_subtilis_mycA_pksL_mod2 LMLNHKIPPLAHLFNKPNPLIPFHS--SPFYVNQEVMDFTPE-----DRPLRGGIS
Microcystis_aruginosa_mycE LILQQGEIPPHLHFQSPNPLINWQD--HPIEIPTQNIWPNNNK-----V--P-IAGVS
Microcystis_aruginosa_mycG LALKYRVIPPTLHFQNPQINFSQ--TPFYINNEAISWT----TKQDKSEKLP RRAGVN
Mycobacterium_leprae_pksE LALQHAYIPRNHFQTNPAIDAST--TRLFVPTESAPWPA-IP-----G--PRRAGVS
Mycobacterium_tuberculosis_mas LSLRHGVPPLLHFNKPNPLIPFHS--SPFYVNQEVMDFTPE-----DRPLRGGIS
Mycobacterium_tuberculosis_ppsC LAVHNGYIPRHLNFHQLTPHASEAA--SRLRIAADGIDWPT-TG-----R--PRRAGVS
Mycobacterium_tuberculosis_ppsD LAVQRGHI PRNLHFTRWNPAIDASA--TRLFVPTESAPWPA-AA-----G--PRRAAVS
Mycobacterium_tuberculosis_ppsE LCLKNKALPATLHYTSPNELRLDQ--SPFVQSKYGPWE----CD-----GVRRAGVS
Mycococcus_xanthus_tal_mod2 LSLRQGEVPPPTLHVTVQNPKLELTD--SPFVIADRLAPW-----SLPGPRAAVS
Nostoc_sp_nosB LALQNKMLPPSLHFETPNPKIDFAN--SPFYVNTTLEWR----T-----NTTPRRAGVS
Pseudomonas_syringae_cfa6 MALRTGTLPRSLHITAPSTEVDWST--GAIELLSERRPWPVNAD-----R--PRRAGIS
Pseudomonas_syringae_cfa7 MALNQGVLPKTLHAQEP SRKIDWSE--QTVRLLHRARPWP-ETD-----Q--PRRAGVS
Sorangium_cellulosum_epoA_mod1 LSLQHGGIIPAHLHAQALNPRISWGD--LRLTVTRARTPWPDPWT-----HT--PRRAGVS
Sorangium_cellulosum_epoC_mod2 LALEHRQLPPSLNFESPNPSIDFAS--SPFYVNTSLKDNW----T-----GSTPRRAGVS
Sorangium_cellulosum_epoF_mod9 LALHHEIPRNLHFHTLNPRIRIEG--TALALATEPVPWP-RAG-----R--PRFAGVS
Stigmatella_aurantiaca_mxaB2_B1 LSLQHKIEIPPHLHFRTPNPHVEWDR--IPARVPERVWPWAHEG-----RRIAGVS
Stigmatella_aurantiaca_mxaC1 LAMRHQSIIPHLHLKQVNPRIQLEG--SALTIPTRLTWPWKASG-----Q--PRRAGVS
Stigmatella_aurantiaca_mxaC2 LALRHEAIPKHLHFKALNPRITIED--TPFVIPTQMPPVPSQ-----R--RRYAAVS
Stigmatella_aurantiaca_mxaC3 LAHRHGVPVPHLHLKLNPHIELNG--FPLDIPTQVTPWPERAR-----PRIAGIS
Stigmatella_aurantiaca_mxaD LALRNETIPPNLHFKRLNPDIVLDG--APVVVPTEPSRWPRAEQ-----PRFAGVS
Stigmatella_aurantiaca_mxaE LSMQHGVAPPIAHFQRLNPRIDFAG--ESITIPALHPWPAREE-----R--KRRGAIS
Stigmatella_aurantiaca_mxaF LSIKNRALPPSLHFETPNPLIPFED--LRLEVQRTLSDWPEPD-----R--KLVAGVS
Streptomyces_natalensis_PimS0 LALRHGVLPTLHVDAPS AKVDWSA--GSVRLLEAREPWPRESG-----R--TRRAGVS
Streptomyces_natalensis_PimS2_mod3 LALRNGQLPRTLHADTPSSHIDWTP--GTVSLLTEPTDWP-EGE-----R--PRRCAVS
Streptomyces_natalensis_PimS2_mod4 MAMRHAAPRSLHIDRPPSHVDWSS--GAIQLLLEPLPWP-ATG-----H--PRRAGVS
S_viridochromogenes_aviM LALDEDRI PASLLDGDPNPEIDWAG--LDIRLATRALPWPPEP-P-----H--PRRAAVS
Xanthomonas_abilineans_xabB_mod2 LMLKHGQAGTRHFSTLNPLIDLRG--TSFEVVAQHRAWPSQVGI---HGTLLPRRAGIS
Xanthomonas_abilineans_xabB_mod3 LQFRHGELVPSLNAFPLNPIYIEF---GRFQVQQQPAPWPR-----RGAQ---PRRAGLS

[481	487]
Bf_PKS7	
Aspergillus_terreus_at1	NFGASG	
Aspergillus_terreus_at4	NFSAAG	
Aspergillus_terreus_at5	NFGASG	
Gm_PKS5	
Ch_PKS9	SFGYGG	
Ch_PKS8	SFGYGG	
Gz_PKS2	SFGVGG	
Gm_PKS2	SFGFGG	
Ch_PKS10	SFGISG	
Ch_PKS5	SFGVGG	
Ch_PKS7	SFGYGG	
Ch_PKS1	SFGYGG	
Didymella_maydis_PKS1	SFGYGG	
Gz_PKS4	GFGMGG	
Gz_PKS7	CFGFGG	
Gm_PKS7	C.....	
Ch_PKS6	SFGFGG	
Gz_PKS1	
Ch_PKS4	STGYGG	
Gm_PKS6	SFGYGG	
Nc_PKS1	NFGFGG	
Bf_PKS8	NFGFGG	
Nc_PKS2	NFGFSG	
Ch_PKS2	NYGFGG	
Aspergillus_terreus_lovF	NFGFGG	
Penicillium_citrinum_mlcB	NFGFGG	
Ch_PKS3	NFGYGG	
Nc_PKS3	NFGYGG	
Bf_PKS2	SFGYGG	

Gz_PKS5	SFGYGG
Gm_PKS1_CAC44633	SFGFGG
Gm_PKS1	SFGFGG
Bf_PKS5	SFGFGG
Gm_PKS9	SFGFGG
Aspergillus_terreus_lovB	SFGFGG
Penicillium_citrinum_mlca	SFGFGG
Gz_PKS9
Ch_PKS16	NFGFGG
Bf_PKS6	SFGFGG
Bf_PKS3	SFGFGG
Ch_PKS17	SFGYVF
Gz_PKS10	SFGFGG
Gm_PKS10	SFGFGG
Nc_PKS4	SFGFGG
Bf_PKS4	SFGFGG
Gz_PKS11
Ch_PKS14	SFGLGG
Ch_PKS12	NFGIGG
Gm_PKS12	SFGIGG
Ch_PKS13_pseudogene	SFRVGG
Ch_PKS11_FUM1	SFGIGG
Gm_FUM1_AAD43562	SFGIGG
Gm_FUM1	SFGIGG
Ch_PKS15	SFGIGG
Nc_PKS5	SFGYGG
Gm_PKS15	SFGYGG
Nc_PKS6	SFGYGG
Gm_PKS14	SFGYGG
Bf_PKS9	SFGYGG
Gm_PKS13	SLGIGG
Bf_PKS10	SFGYGG
Bf_PKS11	SFGYGG
Penicillium_patulum_6MSAS	SYGYGG
Byssochlamys_nivea_6MSAS	SYGYGG
Aspergillus_terreus_pksM	SYGYGG
Aspergillus_parasiticus_pksL2	SYGYGG
Penicillium_griseofulvum_pks2	SYGYSG
Ch_PKS25	SHSYGG
Nc_PKS7	NFSAAG
Colletotrichum_lagenarium_pks1	NFSAAG
Nodulisporium_sp_PKS1	NFSAAG
Xylaria_sp_PKS12	NFSAAG
Bf_PKS12	NFSAAG
Glarea_lozoyensis_PKS1	NFSAAG
Monascus_purpureus_PKS1	NFSAAG
Bf_PKS13	NFSAAG
Ch_PKS18	NFSAAG
Bf_PKS14	NFGAAG
Ch_PKS20	SFGAHG
Aspergillus_nidulans_wA	NFSAAG
Aspergillus_fumigatus_alb1	NFSAAG
Gz_PK12	NFSAAG
Gm_PKS3_CAC88775	NFSAAG
Gm_PKS3	NFSAAG
Gz_PKS12	NFSAAG
Gm_PKS4_CAC92399	NFSAAG
Gm_PKS4	NFSAAG
Aspergillus_parasiticus_pksL1	NFSAAG
Aspergillus_nidulans_pksST	NFSAAG
Gz_PKS13	NFDAAV
Bf_PKS15	SFDAAG
Bf_PKS19	SHGAAG
Ch_PKS22	SYGAAG
Bf_PKS16	SYGAAG
Ch_PKS19	NFGAAG
Ch_PKS23	NYGAAG
Ch_PKS21	NYGASG
Bf_PKS17
Bf_PKS18	NYGASG
Bf_PKS20	NFGASG

Homo_sapiens_FAS	SFGFGG
Gallus_gallus_FAS	SFGFGG
Caenorhabditis_elegans_FAS	SFGFGG
Bombyx_mori_FAS_p270	SFGFGG
Ch_9497_PKS24_3p_of_nps7	AFGIGG
Nc_3_oxoacyl_ACP_synthetase	SFGXGG
Homo_sapiens_3_oxoacyl_ACP_synth	SFGFGG
Glycine_max_beta_ketoacyl_synth	SFGFGG
Streptomyces_coelicolor_act	GSGFGG
Streptomyces_avermitilis_pks9	GSGFGG
Bacillus_subtilis_pksK_mod2	SFGLGG
Bacillus_subtilis_pksK_mod3	SFGIGG
Bacillus_subtilis_pksL_mod1	SFGATG
Bacillus_subtilis_pksL_mod2	SFGAGG
Bacillus_subtilis_pksL_mod3	SFGYSG
Bac_subtilis_mycA_pks1_mod2	SFGFSG
Microcystis_aruginosa_mycE	SFGFSG
Microcystis_aruginosa_mycG	SLGIGG
Mycobacterium_leprae_pksE	SFGLSG
Mycobacterium_tuberculosis_mas	SFGMSG
Mycobacterium_tuberculosis_ppsC	SFGVSG
Mycobacterium_tuberculosis_ppsD	SFGLSG
Mycobacterium_tuberculosis_ppsE	SFGVGG
Myxococcus_xanthus_tal_mod2	AFGLGG
Nostoc_sp_nosB	SFGIGG
Pseudomonas_syringae_cfa6	SFGVSG
Pseudomonas_syringae_cfa7	SFGFSG
Sorangium_cellulosum_epoA_mod1	SFGMSG
Sorangium_cellulosum_epoC_mod2	SFGIGG
Sorangium_cellulosum_epoF_mod9	AFGLSG
Stigmatella_aurantiaca_mxaB2_B1	SFGLSG
Stigmatella_aurantiaca_mxaC1	SFGISG
Stigmatella_aurantiaca_mxaC2	SFGISG
Stigmatella_aurantiaca_mxaC3	AFGLSG
Stigmatella_aurantiaca_mxaD	AFGISG
Stigmatella_aurantiaca_mxaE	SFGISG
Stigmatella_aurantiaca_mxaF	SFGFGG
Streptomyces_natalensis_PimS0	SFGLTG
Streptomyces_natalensis_PimS2_mod3	SFGISG
Streptomyces_natalensis_PimS2_mod4	SFGLSG
S_viridochromogenes_aviM	GFGYGG
Xanthomonas_albilineans_xabB_mod2	SFGFGG
Xanthomonas_albilineans_xabB_mod3	AFGAGG

Bf, *Botrytis fuckeliana*; Ch, *Cochliobolus heterostrophus*; Gm, *Gibberella moniliformis*; Gz, *Gibberella zeae*; Nc, *Neurospora crassa*.

Institution: Albert R. Mann Library [Sign In as Member / Individual](#)[Full Text of this Article](#)**Table 3. Type I PKS genes annotated from the genomes of eight ascomycete fungi**

Fungus	PKS gene*	Domain structure of the PKS protein†	
<i>Schizosaccharomyces pombe</i>	None		
Schizosaccharomycetes;			
Schizosaccharomycetales;			
Schizosaccharomycetaceae			
<i>Saccharomyces cerevisiae</i>	None		
Saccharomycotina;			
Saccharomycetes;			
Saccharomycetales;			
Saccharomycetaceae			
<i>Eremothecium (Ashbya) gossypii</i>	None		
Saccharomycotina;			
Saccharomycetes;			
Saccharomycetales;			
Eremotheciaceae			
<i>Botryotinia fuckeliana</i>	(reducing PKS clade I)	(KS-AT-DH-(ME)-ER-KR-PP)	
(<i>Botrytis cinerea</i>)	<i>BfPKS1</i> - [†] M58U	KS-AT-DH-ME -ER-KR-PP	
Pezizomycotina;	<i>BfPKS2</i> -M37H	KS-AT-DH-ME -ER-KR-PP	

Leotiomyces; Helotiales; Sclerotiniaceae	(reducing PKS clade II) <i>BfPKS3</i> -M4H5 <i>BfPKS4</i> -M4CL	(KS-AT-DH-(ME) -KR-PP- -(CON)-(AMP-PP) KS-AT-DH -KR-PP -CON-AMP-PP KS-AT-DH-ME -KR-PP	
	<i>BfPKS5</i> -M40A	KS-AT-DH-ME -KR-PP -CON-AMP-PP	
	<i>BfPKS6</i> -M39G	KS-AT-DH-ME -KR-PP	
	** <i>BfPKS7</i> -M2TQ	§ KS-AT-DH-ME -KR-PP -CON-AMP-PP	
	(reducing PKS clade III) <i>BfPKS8</i> -M58P	(KS-AT-DH -ER-KR-PP-(PP)) KS-AT-DH -ER-KR-PP-PP	
	(reducing PKS clade IV) <i>BfPKS9</i> -M4RN	(KS-AT-DH-(ME)-ER-KR-PP) KS-AT-DH -ER-KR-PP	
	<i>BfPKS10</i> -M5VH	KS-AT-DH -ER-KR-PP	
	<i>BfPKS11</i> -M48U	KS-AT-DH-ME -ER-KR-PP	
	(nonreducing PKS clade II) <i>BfPKS12</i> -M5QUb	(KS-AT -PP-(PP) -CYC) KS-AT -PP-PP -CYC	
	<i>BfPKS13</i> -M2WM	KS-AT -PP-PP -CYC	
	(nonreducing PKSs basal to clades I and II) <i>BfPKS14</i> -M5DE	(KS-AT -PP-(PP) -(CYC)) KS-AT -PP-PP	
	<i>BfPKS15</i> -M304	KS-AT -PP-PP -CYC	
	(nonreducing PKS clade III) <i>BfPKS16</i> -M3R8	(KS-AT -PP-(PP)-ME-(CYC)) KS-AT -PP-PP -ME	
	<i>BfPKS17</i> -M3N0	KS-AT -PP -ME	
	<i>BfPKS18</i> -M5QUa	KS-AT -PP -ME	
	<i>BfPKS19</i> -M34F	KS-AT- -PP -ME	
	<i>BfPKS20</i> -M5PH	KS-AT- -PP-PP -ME-CYC	
<i>Cochliobolus heterostrophus</i>	(reducing PKS clade I) <i>ChPKS1</i> - [‡] M4FP	(KS-AT-DH-(ME)-ER-KR-PP) KS-AT-DH -ER-KR-PP	

<i>(Bipolaris maydis)</i>	<i>ChPKS3-M3FUa</i>	KS-AT-DH-ME-ER-KR-PP	
Pezizomycotina;	<i>ChPKS4-M3FUb</i>	KS-AT-DH-ME-ER-KR-PP	
Dothideomycetes;	<i>ChPKS5-M4EG</i>	KS-AT-DH-ME-ER-KR-PP	
Pleosporales;	<i>ChPKS6-M578</i>	KS-AT-DH-ME-ER-KR-PP	
Pleosporaceae	<i>ChPKS7-M584</i>	KS-AT-DH -ER-KR-PP	
	<i>ChPKS8-M4VU</i>	KS-AT-DH-ME-ER-KR-PP	
	<i>ChPKS9-M4U9</i>	KS-AT-DH -ER-KR-PP	
	<i>ChPKS10-M43G</i> (reducing PKS clade II)	KS-AT-DH -ER-KR-PP (KS-AT-DH-(ME) -KR-PP -(CON)-(AMP-PP))	
	<i>ChPKS16-M4GJ</i>	KS-AT-DH-ME -KR-PP	
	<i>ChPKS17-M3HR</i> (reducing PKS clade III)	KS-AT-DH-ME -KR-PP -CON (KS-AT-DH -ER-KR-PP-(PP))	
	<i>ChPKS2-M4LU</i>	KS-AT-DH -ER-KR-PP-PP	
	(reducing PKS clade IV)	(KS-AT-DH-(ME)-ER-KR-PP)	
	<i>ChPKS11-M55H</i>	KS-AT-DH-ME-ER-KR-PP	
	<i>ChPKS12-M3U6</i>	KS-AT-DH-ME-ER-KR-PP	
	[¶] <i>ChPKS13-M3SM</i>	KS-AT-DH-ME-ER-KR-PP	
	<i>ChPKS14-M3SV</i>	KS-AT-DH-ME-ER-KR-PP	
	<i>ChPKS15-M356b</i>	KS-AT-DH-ME-ER-KR-PP	
	(nonreducing PKS clade II)	(KS-AT -PP-(PP) -CYC)	
	<i>ChPKS18-M5CW</i>	KS-AT -PP -CYC	
	(nonreducing PKSs basal to clades I and II)	(KS-AT -PP-(PP) -(CYC)) KS-AT -PP	
	<i>ChPKS19-M4H3</i>		
	<i>ChPKS20-M5KW</i>	KS-AT -PP-PP	
	(nonreducing PKS clade III)	(KS-AT -PP-(PP) -ME-(CYC))	
	<i>ChPKS21-M3FU</i>	KS-AT -PP -ME	

	<i>ChPKS22-M3QM</i>	KS-AT -PP-PP-PP-ME	
	<i>ChPKS23-M4MK</i>	KS-AT -PP-PP -ME	
	(fungal NRPS/PKS clade) <i>ChPKS24-M356a</i>	(AMP-PP-KS-AT-DH -KR-PP -DHG) AMP-PP-KS-AT-DH -KR-PP -DHG	
	(fungal 6MSAS clade) <i>ChPKS25-M3HB</i>	(KS-AT-DH -KR-PP) KS-AT-DH -KR-PP	
<i>Gibberella moniliformis</i> <i>(Fusarium verticillioides)</i>	(reducing PKS clade I) <i>GmPKS2-[†]MOWT</i>	(KS-AT-DH-(ME)-ER-KR-PP) KS-AT-DH-ME-ER-KR-PP	
Pezizomycotina;	<i>GmPKS5-MOQ4</i>	KS-AT-DH-ME-ER-KR-PP	
Sordariomycetes;	<i>GmPKS6-M232b</i>	KS-AT-DH -ER-KR-PP	
Hypocreales;	<i>GmPKS7-M1NB</i>	KS-AT-DH -ER-KR-PP	
Nectriaceae	(reducing PKS clade II) <i>GmPKS1-M232a</i>	(KS-AT-DH-(ME) -KR-PP -(CON)-(AMP-PP)) KS-AT-DH-ME -KR-PP -CON-AMP-PP	
	<i>GmPKS8-MODGb</i>	KS-AT-DH-ME -KR-PP	
	<i>GmPKS9-M2SJ</i>	KS-AT-DH-ME -KR-PP -CON-AMP-PP	
	<i>GmPKS10-M3ED</i>	KS-AT-DH-ME -KR-PP -CON-AMP-PP	
	(reducing PKS clade IV) <i>GmPKS11-M2VE</i>	(KS-AT-DH-(ME) -ER-KR-PP) KS-AT-DH-ME-ER-KR -PP	
	<i>GmPKS12-MOUB</i>	KS-AT-DH-ME-ER-KR -PP	
	<i>GmPKS13-MODGa</i>	KS-AT-DH-ME-ER-KR -PP	
	<i>GmPKS14-M2DH</i>	KS-AT-DH -ER-KR -PP	
	<i>GmPKS15-M2KJ</i>	KS-AT-DH-ME-ER-KR -PP	
	(nonreducing PKS clade I) <i>GmPKS3-MOS2</i>	(KS-AT -PP-(PP) -CYC) KS-AT -PP-PP	
	<i>GmPKS4-MYKP</i>	KS-AT PP -CYC	

<i>Gibberella zeae</i> <i>(Fusarium graminearum)</i>	(reducing clade I) <i>GzPKS1</i> - ₉₄₉₀₊₁₈₂₆₅	(KS-AT-DH-(ME) -ER-KR-PP) § KS-AT-DH-ME-ER-KR-PP	
Pezizomycotina;	<i>GzPKS2</i> -17858	§ KS-AT-DH-ME-ER-KR-PP	
Sordariomycetes;	<i>GzPKS4</i> -10828	§ KS-AT-DH -ER-KR-PP	
Hypocreales;	<i>GzPKS5</i> -11130	§ KS-AT-DH-ME-ER-KR-PP	
Nectriaceae;	<i>GzPKS7</i> -8578	§ KS-AT-DH-ME -ER-KR-PP	
	** <i>GzPKS8</i> -12692	§ KS-AT-DH-ME-ER-KR-PP	
	(reducing PKS clade II) <i>GzPKS9</i> -5148	(KS-AT-DH-(ME) -KR-PP -(CON)-(AMP-PP)) § KS-AT-DH -KR-PP	
	<i>GzPKS10</i> -7224	§ KS-AT-DH -KR-PP -CON-AMP-PP	
	(reducing PKS clade IV) ** <i>GzPKS6</i> -4447 <i>GzPKS11</i> -3270	(KS-AT-DH-(ME) -ER-KR-PP) § KS-AT-DH-ME-ER-KR-PP § KS-AT-DH-ME-ER-KR-PP	
	(nonreducing PKS clade I) <i>GzPKS3</i> -8340+6005 <i>GzPKS12</i> -8340+6550	(KS-AT -PP-(PP) -CYC) § KS-AT -PP -CYC § KS-AT -PP-TE	
	(nonreducing PKSs basal to clades I and II) <i>GzPKS13</i> -7923	(KS-AT -PP-(PP) -(CYC)) § KS-AT -PP	
	<i>GzPKS14</i> -15081+19966	§ KS-AT -PP	
	(nonreducing PKS clade III)	(KS-AT -PP-(PP)-ME -(CYC))	
	<i>GzPKS15</i> -13158	§ KS-AT -PP	
	** <i>GzPKS16</i> -18966	§ KS-AT -PP -ME	

<i>Neurospora crassa</i>	(reducing PKS clade I)	(KS-AT-DH-(ME) -ER-KR-PP)	
Pezizomycotina;	<i>NcPKS3</i> -NCU04865.1	KS-AT-DH-ME-ER-KR-PP	
Sordariomycetes;	(reducing PKS clade II)	(KS-AT-DH-(ME) -KR-PP -(CON)-(AMP-PP))	
Sordariales;	<i>NcPKS4</i> -NCU08399.1	KS-AT-DH-ME-ER-KR-PP	
Sordariaceae;	(reducing PKS clade III)	(KS-AT-DH -ER-KR-PP-(PP))	
	<i>NcPKS1</i> -NCU06013.1	KS-AT-DH -ER-KR-PP	
	<i>NcPKS2</i> -NCU05011.1	KS-AT-DH -ER-KR-PP	
	(reducing PKS clade IV)	(KS-AT-DH-(ME) -ER-KR-PP)	
	<i>NcPKS5</i> -NCU09638.1	KS-AT-DH -ER-KR-PP	
	<i>NcPKS6</i> -NCU02918.1	KS-AT-DH-ME-ER-KR-PP	
	(nonreducing PKS clade II)	(KS-AT -PP-(PP) -CYC)	
	<i>NcPKS7</i> -NCU03584.1	KS-AT- PP-PP -CYC	

*Genes within a species are listed by their subclades as they appear in the protein genealogy in Fig. 1.

†KS, ketoacyl synthetase [PKS-associated domain pfam00109.6 (N-terminal) + pfam2801.6 (C-terminal)]; AT, acyl transferase (PKS-associated domain pfam 00698.6); DH, dehydratase (PKS-associated domain, C terminus of COG3321.1, which includes KS and AT domains); ER, enoyl reductase (PKS-associated domain ~COG0604.1); KR, ketoreductase (PKS-associated domain ~COG1028.1); PP, acyl carrier protein of PKSs and thiolation of NPSs (attachment site of phosphopantetheine, pfam 00550.6); ME, methyltransferase (PKS-associated domain COG2226.1); CYC, Claisen-type cyclase; AMP, adenylation (NPS-associated domain COG1020.1); CON, condensation (NPS-associated domain pfam 00668.6); DHG, dehydrogenase (NPS-associated domain COG3320.1).

‡Celera/TMRI assembly metacontig numbers from 5× coverage of *B. fuckeliana*, *C. heterostrophus*, and *G. moniliformis* genomes.

§Modular construction is predicted, based on placement of the gene fragment in the protein genealogy; prediction is identical to its orthologs when they occur or includes all possible domains for those genes without an orthologous match in the genealogy.

¶Pseudogene with domain residues punctuated by stop codons.

||TMRI assembly contig numbers from 2× coverage of the *G. zeae* genome.

**Gene drawn onto gene genealogy and marked with a single asterisk in Fig. 1, as available sequence is a C-terminal fragment of the KS domain (alignment used for phylogenetic analysis includes only N-terminal fragments).

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