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(54) **ENGINEERING THE PRODUCTION OF A CONFORMATIONAL VARIANT OF OCCIDIOFUNGIN THAT HAS ENHANCED INHIBITORY ACTIVITY AGAINST FUNGAL SPECIES**

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(51) **Int. Cl.**

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**A61K 38/12** (2006.01)  
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**A01N 63/00** (2006.01)  
**A01N 63/02** (2006.01)  
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**C07K 7/56** (2006.01)

(52) **U.S. Cl.**

CPC ..... **C07K 7/54** (2013.01); **A01N 43/713** (2013.01); **A01N 63/00** (2013.01); **A01N 63/02** (2013.01); **A61K 38/12** (2013.01); **C07K 7/56** (2013.01); **C12N 9/16** (2013.01); **C12Y 301/02** (2013.01)

(58) **Field of Classification Search**

CPC ..... **C07K 7/54**; **A01N 43/713**; **A01N 63/00**; **A61K 38/12**

See application file for complete search history.

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(57) **ABSTRACT**

Occidiofungin is a cyclic nonribosomally synthesized antifungal peptide with submicromolar activity. This invention is directed to compositions enriched for particular occidiofungin diastereomers/conformers, methods of making compositions enriched for particular diastereomers/conformers and microorganisms suitable for producing enriched compositions of particular diastereomers/conformers. Methods of treating fungal infections or plants infected by fungi are also provided.

**1 Claim, 14 Drawing Sheets**  
**(4 of 14 Drawing Sheet(s) Filed in Color)**

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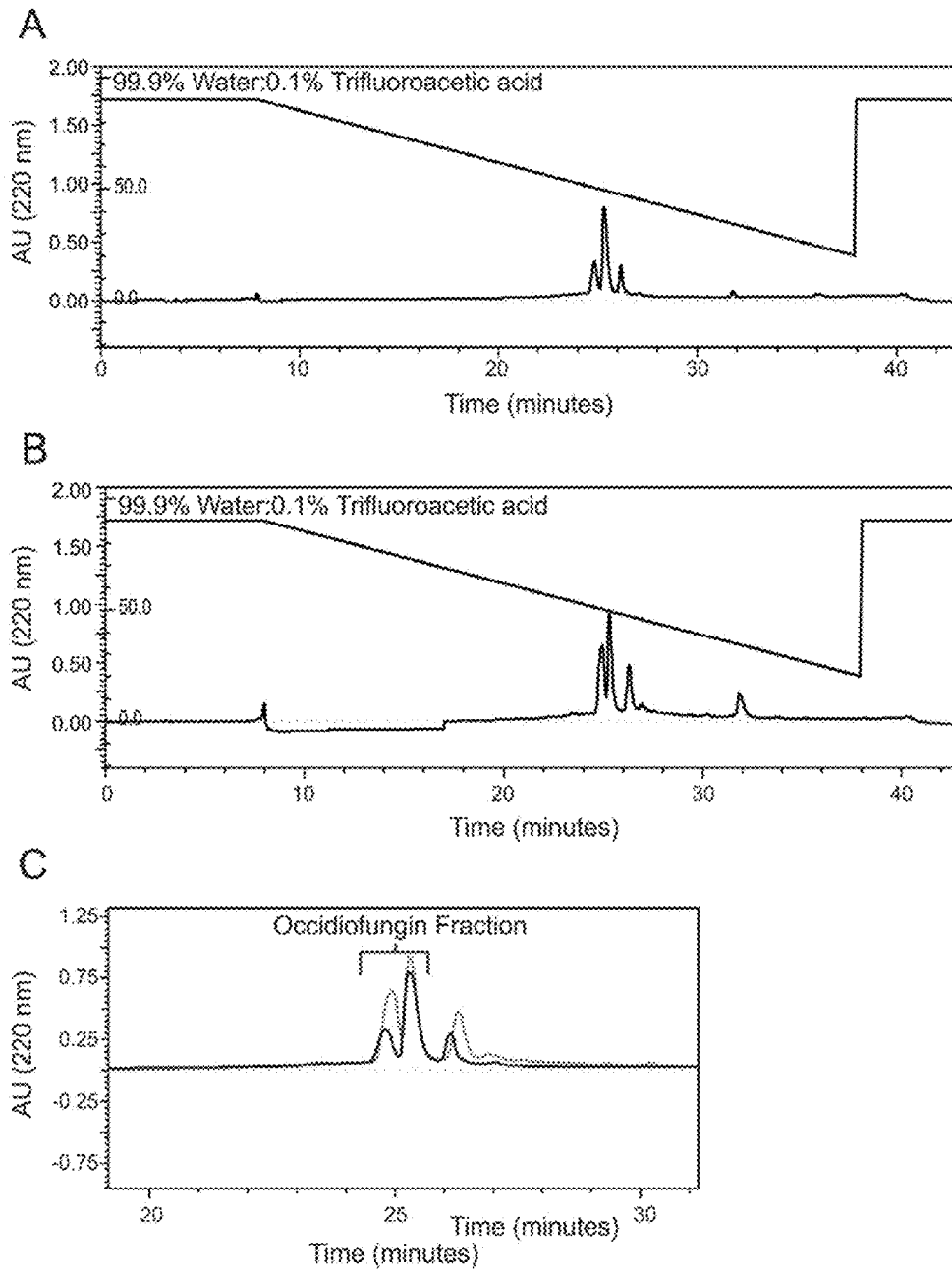


FIGURE 2

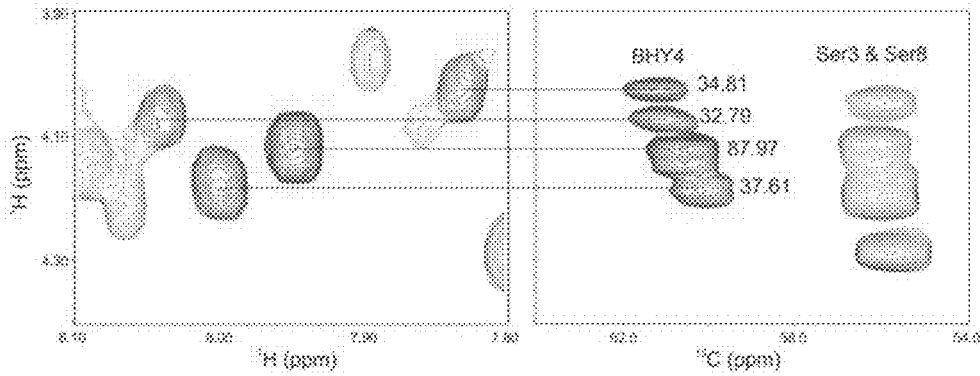


FIGURE 3

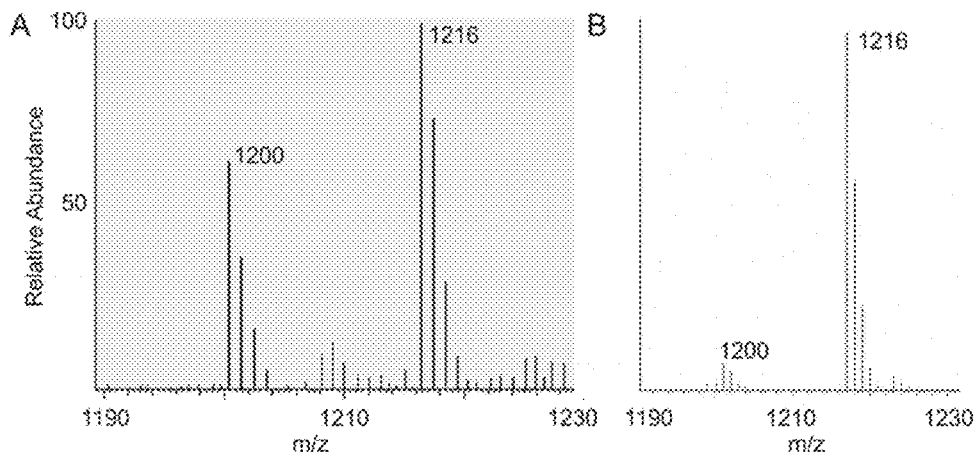


FIGURE 4



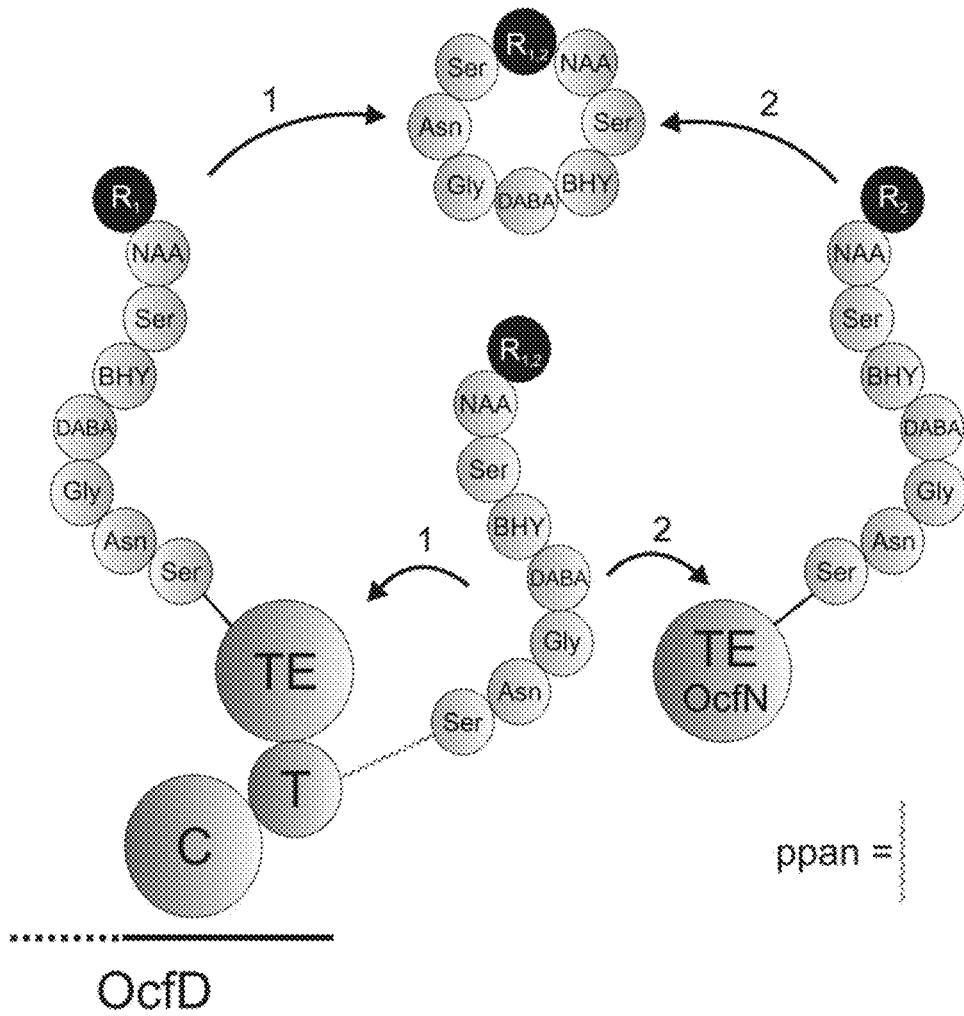


FIGURE 6

A

Occidiofungin MICs

Isolate	Wild-Type MIC (µg/mL)	<i>ocfN</i> mutant MIC (µg/mL)
<i>Candida albicans</i> 66027	1.0	1.0
<i>Candida albicans</i> LL	0.5	1.0
<i>Candida albicans</i> TE	0.5	1.0
<i>Candida glabrata</i> 66032	0.5	1.0
<i>Candida parapsilosis</i> 90018	1.0	2.0
<i>Candida tropicalis</i> 66029	0.5	1.0

B

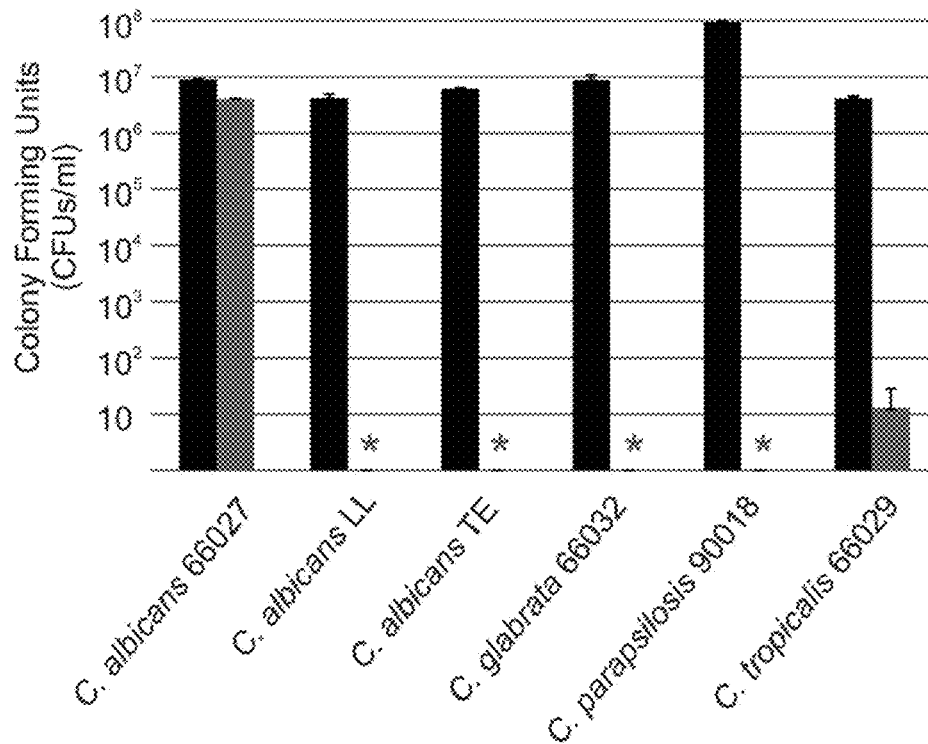


FIGURE 7



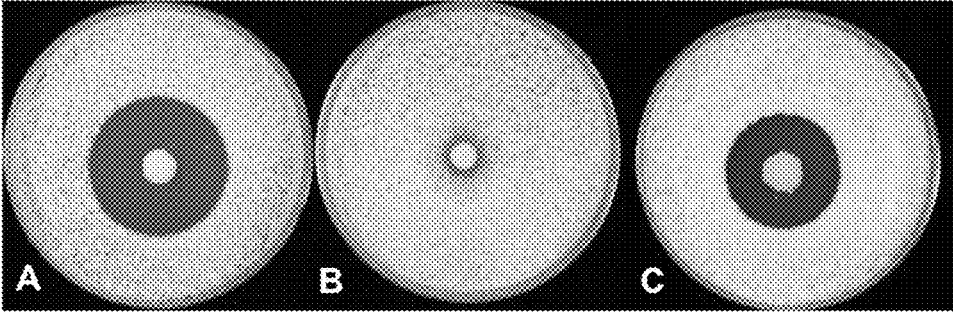
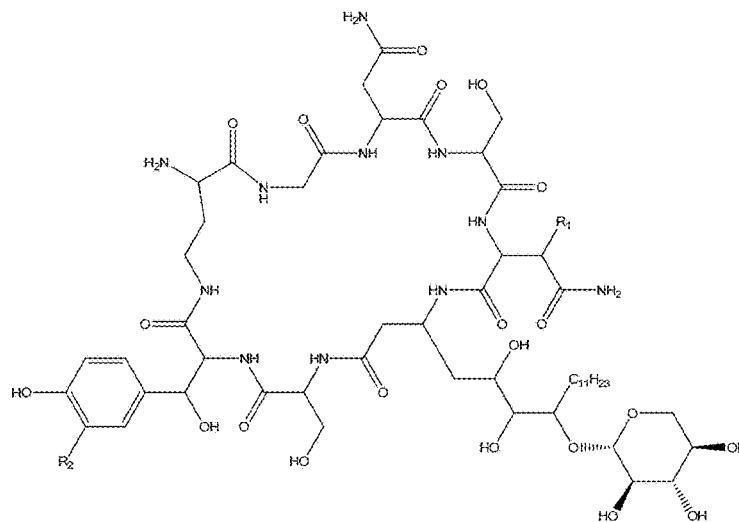
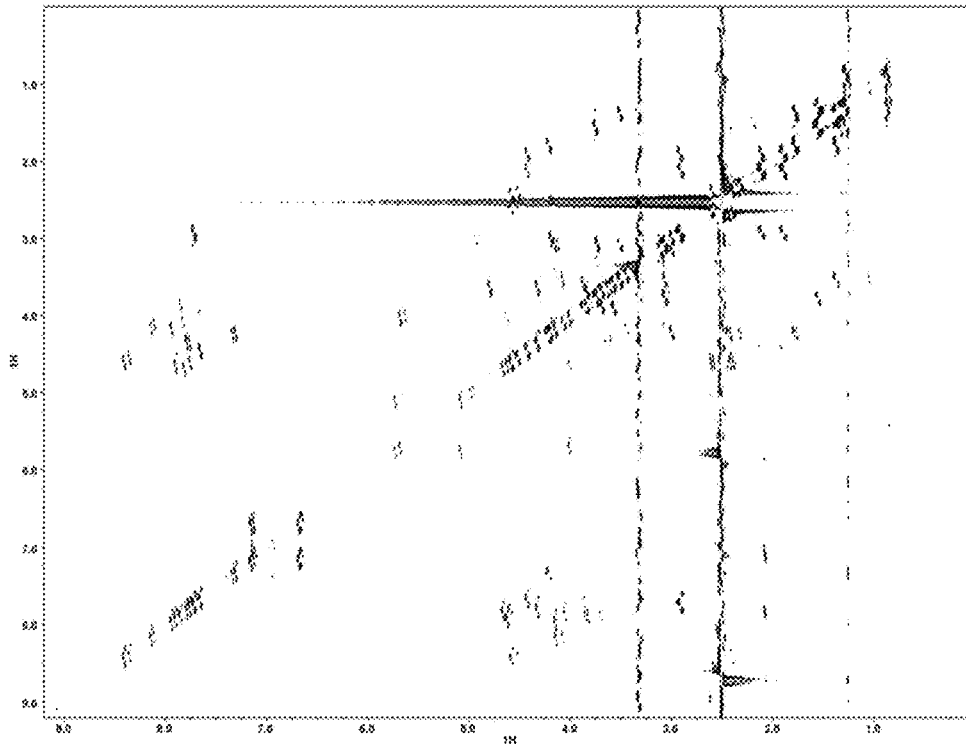
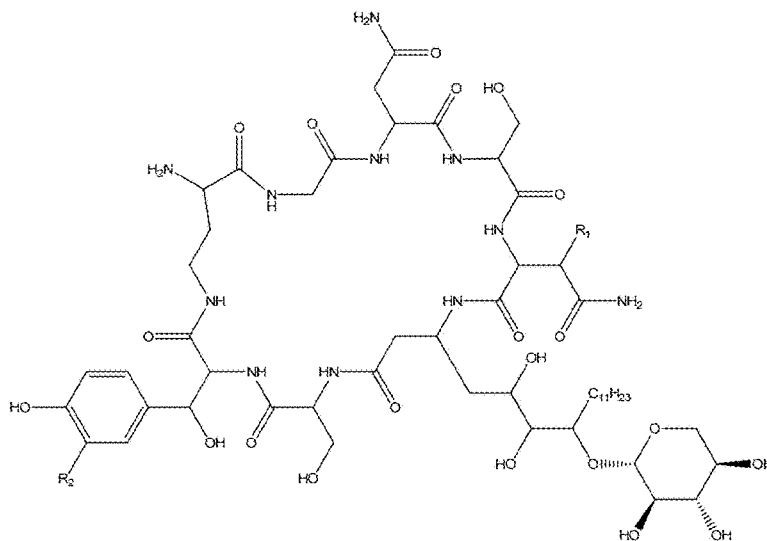
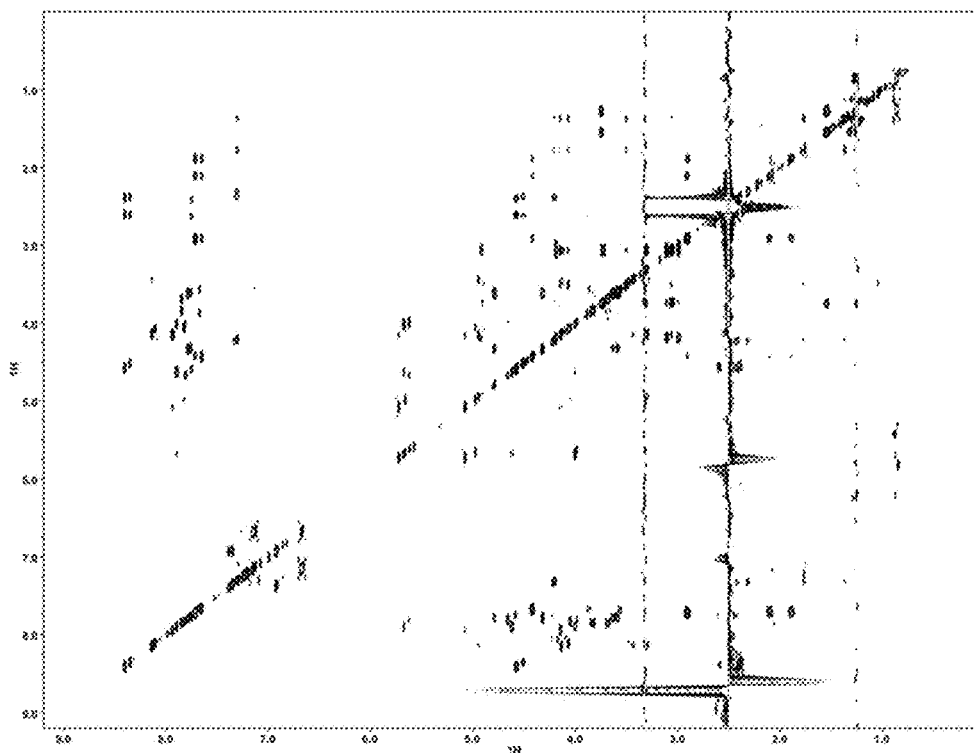


FIGURE 8



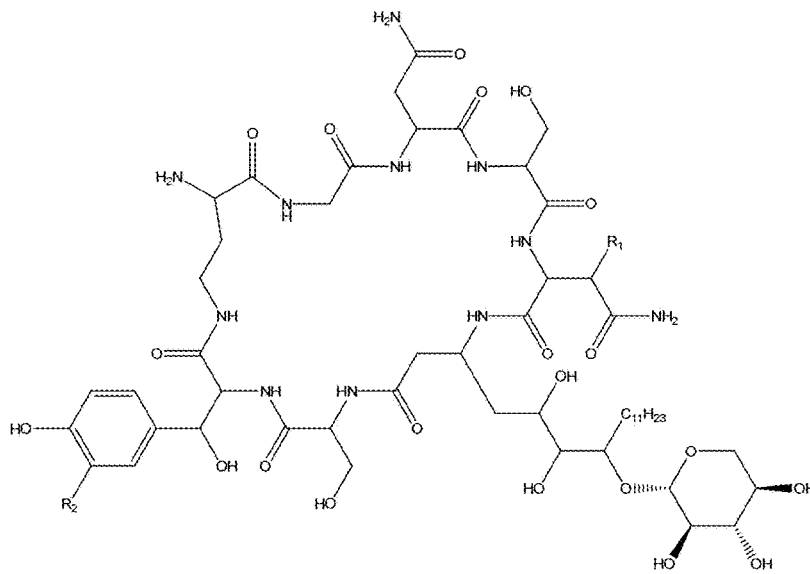
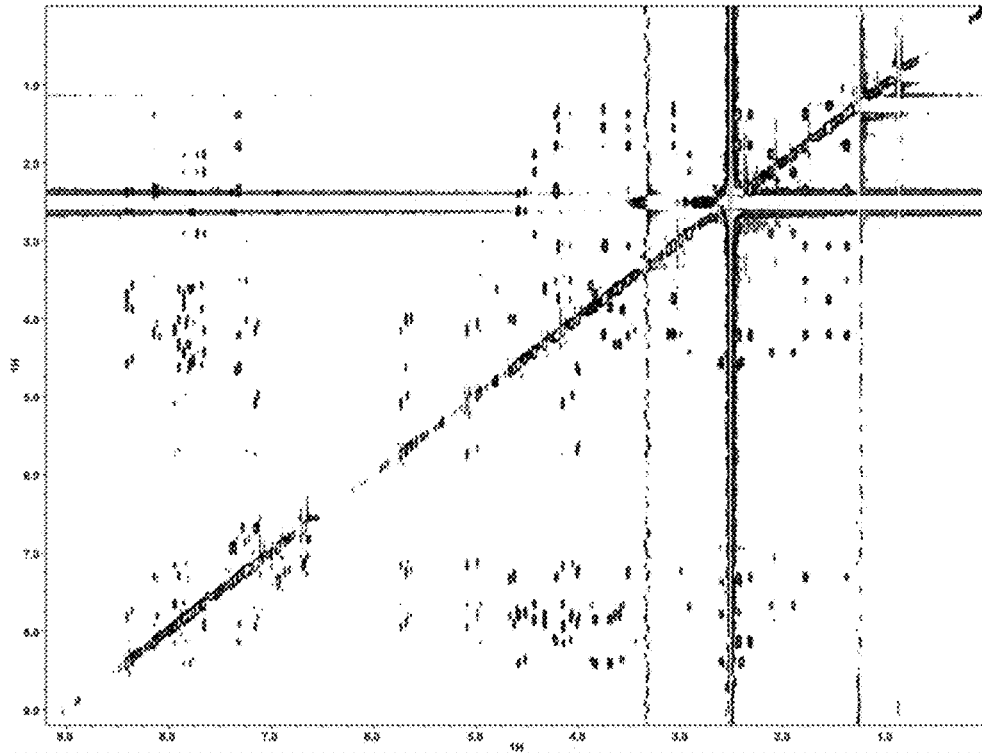
Occidiofungin: R1 (-H or -OH); R2 (-H or -Cl)

FIGURE 9



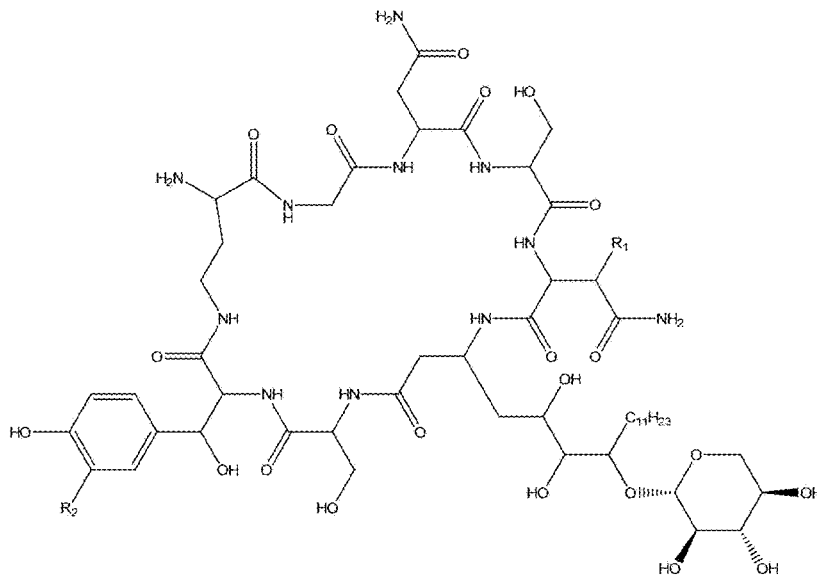
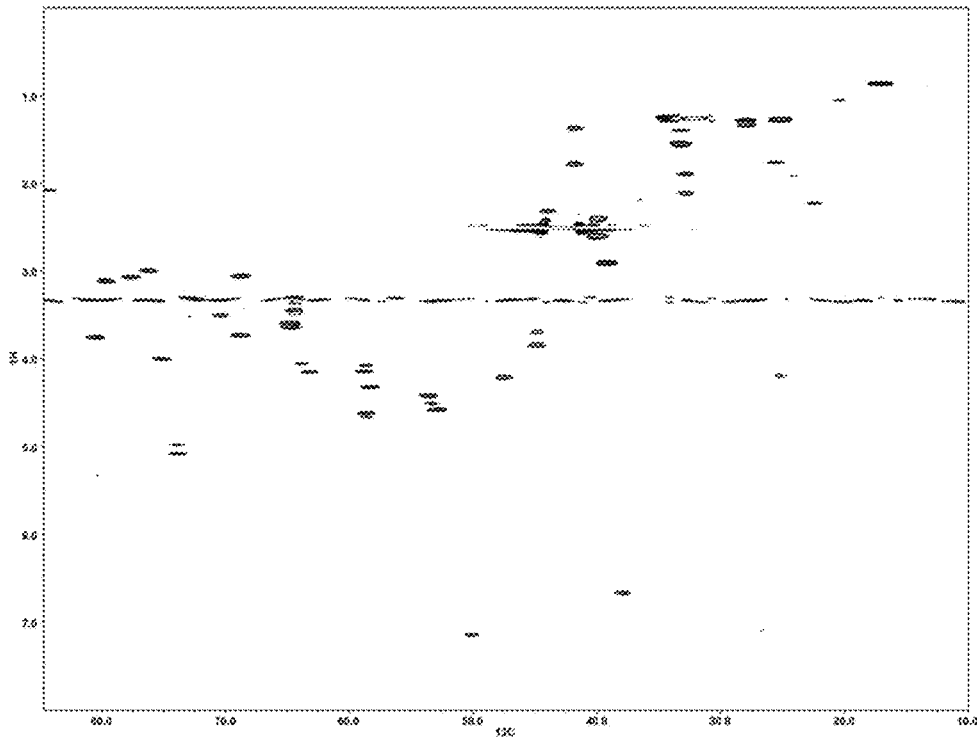
Occidiofungin: (R<sub>1</sub>, -H or -OH); (R<sub>2</sub>, -H or -Cl)

FIGURE 10



Occidiofungin: (R1,-H or -OH); (R2,-H or -Cl)

FIGURE 11



Occidiofungin: (R<sub>1</sub>, -H or -OH); (R<sub>2</sub>, -H or -Cl)

FIGURE 12

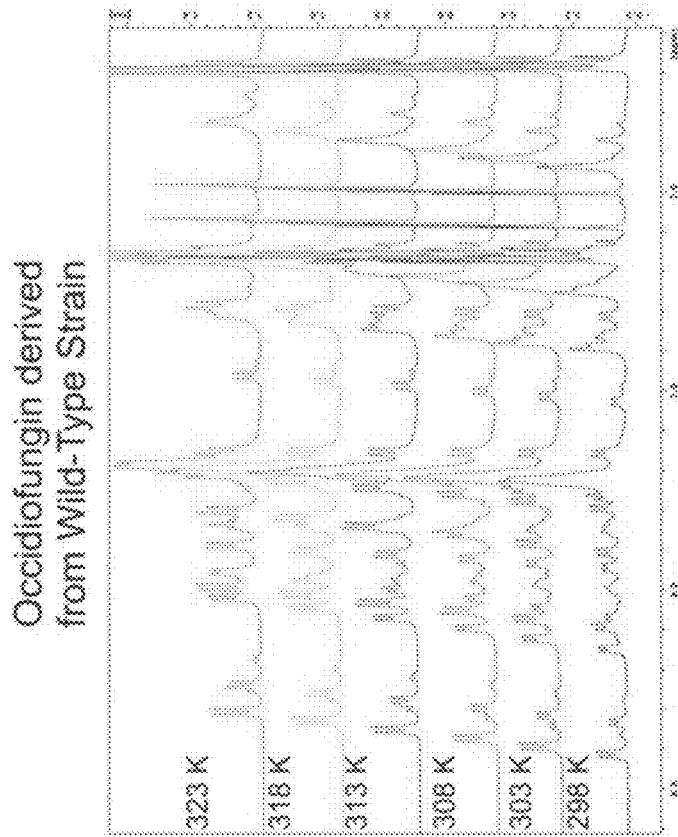


FIGURE 13B

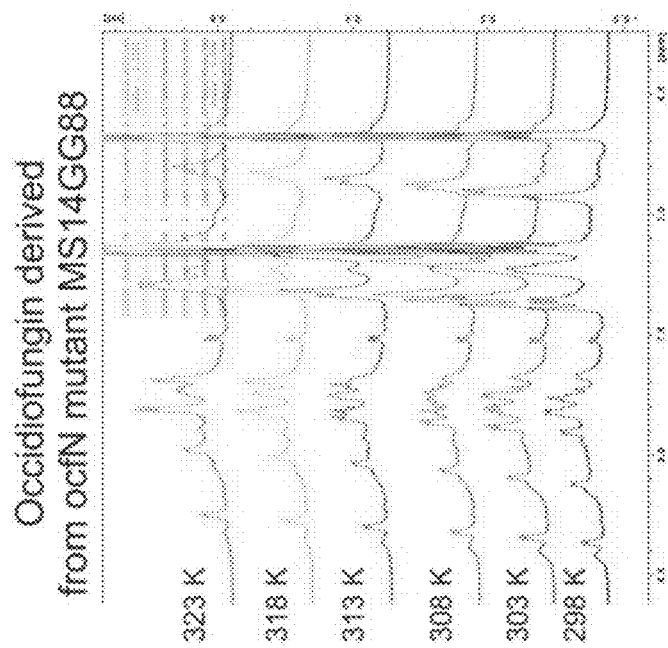


FIGURE 13A

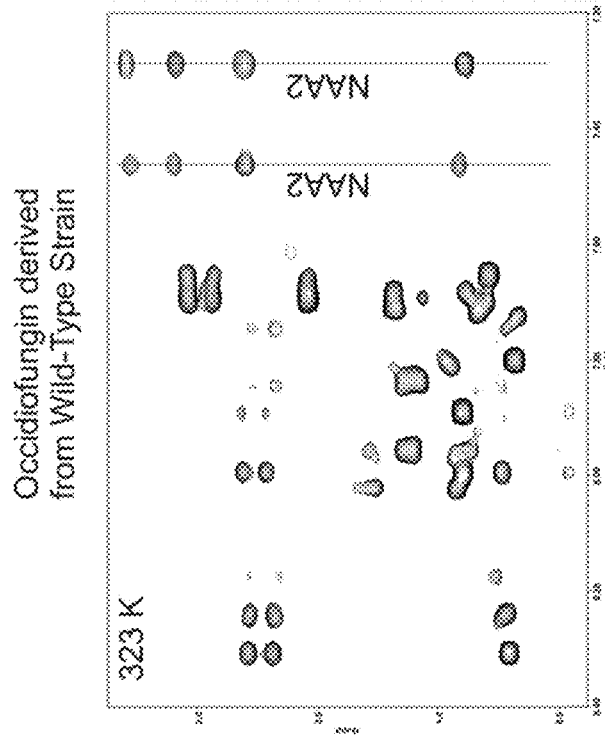


FIGURE 14B

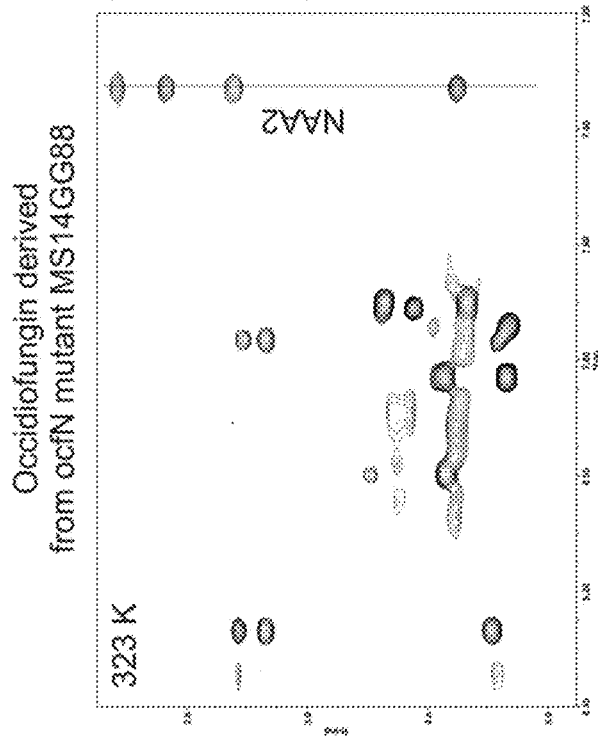


FIGURE 14A

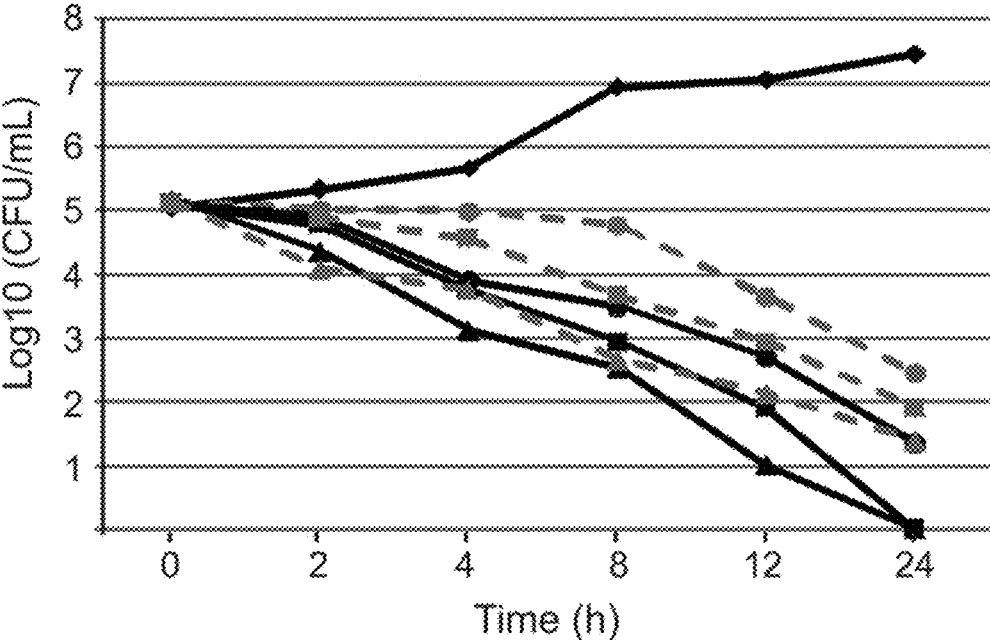


FIGURE 15



1

**ENGINEERING THE PRODUCTION OF A  
CONFORMATIONAL VARIANT OF  
OCCIDIOFUNGIN THAT HAS ENHANCED  
INHIBITORY ACTIVITY AGAINST FUNGAL  
SPECIES**

CROSS-REFERENCE TO RELATED  
APPLICATION

This application claims the benefit of U.S. Provisional Application Ser. No. 61/731,105, filed Nov. 29, 2012, the disclosure of which is hereby incorporated by reference in its entirety, including all figures, tables and amino acid or nucleic acid sequences.

The Sequence Listing for this application is labeled "Seq-List.txt" which was created on Nov. 26, 2013 and is 264 KB. The entire contents of the sequence listing is incorporated herein by reference in its entirety.

This invention was made with government support under 0204332 awarded by the National Institute of Food and Agriculture, USDA. The government has certain rights in the invention.

BACKGROUND OF THE INVENTION

Nonribosomal peptide synthetases (NRPSs) produce a wide array of small and structurally complex peptides that have therapeutic potential. The system enables the incorporation of nonproteinogenic amino acids into the polypeptide. Polyketide synthetases (PKSs) are a family of enzymes or enzyme complexes that produce polyketides. Integration of PKSs into the NRPSs system further increases the variety of polypeptides that can be produced by these systems. Recent studies are aimed at exploiting NRPSs for producing peptide libraries that can be screened for therapeutic applications.<sup>1-9</sup>

Unlike linear peptides, cyclic peptides are restrained to fewer conformations that facilitate their interaction with their molecular target.<sup>10-18</sup> These structural constraints provide resistance to proteases, extreme pH, and temperature.<sup>10, 19</sup> These attributes make them one of the most promising scaffolds for pharmacophores. Synthetic design of cyclic peptides is hindered by regioselectivity.

Classical total synthesis of peptides by solid phase or solution phase peptide synthesis followed by subsequent cyclization reactions requires the addition and removal of protecting groups at the right stages to drive the cyclization among the correct residues.<sup>8</sup> Even with these considerations, proper cyclization is hindered by intermolecular interactions and entropically disfavoured pre-cyclization conformations resulting in a vast mixture of compounds or low yields. Microorganisms ensure the formation of a functional cyclic peptide conformation by enzymatically catalyzing the cyclization and release of the peptide with regioselectivity using a cyclase thioesterase.<sup>1, 7</sup> The cyclase thioesterase is often located at the C-terminal end of the last NRPS involved in the synthesis of the peptide and is referred to as the TE (Thioesterase) domain.

The TE domain can hydrolyze the bound peptide as a linear peptide or it can catalyze an intramolecular reaction resulting in the formation of a cyclic peptide. At present, very little is known about the cyclization mechanism of peptides. The crystal structure of the surfactin peptide cyclase provided the first basic understanding of its mechanism of action.<sup>20, 21</sup> The peptidyl chain bound to 4-phosphopantetheine cofactor (ppan) that is attached to the thiolation (T)-domain is transferred to a serine in the adjacent TE domain. Ser80 is part of a catalytic triad of residues (His

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207 and Asp107) in the surfactin cyclase. His207 and Asp107 activate the Ser80, facilitating the transfer of the peptidyl chain to the TE domain. Once the peptide is transferred to the TE domain, the cyclase binding pocket enables proper orientation and cyclization of the peptide substrate. The enzyme was found to share structural homology to  $\alpha,\beta$ -hydrolase family. The lack of water in the binding cleft of the cyclase, which prevents hydrolysis, is the significant alteration from the hydrolase family that gives the cyclase thioesterase its ability to form cyclic peptides.

Occidiofungin is a broad spectrum nonribosomally synthesized cyclic antifungal peptide that has submicro/nanomolar activity and low toxicity.<sup>19, 22-26</sup> An interesting feature in occidiofungin's biosynthetic pathway is the presence of two putative thioesterases. One is present as an independently expressed thioesterase, OcfN, and the other is a C-terminal TE domain of OcfD. There remains a need for the production of anti-fungal agents that have increased cidal activity against various fungi.

BRIEF SUMMARY OF THE INVENTION

This invention relates to antifungal compounds and their therapeutic use in the prevention or treatment of fungal infections and diseases. Particularly, various aspects of the invention provide compositions enriched for occidiofungin diastereomers/conformers that have higher activity against fungal infections or diseases (in mammals or plants).

Other aspects of the invention provide for compositions enriched for particular diastereomers/conformers produced by genetic modification of occidiofungin producing microorganisms such that the production of a particular occidiofungin diastereomer/conformer is favored. Thus, the invention relates to methods of making such occidiofungin diastereomers/conformers, compositions enriched for such diastereomers/conformers and methods of using compositions comprising occidiofungin diastereomers/conformers disclosed herein as fungicides for animals and plants. The invention further relates to the microorganisms that produce compositions enriched for occidiofungin enriched for occidiofungin diastereomers/conformers corresponding to diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations). Methods of increasing the production of occidiofungin diastereomers/conformers corresponding to diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations) in microorganisms and production systems are also provided.

As discussed above, one aspect of the invention provides compositions enriched for occidiofungin diastereomers/conformers, in particular the occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations). Thus, the invention provides compositions enriched for such antifungal diastereomers/conformers for treating fungal infection. In certain embodiments of this aspect of the invention, pharmaceutical and agricultural compositions that contain a composition enriched for diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations) are provided. Compositions enriched for a particular occidiofungin diastereomer/conformer can also be produced by the genetically modified microorganisms discussed below (e.g., microorganisms in which the function of ocfD and/or ocfN has been altered in order to favor the production of a particular diastereomer/conformer).

Novel antifungals are needed because of the importance of fungal infections in immunocompromised patients, and the limitations of currently-available antifungal agents regarding their spectra of activity and toxicities. In addition, new antifungals are crucial for food preservation and production of a sufficient and affordable food supply. In this context, this application relates to the disclosure of a composition enriched for occidiofungin diastereomers/conformers having increased antifungal activity as compared to occidiofungin compositions produced by *Burkholderia contaminans* MS14 (disclosed in U.S. Patent Application Publication 2011/0136729, the disclosure of which is hereby incorporated by reference in its entirety). Diastereomers/conformers have been characterized by a number of techniques, including COSY, TOCSY, NOESY, ROESY, and HSQC 2D NMR spectroscopy experiments.

The antifungal activity of the disclosed occidiofungin diastereomers/conformers (diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations)) provides for compositions having greater antifungal activity as compared to as compared to occidiofungin compositions produced by *Burkholderia contaminans* MS 14 when cultured under the same conditions.

The phrase “enriched for the disclosed occidiofungin diastereomers/conformers” is intended to convey that the a composition contains the disclosed occidiofungin diastereomers/conformers (diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations)) in an amount higher/greater than that produced by *Burkholderia contaminans* MS14 as disclosed in the examples provided herein (in which approximately 36% of total amount of occidiofungin corresponds to occidiofungin diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations)). Thus, the phrase indicates that at least 37% of the total amount of occidiofungin diastereomers/conformers present within an enriched composition are the disclosed diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations)). In various embodiments, compositions “enriched for the disclosed occidiofungin diastereomers/conformers” contain at least 40%, 41%, 42%, 43%, 44%, 45%, 46%, 47%, 48%, 49%, 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, 60%, 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, 70%, 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, 79%, 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98% or 99% of the disclosed diastereomers/conformers in relation to the total amount of occidiofungin diastereomers/conformers in the composition.

The phrase “enriched for a particular occidiofungin diastereomers/conformer” is intended to convey that a composition contains the an occidiofungin diastereomer/conformer that is produced by a microorganism in which the activity of the ocfD and/or ocfN thioesterase has been altered such that the production of a particular conformer is favored.

#### BRIEF DESCRIPTION OF THE DRAWINGS

The patent or application file contains at least one drawing executed in color. Copies of this patent or patent application publication, with color drawing(s), will be provided by the Office upon request and payment of the necessary fee.

FIG. 1. Covalent structure of occidiofungin. R1 and R2 represent the locations where a hydroxyl or chlorine is added, respectively (R1, —H or —OH; R2, —H or —Cl).

FIGS. 2A-2C. RP-HPLC Chromatograms. A. Chromatogram of the final purification step of the wild-type occidiofungin fraction at 220 nm using a 4.6×250 mm C18 column. B. Chromatogram of the final purification step of ocfN mutant occidiofungin fraction at 220 nm using a 4.6×250 mm C18 column. C. Overlay of the wild-type (black) and the mutant (grey) fractions of occidiofungin.

FIG. 3. TOCSY (left panel) and HSQC (right panel) spectra of BHY4 in the wild-type sample. The proportions of Asn1 and BHN1 variants were determined by the measurement of the <sup>13</sup>C-HSQC Ha-Ca cross peak intensities of BHY4 in the HSQC spectra. These values are listed next to their corresponding peaks in the right panel. The peaks in red and green represent the BHY4 peaks associated with BHN1 and Asn1 variants, respectively. Based on the calculation of their relative proportions, i.e. (34.81+87.97 for the BHY4 peaks found in the BHN1 conformational variants) and (32.79+37.61 for the BHY4 peaks found in the Asn1 conformational variants), the approximate proportion of the Asn1 variants could be calculated as (32.79+37.61)/(34.81+87.97)+(32.79+37.61).

FIGS. 4A-4B. ESI mass spectrometry. A. EST mass spectrometry data of purified wild-type occidiofungin fraction. B. ESI mass spectrometry data of purified ocfN mutant occidiofungin fraction.

FIGS. 5A-5C. TOCSY fingerprint region (NH correlations). A. NH correlations in the wild-type sample. The two BHN1 and four Asn1 spin systems present in the wild-type sample are colored red. B. NH correlations in the ocfN mutant sample. C. Overlay of the NH correlations found in the wild-type and ocfN mutant samples. NH correlations that are not present in the ocfN mutant sample are colored green.

FIG. 6. Schematic of occidiofungin ring closure. The completely synthesized eight amino acid linear peptide is bound by a 4-phosphopantetheine cofactor (ppan) linker to the thiolation (T) domain. The peptide varies by the presence or absence of a hydroxyl group on the beta carbon of Asn1. The TE domain of OcfD is capable of forming the cyclic peptide of both variants in the absence of a functional OcfN cyclase thioesterase. However, it is not as efficient at producing the Asn1 cyclic peptide variant as OcfN. In the presence of a functional OcfN cyclase thioesterase, a novel diastereomers of occidiofungin is formed by the selective ring closure of the Asn1 cyclic peptide. R1 and R2 are BHN1 and Asn1, respectively.

FIGS. 7A-7B. Comparison of the bioactivity from the wild-type and ocfN mutant occidiofungin fractions. A. MICs of wild-type and ocfN mutant fraction determined by CLSI M27-A3 method in RPMI 1640. B. Comparison of the CFUs in the MIC wells of wild-type fraction to the corresponding well having the same concentration of the ocfN mutant occidiofungin fraction. Asterisks represent no detectable colonies in the MIC wells of the wild-type occidiofungin fraction. Black and grey bars are ocfN mutant and wild-type fractions, respectively. Standard deviations for the CFU measurements are presented.

FIGS. 8A-8C. Potato dextrose agar plates were inoculated with each of the strains and incubated for 3 days at 28° C. The plates were oversprayed with the indicator fungus *Geotrichum candidum* and incubated overnight. A: The wild-type strain MS14; B: Negative control MS14GG78 (ocfJ::nptII); C: MS14GG88 (ocfN::nptII).

FIG. 9. COSY60 NMR Spectrum of Occidiofungin from ocfN mutant MS14GG88 recorded at 600 MHz in DMSO-d<sub>6</sub>.

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FIG. 10. TOCSY60 NMR Spectrum of Occidiofungin from ocfN mutant MS14GG88 recorded at 600 MHz in DMSO-d<sub>6</sub>.

FIG. 11. NOESY400 NMR Spectrum of Occidiofungin from ocfN mutant MS14GG88 recorded at 600 MHz in DMSO-d<sub>6</sub>.

FIG. 12. <sup>13</sup>C-HSQC NMR Spectrum of Occidiofungin from ocfN mutant MS14GG88 recorded at 600 MHz in DMSO-d<sub>6</sub>.

FIGS. 13A-13B. One-dimensional NMR temperature titration curves for occidiofungin derived from ocfN mutant MS14GG88 (FIG. 13A) and wild-type strain MS14 (FIG. 13B).

FIGS. 14A-14B. TOCSY fingerprint region (NH correlations) for occidiofungin derived from ocfN mutant MS14GG88 (FIG. 14A) and wild-type strain MS14 (FIG. 14B) at 50° C.

FIG. 15. Time-kill experiments performed against *Candida glabrata* ATCC66032. Solid black lines and dashed grey lines correspond to samples treated with occidiofungin derived from wild-type strain MS14 and ocfN mutant MS14GG88, respectively. Circles, squares, and triangles

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represent samples treated with 0.5, 1.0, and 2.0 µg/mL of occidiofungin, respectively. The diamond represents the sample treated with the blank control.

#### BRIEF DESCRIPTION OF THE SEQUENCES

SEQ ID NOs: 1-2: PCR primer sequences

SEQ ID NO: 3: amino acid sequence for OcfN (thioesterase; thioesterase motif of -G-X-S-X-G- underlined (X is any amino acid))

```
MRLICFPYAGGSAAVYRTLQASLPGIEVCRHELAGRGRSRLSEPAVRDMA
TLVDTLCDLDDCFDRPFALLGHSMGAAIAAELALRLPAHARPNLRHLF
VSARAAPGKERHRRMQALDDRAFIDALREMGGTPKAVLDNSELMALL
MPALRADFTMIENHRPVPGPRLAVDITAFAGRADKEIPVDAVAGWGAAT
TGRFDHFVIEGDHFFLRNEMRTMAGIIAARMRPEHAASSALQA
```

SEQ ID NO: 4: amino acid sequence for OcfD (thioesterase motif of -G-X-S-X-G- underlined (X is any amino acid))

```
MQDNNVLVTD RESLSRVAGV YGIAAYAPSQ QPGRPLTRSV RLTPASLDLL
RRIGDGELAE FAVAAAGIAF LLWKYFRIPV TVLGTPLAG HPSARAAIVP
LIIEVRPDER IEDYLSRVAG IVEDSYAEPR PPLETLVRNE KDMALALQTK
VALADDRVHH APTGRDDDLQ LHLRLARGEI ELRYSGAIEP FIIDGFAGSL
AAVLEAFEHL DGAVGDIEAA PPEQGPLLAA FNETATAGPS HPTVAMFEA
QVARTPTAPA LVTDSLSMTY ADLNARANSL AHHLREHHGV GPESLVGIML
DRSEWMIVAI LGILKAGAAF VPLDPAYPAE RINHILGDTG LSLLVTSQSSQ
LAQWYEFSGV TLLLDQELPG WQPLPDNPPH RAEPALHAYV LYTSGSTGKP
KGCLLEHRNL AHYIAWAAGY YPPESTTGSF GLYSSLCFDF TLTNIFCPLV
RGKTLRIYPQ SESIDTILAR MFQPGSGVDT LKLTPTHIHL LEYMNLRASG
VRKVIVGGEE LTPQHIAATLR KIDPAIEIYN EYGPTAATVG CIVERVEDAP
PTVLIGRPIA DTRVYMLDDA LRPVPLGVPG EICLAGAGLA RGYHQRPDVT
AAKFVEHFFP GEARIYRTGD IGRWLPDGRI QCYGRVDHQV KIRGHRVELG
EIEAAIAAHE DVVGAAMVLR ESAHGVRKLA AYVKGAAASLS VPNLRAYLAG
KLPDYMVPSD IPIAEFPLN ANGKLDRLPAL LALEPAAPE EAPLDATPIQ
RELVRIWRDV LDNPAVDLAG RFFDYGGDSL QAMQLVSRIV SSFSVEIGID
AIPFELQTISA VSDLIEASSP HPGSTAGAIP PRSRANDLPL SFPQORLWFL
AQLEGPSATY NISSALRFEG ELDVARLRFA VSEISRHEI LRTTFPAVDG
RGVQRIAPPA PVALDVVDVA SESDTLALLA EEADRPPDLA AGPLYRVVLY
RVHERLHVFG IVMHHIVSDA WSSGILIGEL AALYAGESLP ELAVQYADYA
VWQHERLASA DTHRELALLS AALADAPDLI ELPTDRPRPA VQQFRGAVLP
FQLSAERADG LRAIARASGT STFMVLAAY ALLLSRYSNQ QDLVIGSPIA
NRRSSMTEPL IGFANMLAL RVDLSGNPTF GDLLARVKRV ALDGYSRQEI
PFEQVDSLE LERNLGRTPV FQVVFAYEKA QPRAVSFPGL VATFVAVETH
TAKFDLTLHV QDADDGLAGS LEYNLDLFDATAIDRMAEHF RTLVDVAIAD
PDRPLGALS SNAERNLLT VEWNRDTDF GEDAAQPLHR LFEQQVERTP
```

-continued

DAVAIVFDOT ALTYAELNLR ANRLAHLVA LGVGPDSL VGAMERSLDM  
 VALLAILKAG GAYVPVDPDY PAERVRFMID HAQLRWLLTQ QHLHDALPDT  
 DAHVIVVDRD SLDLDAATS NPAPALNGDN LAYMIYTS GS TGRPKGALNT  
 HRAITNRILW MQHAYALDAD DAVLQKTPFS FDVSVWELFW PLVTGARLVF  
 ARPGGQRETD YLVELIERER ITTIHFVPSM LRAFLDHPDL DAHCASLRRV  
 VCSGEALPHD LQQRCLERLD VKLYNLYGPT EAAVDVTAW E CRRDDPHRIV  
 PIGRPIANTR LYIVDAQMQP TPIGVAGELL IGGTPVGRGY HGEPELSAEK  
 FIADPFSADP LARLYRTGDL ARYRPDGNIE FLGRIDHQIK LRGLRIEPE  
 IEAALRAHPS VDDCVVIKT EGARTPLIAY VATAAPDIAD LRGYLGKLA  
 DYMVPSQFFA LESLPLPNG KINKALPLP ADRGDAAQPH APAVTPREIL  
 LASICIDLQ LPSVGIHDF FELGGDSILS IQVIARANQA GLRVTAKQLF  
 QYQTIAQLAA APEERAACAP TLSPLGDAPL TPVQHWFPEQ EIDAPSHYNQ  
 TVLIQVPADI DASRLADAFR QVYEHHDALR LRFSDAGRW TQQVVGEM  
 PALFAKQVIA DDAGERLAAM RAAAADAERG IDITHGPLLA ARLFCLADEP  
 LARLFVSIHH LAVDGVSWRV LLEDLHAAYH GQPLPGKTS FREWALHQQ  
 LARSPAIGDE ARLWQALLAQ PVEPMPVDYP GTGAANNAV DASSVSFELG  
 EADTTALLR LPRAYDTRIN DVLLVALAQA CSMVTGNTRT RIDLESHGRH  
 VSDAPLDLTR TVGWFTSIYP VVLDADAMHA PEQALRAARQ QLRRIPADGL  
 GYSLRLYQSP DAAVRDSLAA LPKADILFNY HGQLDVLRLQ SDGWRPAAED  
 LGSRLRAGRSQ RTHAFEIVAA VADGKLQVDW RYGERLHRRQ TVENLAAHFR  
 DRLLDFAASV PDTAADDIED SYPLSSLQQG ILFHSLYDLD PAAYFQQFSF  
 VVSGPLQVPA LRQAWANALA RHAVLRTAFA WADRHPVQT VRHTVDLPWT  
 FLDWRHRDAS RRAQDFDAFL ADDRRRGFDL QRAPLFRCTL IQETDTRHRF  
 CWSAHHIILD GWSTATLMKE VFDDYLSLAR TGMPAVAASA PGYRAYIDWL  
 ARHPRSADET WWRAELAGFK AATPVAASPA RQATGDAPRQ DKRRTQQFLL  
 DEALAARLQT LTRTHRVTLN VLIRAVWALV LRRHAGTDDV VFGVTVSGRP  
 PMLDGVESIV GLFINTLPLR LRIAPERPFI EWLAEVHAAQ TAMEPHSYSS  
 LVDIQSWSEL PAGDSLFDL LVFENFPVAA APDLGPDDIE ILDTRAFAES  
 NYPLTLTVHP NERIGFHISH DAHRIAPEVV RQMLDTRLTL LERFAENPGQ  
 LTGQLADPPA ADGRPSAPRS GAGPAIEAAA GAAAAARAVA HAADESTLLE  
 IWRRIKRD IAVSDNYFDL GGHSIIAIQL MAHVEKAPDR RLPISCLFEN  
 PTIEKLAAL AAKEPSAPAG GLVPIRDGGP AAPLFLPGA GGNVYFRPL  
 ANHLSGAHAI HGLEALGLDG ACEPLTRVED IAARHIERIW PLVGAGPYL  
 AGHSFGAHVA LEMSRLVAK GADVLLAIF DASAPIDSSA ATYWQDWDT  
 EWLVAIAHEI GTFLGDLQV TRADLVHLDP DGQAGLILR IGDRGSWFAD  
 AGSRLRAYL RVYQANFKSH YAPHATPLPV PIALFRSTER DPGDYAPSS  
 IAQLRLDATW GWSRFSAPV AVTDVPGDHL TMLLDPHAGV LAAHVNSFLE KTPS

SEQ ID NOs: 5-23: polynucleotide and polypeptides associated with GenBank Accession No. EU938698.5.

#### DETAILED DESCRIPTION OF THE INVENTION

This invention relates to antifungal compounds and their therapeutic use in the prevention or treatment of fungal infections and diseases. Particularly, various aspects of the invention provide compositions enriched for occidiofungin diastereomers/conformers that have higher activity against fungal infections or diseases. Thus, the invention relates to methods of making such occidiofungin diastereomers/conformers, compositions enriched for such diastereomers/conformers and methods of using compositions comprising occidiofungin diastereomers/conformers disclosed herein as fungicides for animals and plants. The invention further relates to the microorganisms that produce compositions enriched for occidiofungin enriched for occidiofungin diastereomers/conformers corresponding to diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations). Methods of increasing the production of occidiofungin diastereomers/conformers corresponding to diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations) in microorganisms and productions systems are also provided.

As discussed above, one aspect of the invention provides compositions enriched for occidiofungin diastereomers/conformers, in particular the occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations observed under the following conditions: 2 mM samples of occidiofungin diastereomers in dimethylsulfoxide (DMSO-d<sub>6</sub>, Cambridge Isotopes) subjected to 2-D TOCSY, spectra collected at 323 K with a mixing time of 60 milliseconds and data processing using NMRPipe with 45 degree sinebell squared shifts in both dimensions). Thus, the invention provides compositions enriched for such antifungal diastereomers/conformers for treating fungal infection. In certain embodiments of this aspect of the invention, pharmaceutical and agricultural compositions that contain a composition enriched for diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations) are provided.

Another aspect of the invention provides for compositions that are enriched for a particular occidiofungin diastereomer/conformer. In this aspect of the invention, the activity of the ocfD and/or ocfN thioesterases is altered such that the activity of one of the thioesterases is decreased (or eliminated) and the activity of the second thioesterase remains functional or is increased. Thus, microorganisms can be genetically manipulated such that OcfD thioesterase activity is decreased or eliminated and the thioesterase activity of OcfN is increased or maintained at unaltered (e.g., levels of activity as observed in *Burkholderia contaminans* MS14 or microorganisms engineered with the biosynthetic pathway for the production of occidiofungin). Alternatively, microorganisms can be genetically manipulated such that OcfN activity is decreased or eliminated and the thioesterase activity of OcfD is increased or unaltered.

Compositions comprising occidiofungin diastereomers/conformers as disclosed herein may be formulated prior to administration in an agriculturally acceptable carrier, for example in an aqueous carrier, medium or suitable diluent, such as saline or other buffer. The formulated compositions may also be in the form of a dust or granular material, or a

suspension in oil (vegetable or mineral), water or oil/water emulsions, a wettable powder, or in combination with any other carrier material suitable for agricultural application. Suitable agricultural carriers can be solid or liquid and are well known in the art. The term “agriculturally-acceptable carrier” covers all adjuvants, e.g. inert components, dispersants, surfactants, tackifiers, binders, etc. that are ordinarily used in the formulation of agricultural compositions; these are well known to those skilled in formulation of agricultural compositions.

A pharmaceutical composition contains a desired amount of an occidiofungin diastereomers/conformers as disclosed herein. Thus, the pharmaceutical composition can comprise occidiofungin diastereomers/conformers having the total correlation spectroscopy (TOCSY) fingerprint identified in FIG. 5C as the green NH correlations or the pharmaceutical composition can comprise a particular occidiofungin diastereomer/conformer. Either of these pharmaceutical compositions can be in the form of, for example, a tablet, a caplet, a pill, a hard or soft capsule, a lozenge, a cachet, a dispensable powder, granules, a suspension, an elixir, a dispersion, a liquid, or any other form reasonably adapted for administration. If intended for parenteral administration, it can be in the form, for example, of a suspension or transdermal patch. Presently preferred are oral dosage forms that are discrete dose units each containing a predetermined amount of the disclosed occidiofungin diastereomers/conformers.

Pharmaceutically acceptable carriers that may be used in these compositions include, but are not limited to, ion exchangers, alumina, aluminum stearate, lecithin, serum proteins, such as human serum albumin, buffer substances such as phosphates, glycine, sorbic acid, potassium sorbate, partial glyceride mixtures of saturated vegetable fatty acids, water, salts or electrolytes, such as protamine sulfate, disodium hydrogen phosphate, potassium hydrogen phosphate, sodium chloride, zinc salts, colloidal silica, magnesium trisilicate, polyvinyl pyrrolidone, cellulose-based substances, polyethylene glycol, sodium carboxymethylcellulose, polyacrylates, waxes, polyethylene-polyoxypropylene-block polymers, polyethylene glycol and wool fat.

The compositions of the present invention may be administered orally, parenterally, by inhalation spray, topically, rectally, nasally, buccally, vaginally or via an implanted reservoir. The term “parenteral” as used herein includes subcutaneous, intravenous, intramuscular, intra-articular, intra-synovial, intrasternal, intrathecal, intrahepatic, intralésional and intracranial injection or infusion techniques. Preferably, the compositions are administered orally, intraperitoneally or intravenously.

Sterile injectable forms of the compositions of this invention may be aqueous or an oleaginous suspension. These suspensions may be formulated according to techniques known in the art using suitable dispersing or wetting agents and suspending agents. The sterile injectable preparation may also be a sterile injectable solution or suspension in a non-toxic parenterally acceptable diluent or solvent, for example as a solution in 1,3-butanediol. Among the acceptable vehicles and solvents that may be employed are water, Ringer’s solution and isotonic sodium chloride solution. In addition, sterile, fixed oils are conventionally employed as a solvent or suspending medium. For this purpose, any bland fixed oil may be employed including synthetic mono- or diglycerides. Fatty acids, such as oleic acid and its glyceride derivatives are useful in the preparation of injectables, as are natural pharmaceutically-acceptable oils, such as olive oil or castor oil, especially in their polyoxyethylated versions. These oil solutions or suspensions may also contain a

long-chain alcohol diluent or dispersant, such as carboxymethyl cellulose or similar dispersing agents that are commonly used in the formulation of pharmaceutically acceptable dosage forms including emulsions and suspensions. Other commonly used surfactants, such as Tweens, Spans and other emulsifying agents or bioavailability enhancers which are commonly used in the manufacture of pharmaceutically acceptable solid, liquid, or other dosage forms may also be used for the purposes of formulation.

The compositions of this invention may be orally administered in any orally acceptable dosage form including, but not limited to, capsules, tablets, aqueous suspensions or solutions. In the case of tablets for oral use, carriers commonly used include lactose and corn starch. Lubricating agents, such as magnesium stearate, are also typically added. For oral administration in a capsule form, useful diluents include lactose and dried cornstarch. When aqueous suspensions are required for oral use, the active ingredient is combined with emulsifying and suspending agents. If desired, certain sweetening, flavoring or coloring agents may also be added.

Alternatively, the compositions of this invention may be administered in the form of suppositories for rectal administration. These can be prepared by mixing the agent with a suitable non-irritating excipient that is solid at room temperature but liquid at rectal temperature and therefore will melt in the rectum to release the drug. Such materials include cocoa butter, beeswax and polyethylene glycols. The compositions of this invention may also be administered topically, ophthalmically, by nasal aerosol or inhalation. Such compositions are prepared according to techniques well-known in the art of pharmaceutical formulation.

Compositions disclosed herein can be used to treat fungal infections in immunocompromised patients or patients having fungal infections. Thus, another aspect of the invention provides for administering compositions enriched for occidiofungin diastereomers/conformers (e.g., those corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations) having increased antifungal activity as compared to occidiofungin compositions produced by *Burkholderia contaminans* MS14 (disclosed in U.S. Patent Application Publication 2011/0136729, the disclosure of which is hereby incorporated by reference in its entirety). These diastereomers/conformers have been characterized by a number of techniques, including COSY, TOCSY, NOESY, ROESY, and HSQC 2D NMR spectroscopy experiments.

The antifungal activity of the disclosed occidiofungin diastereomers/conformers (diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations)) provides for compositions having greater antifungal activity as compared to as compared to occidiofungin compositions produced by *Burkholderia contaminans* MS 14 when cultured under the same conditions. The phrase "enriched for the disclosed occidiofungin diastereomers/conformers" is intended to convey that the composition contains disclosed occidiofungin diastereomers/conformers (diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations)) in amounts higher than that produced by a reference strain (e.g., *Burkholderia contaminans* MS14 as disclosed in the examples provided herein). Thus, the phrase indicates that at least 37% of the total amount of occidiofungin diastereomers/conformers present within an enriched composition are the disclosed diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations)). In various embodiments, compositions

"enriched for the disclosed occidiofungin diastereomers/conformers" contain at least 40%, 41%, 42%, 43%, 44%, 45%, 46%, 47%, 48%, 49%, 50%, 51%, 52%, 53%, 54%, 55%, 56%, 57%, 58%, 59%, 60%, 61%, 62%, 63%, 64%, 65%, 66%, 67%, 68%, 69%, 70%, 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, 79%, 80%, 81%, 82%, 83%, 84%, 85%, 86%, 87%, 88%, 89%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98% or 99% of the disclosed diastereomers/conformers in relation to the total amount of occidiofungin diastereomers/conformers in a composition.

As discussed above, one aspect of the invention provides microorganisms capable of producing compositions enriched for occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations). In this aspect of the invention, microorganisms are transformed with the genes associated with the biosynthesis of occidiofungin. These genes and open reading frames (ORFs) are disclosed in disclosed in U.S. Patent Application Publication 2011/0136729, the disclosure of which is hereby incorporated by reference in its entirety; Gu et al., *Appl. Environ. Microbiol.*, 2011, 77:(17):6189-6198 which is also incorporated by reference in its entirety and GenBank Accession No. EU938698.5, which is also hereby incorporated by reference in its entirety and is also provided on pages 29-72 (SEQ ID NOs: 5-23). These transformed microorganisms are further manipulated genetically such that the microorganisms exhibit an increase in the level of OcfN thioesterase (SEQ ID NO: 3) activity. An increase in the level of OcfN thioesterase activity can be achieved by means of expressing the ocfN gene in a multicopy plasmid with a native promoter or any other promoter sequence. Another way to increase the expression of the ocfN gene within the cell is to chromosomally integrate additional copies of the ocfN gene using transposons. Yet a further means to increase ocfN thioesterase activity is to substitute the native promoter associated with the ocfN gene with a promoter that increases expression of the gene (relative to the native promoter). In certain embodiments of this aspect of the invention, the thioesterase activity of OcfD can be decreased or eliminated by a point mutation of the catalytic serine at position 2954 of SEQ ID NO: 4, insertional mutation or point mutation of amino acids within the thioesterase motif (in addition to the substitution of the serine residue) found in ocfD to reduce or eliminate its activity, deletion of the catalytic serine or other portions of SEQ ID NO: 4 (e.g., portions or the entirety of the thioesterase motif in SEQ ID NO: 4) or truncation SEQ ID NO: 4 such that thioesterase activity is reduced or eliminated (in addition to increasing the level of OcfN thioesterase activity) within the genetically modified microorganisms.

Another aspect of the invention provides for compositions enriched for a particular Occidiofungin diastereomer/conformer. In this aspect of the invention, transformed microorganisms are manipulated genetically such that the microorganisms exhibit an increase in the level of OcfD thioesterase (SEQ ID NO: 4) activity. An increase in the level of OcfD thioesterase activity can be achieved by means of expressing the ocfD gene in a multicopy plasmid with a native promoter or any other promoter sequence. Another way to increase the expression of the ocfD gene within the cell is to chromosomally integrate additional copies of the ocfD gene using transposons. Yet a further means to increase ocfN thioesterase activity is to substitute the native promoter associated with the ocfD gene with a promoter that increases expression of the gene (relative to the native promoter). In certain embodiments of this aspect of the invention, the

thioesterase activity of OcfN can be decreased or eliminated by a point mutation of the catalytic serine at position 73 of SEQ ID NO: 3, insertional mutation or point mutations of other amino acids within the thioesterase motif (in addition to the substitution of the serine residue) of the thioesterase to reduce or eliminate its activity, deletion of the catalytic serine or other portions of SEQ ID NO: 3 (e.g., portions or the entirety of the thioesterase motif in SEQ ID NO: 3), truncation SEQ ID NO: 3 such that thioesterase activity is reduced or eliminated or deletion of ocfN in its entirety (in addition to increasing the level of OcfD thioesterase activity) within the genetically modified microorganisms. Where the biosynthetic pathway for occidiofungin biosynthesis is engineered into a microorganisms, once can, of course, omit ocfN to achieve the same effect as the mutation or deletion of ocfN as discussed above.

Thus, microorganisms such as bacterial cells, fungal cells and yeast can be transformed with genes encoding the occidiofungin biosynthetic pathway and genetically manipulated, as discussed above, such that the cells have increased OcfN activity and/or decreased OcfD activity as compared to reference bacterial, fungal or yeast cells. Alternatively, microorganisms such as bacterial cells, fungal cells and yeast can be transformed with genes encoding the occidiofungin biosynthetic pathway and genetically manipulated, as discussed above, such that the cells have increased OcfD activity and/or decreased OcfN activity as compared to reference bacterial, fungal or yeast cells. Such cells can then be used to produce compositions enriched for occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations) or to produce compositions enriched for a particular occidiofungin diastereomer/conformer. The phrase "reference bacterial, fungal or yeast cells" refers to bacterial, fungal or yeast cells containing the genes associated with the biosynthetic pathway for the production of occidiofungin and where the function/activity of OcfN and/or OcfD has not been altered as disclosed herein. Thus, the phrase "reference bacterial, fungal or yeast cells" refers to cells containing, for example, polynucleotide (SEQ ID NO: 23 encoding the open reading frames (ORFs; SEQ ID NOs: 5-22)) disclosed in GenBank Accession No. EU938698.5. For the comparison of compositions comprising particular occidiofungin conformers (or compositions enriched for occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations)), compositions containing the diastereomers/conformers are obtained from cells genetically manipulated to have increased ocfN activity and/or decreased OcfD activity (or increased OcfD activity and/or decreased ocfN activity) and compared to compositions containing occidiofungin produced by reference bacterial, fungal or yeast cells cultured under similar or the same conditions (e.g., the same temperature and medium).

Bacterial cells can be selected Gram negative bacteria or Gram positive bacteria. In this aspect of the invention, the Gram-negative bacterial cell can be selected from the group consisting of *Escherichia*, *Zymomonas*, *Acinetobacter*, *Glucobacter*, *Geobacter*, *Shewanella*, *Salmonella*, *Enterobacter* and *Klebsiella*. Gram-positive bacteria can be selected from the group consisting of *Bacillus*, *Clostridium*, *Corynebacterial*, *Lactobacillus*, *Lactococcus*, *Oenococcus*, *Streptococcus* and *Eubacterial* cells. Various thermophilic bacterial cells, such as *Thermoanaerobes* (e.g., *Thermoanaerobacterium saccharolyticum*), *Bacillus* spp., e.g., *Bacillus coagulans* strains, *Bacillus licheniformis* strains, *Bacil-*

*lus subtilis* strains, *Bacillus amyloliquifaciens* strains, *Bacillus megaterium* strains, *Bacillus maceans* strains, *Paenibacillus* spp. strains or *Geobacillus* spp. such as *Geobacillus stearothermophilus*.

Yeast cells suitable for use in this aspect of the invention may be a *Candida*, *Hansenula*, *Kluveromyces*, *Pichia*, *Saccharomyces*, *Schizosaccharomyces*, or *Yarrowia* cell such as a *Kluveromyces lactis*, *Saccharomyces carlsbergensis*, *Saccharomyces cerevisiae*, *Saccharomyces diastaticus*, *Saccharomyces douglasii*, *Saccharomyces kluyveri*, *Saccharomyces norbensis*, *Saccharomyces oviformis*, or *Yarrowia lipolytica* cell. In this aspect of the invention, the yeast cell must be resistant to the effects of occidiofungin to be a viable production system for compositions enriched for occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations).

In other embodiments of this aspect of the invention, fungal cells can be manipulated to produce compositions enriched for occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations). "Fungi" as used herein includes the phyla *Ascomycota*, *Basidiomycota*, *Chytridiomycota*, and *Zygomycota*, *Oomycota* and all mitosporic fungi. A fungal cell may be a yeast cell. "Yeast" as used herein includes ascosporegenous yeast (*Endomycetales*), basidiosporegenous yeast, and yeast belonging to the Fungi Imperfecti (*Blastomycetes*). The fungal host cell may be a filamentous fungal cell. "Filamentous fungi" include all filamentous forms of the subdivision *Eumycota* and *Oomycota* (as defined by Hawksworth et al., 1995, supra). The filamentous fungi are generally characterized by a mycelial wall composed of chitin, cellulose, glucan, chitosan, mannan, and other complex polysaccharides. Vegetative growth is by hyphal elongation and carbon catabolism is obligately aerobic. In contrast, vegetative growth by yeasts such as *Saccharomyces cerevisiae* is by budding of a unicellular thallus and carbon catabolism may be fermentative. The filamentous fungal host cell may be an *Acremonium*, *Aspergillus*, *Aureobasidium*, *Bjerkandera*, *Ceriporiopsis*, *Chrysosporium*, *Coprinus*, *Coriolus*, *Cryptococcus*, *Filibasidium*, *Fusarium*, *Humicola*, *Magnaporthe*, *Mucor*, *Myceliophthora*, *Neocalimastix*, *Neurospora*, *Paecilomyces*, *Penicillium*, *Phanerochaete*, *Phlebia*, *Piromyces*, *Pleurotus*, *Schizophyllum*, *Talaromyces*, *Thermoascus*, *Thielavia*, *Tolyocladium*, *Trametes*, or *Trichoderma* cell. For example, the filamentous fungal host cell may be an *Aspergillus awamori*, *Aspergillus foetidus*, *Aspergillus fumigatus*, *Aspergillus japonicus*, *Aspergillus nidulans*, *Aspergillus niger*, *Aspergillus oryzae*, *Bjerkandera adusta*, *Ceriporiopsis aneirina*, *Ceriporiopsis caregiea*, *Ceriporiopsis gilvescens*, *Ceriporiopsis pannocinta*, *Ceriporiopsis rivulosa*, *Ceriporiopsis subrufa*, *Ceriporiopsis subvermisporea*, *Chrysosporium inops*, *Chrysosporium keratinophilum*, *Chrysosporium lucknowense*, *Chrysosporium merdarium*, *Chrysosporium pannicola*, *Chrysosporium queenlandicum*, *Chrysosporium tropicum*, *Chrysosporium zonatum*, *Coprinus cinereus*, *Coriolus hirsutus*, *Fusarium bactridioides*, *Fusarium cerealis*, *Fusarium crookwellense*, *Fusarium culmorum*, *Fusarium graminearum*, *Fusarium graminum*, *Fusarium heterosporum*, *Fusarium negundi*, *Fusarium oxysporum*, *Fusarium reticulatum*, *Fusarium roseum*, *Fusarium sambucinum*, *Fusarium sarcochroum*, *Fusarium sporotrichioides*, *Fusarium sulphureum*, *Fusarium torulosum*, *Fusarium trichothecioides*, *Fusarium venenatum*, *Humicola insolens*, *Humicola lanuginosa*, *Mucor miehei*, *Myceliophthora ther-*

*mophila*, *Neurospora crassa*, *Penicillium purpurogenum*, *Phanerochaete chrysosporium*, *Phlebia radiata*, *Pleurotus eryngii*, *Thielavia terrestris*, *Trametes villosa*, *Trametes versicolor*, *Trichoderma harzianum*, *Trichoderma koningii*, *Trichoderma longibrachiatum*, *Trichoderma reesei*, or *Trichoderma viride* cell.

Fungal cells may be transformed by a process involving protoplast formation, transformation of the protoplasts, and regeneration of the cell wall in a manner known per se. Suitable procedures for transformation of *Aspergillus* and *Trichoderma* host cells are described in EP 238023, Yelton et al., 1984, Proc. Natl. Acad. Sci. USA 81: 1470-1474, and Christensen et al., 1988, Bio/Technology 6: 1419-1422. Suitable methods for transforming *Fusarium* species are described by Malardier et al., 1989, Gene 78: 147-156, and WO 96/00787. Yeast may be transformed using the procedures described by Becker and Guarente, In Abelson, J. N. and Simon, M. I., editors, Guide to Yeast Genetics and Molecular Biology, Methods in Enzymology, Volume 194, pp 182-187, Academic Press, Inc., New York; Ito et al., 1983, J. Bacteriol., 153: 163; and Hinnen et al., 1978, Proc. Natl. Acad. Sci. USA 75: 1920.

In another embodiment of the present invention, the native promoter of the *ocfN* gene within *Burkholderia contaminans* MS14 can be replaced by promoter elements known to enhance the level of gene expression, thereby increasing *OcfN* thioesterase activity within *Burkholderia contaminans* MS14. *Burkholderia contaminans* MS14 can also be genetically modified by other techniques to produce compositions enriched for occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations). Genetic modifications that *ocfN* thioesterase activity include the introduction of multicopy plasmids comprising a native promoter or any other promoter sequence operably linked to an *ocfN* gene into *Burkholderia contaminans* MS14, integration of additional copies of the *ocfN* gene operably linked to a promoter into the chromosome of *Burkholderia contaminans* MS14 using transposon mutagenesis or by replacement of the native *ocfN* promoter in *Burkholderia contaminans* MS14 with a promoter that increases the expression of *ocfN* transcripts relative to the native promoter sequence.

Another aspect of the invention provides for the introduction of a point mutation into the nucleotide sequence encoding *OcfD*, the truncation of *ocfD* (or introduction of a frameshift mutation) such that the thioesterase activity is reduced or eliminated or the deletion of the segment of the *ocfD* gene encoding the catalytic serine in order to increase the amounts of occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the TOCSY fingerprint identified in FIG. 5C (the green NH correlations) produced by *Burkholderia contaminans* MS14 or by microorganisms genetically modified to produce occidiofungin (e.g., microorganisms into which the biosynthetic pathway for occidiofungin production have been introduced). In this aspect of the invention, a point mutation is introduced into the catalytic serine in the thioesterase domain of *ocfD* in order to reduce its activity. This amino acid is found at position 2954 of SEQ ID NO: 4. For example, the serine can be mutated into an alanine, glycine or proline residue (with glycine or alanine being preferred in this context). Certain embodiments of this aspect of the invention also provide for genetic modification of the microorganisms such that *ocfN* activity is increased as well (e.g., the level of *OcfN* thioesterase activity can be increased by means of expressing the *ocfN* gene in a multicopy plasmid

with a native promoter or any other promoter sequence, chromosomal integration of additional copies of the *ocfN* gene using transposons or other means or substitution of the native promoter associated with the *ocfN* gene with a promoter that increases expression of the gene (relative to the native promoter)).

Another aspect of the invention provides for the introduction of a point mutation into the nucleotide sequence encoding *OcfN*, the truncation of *ocfN* (or introduction of a frameshift mutation) such that the thioesterase activity is reduced or eliminated, the deletion of the segment of the *ocfN* gene encoding the catalytic serine or chromosomal deletion of *ocfN* within a microorganism (e.g., *Burkholderia contaminans* MS14) in order to increase the amounts a particular occidiofungin diastereomer/conformer produced by a microorganism. As would be apparent to one skilled in the art, a similar effect can be obtained by transforming a microorganism with the genes encoding the occidiofungin biosynthetic pathway, with the exception of *ocfN* gene. In this aspect of the invention, a point mutation is introduced into the catalytic serine in the thioesterase domain of *OcfN* in order to reduce its activity. This amino acid is found at position 73 of SEQ ID NO: 3. For example, the serine can be mutated into an alanine, glycine or proline residue (with glycine or alanine being preferred in this context). Certain embodiments of this aspect of the invention also provide for genetic modification of the microorganisms such that *OcfD* activity is increased as well (e.g., the level of *OcfD* thioesterase activity can be increased by means of expressing the *ocfD* gene in a multicopy plasmid with a native promoter or any other promoter sequence, chromosomal integration of additional copies of the *ocfD* gene using transposons or other means or substitution of the native promoter associated with the *ocfD* gene with a promoter that increases expression of the gene (relative to the native promoter)).

#### Materials and Methods

Proportion of Occidiofungin Variants in the Sample. The C-terminal TE domain of *OcfD* and the *OcfN* cyclase thioesterase in the occidiofungin biosynthetic gene cluster are both predicted to be involved in the termination of synthesis and formation of the cyclic peptide. Given that the N-terminal end of the linear peptide is an Asn or BHN, we hypothesized that each thioesterase was required for cyclization of the Asn1 and BHN1 variants. The Asn1 and BHN1 variants of occidiofungin are not separable by RP-HPLC (reverse phase high performance liquid phase chromatography), thus, both variants are present in the purified fraction (FIG. 2). The final RP-HPLC step in the purification process reveals the presence of three peaks. Occidiofungin samples elute as a doublet peak before the third peak. Both the wild type strain MS14 and the *ocfN* mutant MS14GG88 have the same chromatographic profile as observed in the last purification step. Occidiofungin peaks were confirmed by MALDI-TOF and bioassays. It is important to note that the presence of the doublet peak is not associated with the presence of Asn1 or BHN 1. Each peak of the doublet contains both the Asn1 and BHN 1 variants.

The relative proportion of the Asn1 and BHN1 variants could not be directly compared, because direct measurement of the Asn1 peak intensities could not be done due to the peaks overlapping with Asn7. The relative proportion of the Asn1 and BHN 1 variants in the wild-type fraction was determined by measuring the <sup>13</sup>C-HSQC Ha-Ca cross peak intensities of each BHY4 peak in the data set,<sup>27, 28</sup> given that each of the BHY4 peaks could be attributed to either the



Asn1 or BHN1 variant. Based on the Ha-Ca cross peak intensities for BHY4 in HSQC spectrum, the Asn1 and BHN1 variants was determined by measuring the  $^{13}\text{C}$ -HSQC Ha-Ca cross peak intensities of each BHY4 peak in the data set<sup>27, 28</sup>, and was determined to be approximately 36% and 64% of the total amount of occidiofungin, respectively (FIG. 3). The peaks in red and green represent the BHY4 peaks associated with BHN1 and Asn1 variants, respectively. A similar ratio was also observed in the relative abundance of each peak in the ESI-MS spectrum (FIG. 4A). Furthermore, the  $^{13}\text{C}$ -HSQC Ha-Ca cross peak intensities for the BHN1 peaks in the spectra were determined to be 90.50 and 38.65, which support the intensities measured for BHY4 peaks corresponding to the BHN1 conformational variants.

Mutagenesis of the *ocfN* gene was conducted via a marker exchange procedure as described previously<sup>22</sup>, to generate the mutant MS14GG88. The percentage of Asn1 to BHN1 variants in the *ocfN* mutant MS14GG88 fraction could be determined by measuring the proportion of each BHN1 variant using the HSQC data set and by the integration of the HN of Asn1 and BHN1 in the  $^1\text{H}$  NMR spectra. Asn1 and BHN1 variants are approximately 20% and 80% of the total amount of occidiofungin, respectively. The ESI-MS spectrum also shows a lower relative abundance for the Asn1 variant (1200.39 Da) compared to the BHN1 variant (1216.41 Da) (FIG. 4B).

Comparison of Wild-Type and *ocfN* Mutant NMR Spectra. Occidiofungin has a complex spectrum for a peptide of only eight amino acids (FIG. 5A and Table 1). The NMR spectrum represents an average of the conformers on the NMR time scale. Conformers in slow exchange on the NMR time scale may result in multiple spin systems for each amino acid. In some situations, multiple conformers are known to arise for cyclic peptides due to slow interconverting conformational families.<sup>29, 30</sup> Despite the conformational restrictions brought about by the ring closure, occidiofungin still has a significant amount of conformational freedom. Both Asn1 and BHN1 variants are visibly present in the wild-type fraction, which are colored red in FIG. 5A. The TOCSY fingerprint region (NH correlations) is not as complex for the *ocfN* thioesterase MS14GG88 mutant spectra (FIG. 5B). A significant number of spin systems found in the wild-type spectra are absent in the *ocfN* thioesterase mutant spectra. Our experiments show that the TE domain on the C-terminal region of *OcfD* is able to perform the peptide macrocyclization of both the Asn1 and BHN1 variants. Although, there is only one amide spin system for Asn1 produced by *OcfD*. Whereas, the loss of *OcfN* results in the disappearance of the other three Asn1 amide spin systems.

An overlay of the wild-type and *ocfN* mutant NMR spectra shows the amino acid spin systems in green that are absent in the mutant spectra (FIG. 5C). These spin systems are for Asn7, Ser8, Asn1, Novel Amino Acid 2 (NAA2), Ser3, BHY4, and Gly6. The loss of these spin systems suggests that the complex spin system observed for the wild-type occidiofungin fraction is not only due to interconverting conformational families, but is the result of distinct diastereomers formed by the regiospecific activity of the *OcfN* cyclase and *OcfD* TE domain. Dramatic chemical shifts observed, such as the 2 ppm shift for HN of the NAA2, support the formation of a structurally unique conformer of occidiofungin. A unique conformer is further supported by the subsequent loss of a NAA2 spin system in the *ocfN* mutant NMR spectra. Furthermore, the presence of both Asn1 and BHN1 spin systems in the mutant spectra along with the absence of the amide spin systems shown in green indicate that the additional spin systems are not due to the

presence of the  $\beta$ -hydroxyl on Asn1. The additional spin systems are due to the formation of unique diastereomer produced by *OcfN* cyclase thioesterase. To further test for the formation of a configurational isomer versus an interchangeable conformational isomer, one dimensional NMR temperature titrations were performed. Amide and aromatic regions revealed little change in the complexity of peaks present with the occidiofungin derived from *ocfN* mutant MS14GG88 or wild-type strain MS14 (FIGS. 13A-13B). Given that NAA2 spin systems are a good indicator for the presence of both diastereomers in the wild-type spectrum, we collected TOCSY spectra for occidiofungin derived from *ocfN* mutant MS14GG88 or wild-type strain at 50° C. (FIGS. 14A-14B). There was no loss or addition of a spin system for NAA2 in the mutant spectrum. Furthermore, both spin systems for NAA2 remained in the wild-type spectrum. This data supports that the stereoisomers are non-interchangeable isomers, supporting their classification as a diastereomers (configurational isomers) rather than a conformational isomer.

Model for the Coordinated Function of Two Cyclase Thioesterases. There was no loss of an amide spin system for a BHN1 in the *ocfN* mutant NMR spectra. This suggests that *OcfN* thioesterase has a substrate requirement for the peptide containing Asn1, since there is no concomitant loss of a BHN1 spin system with the observed loss of the Asn1 spin systems. The C-terminal TE domain of *OcfD* has a preference for the peptide containing the BHN1, but is capable, albeit at a lower efficiency of cyclizing the Asn1 variant. This provides an interesting scenario for the activity of the two thioesterases (FIG. 6). Both thioesterases contain the GX SXG motif, which is important for the catalytic transfer of the peptide from the T domain to the cyclase. This suggests that substrate recognition occurs prior to the catalytic transfer of the peptide to the cyclase. Presumably, *OcfN* cyclase has a higher affinity or better access for the Asn1 peptide product given that the proportion of the Asn1 cyclic peptide product produced by *OcfD* compared to the BHN1 product is reduced in the wild-type fraction. Therefore the biosynthesis of occidiofungin utilizes the structural differences between Asn and BHN to increase the conformational biodiversity of occidiofungin. The increase in conformational diversity is accomplished by the regiospecific activity of each cyclase, presumably by differences in their binding clefts that helps orientate the peptide before cyclization.

Comparison of the Bioactivity of the Wild-Type and *ocfN* Mutant Product. To determine whether the increase in conformational diversity is important for bioactivity, minimum inhibitory concentrations were determined against medically relevant *Candida* species (FIG. 7A). There was a 2-fold decrease in the minimum inhibitory concentration (MIC) with the purified *ocfN* mutant product with respect to the wild-type product against *Candida albicans* LL, *Candida albicans* TE, *Candida glabrata* ATCC66032, *Candida parapsilosis* ATCC90018, and *Candida tropicalis* ATCC66029. There was no difference in the MIC for *Candida albicans* ATCC66027. Colony forming units (CFUs/mL) were determined for the MIC wells of wild-type product for each *Candida* species and compared to the corresponding well containing the same concentration of the *ocfN* mutant product (FIG. 7B). Following exposure to the same concentration of wild-type and *ocfN* mutant products, these results show a 5 to 7-log decrease in cell density of the *Candida* species treated with wild-type product. The differences in activity are also visualized by the rate of cell death. Time-kill experiments were performed against *Candida glabrata* ATCC66032. There was a ten-fold difference in

yeast present at 4 and 8 hours when cells were treated with 0.5  $\mu\text{g}/\text{mL}$  of occidiofungin derived from ocfN mutant MS14GG88 or wild-type strain (FIG. 15). Furthermore, a slower rate of cell death was also observed for yeast treated with occidiofungin derived from ocfN mutant MS14GG88 at 1.0 and 2.0  $\mu\text{g}/\text{mL}$ . Given that the cyclic occidiofungin variants produced by OcfN constitute less than half of the total structural variants, a 2-fold loss in activity suggests that the configurational isomer made by OcfN are 4-fold more active than the stereoisomer produced by OcfD against five of the *Candida* species tested. Another possible explanation for the observed differences in activity could be attributed to possible synergism between the configurational isomers produced by each cyclase thioesterase. Furthermore, the antifungal activity of the ocfN mutant (MS14GG88:  $8.79 \pm 0.38$  mm) was also significantly reduced ( $P < 0.05$ ) compared to wild-type activity (inhibitory zone radius  $\pm$  SEM:  $13.00 \pm 0.58$  mm) in an overlay assay against *Geotrichum candidum* (FIG. 8).

General Discussion. The findings from this study include experiments showing the following: the relative proportion of the Asn1 and BHN1 variants in the purified fraction; distinct differences in spin systems for the wild-type and ocfN mutant products; proposed model for the coordinated function of two cyclase thioesterases; and demonstrated differences in biological activity of wild-type and ocfN mutant products against therapeutically relevant *Candida* species. Expanding the conformational repertoire of cyclic peptide natural products can be beneficial to microorganisms. These data suggest that the bacterium *Burkholderia contaminans* MS14 is benefited by maintaining two distinct cyclase thioesterases that improves the spectrum of activity of occidiofungin.

Our data support the observation that cyclase thioesterase substrate recognition occurs prior to the catalytic transfer of the peptide. The presence or absence of a hydroxyl group on the beta carbon of the N-terminal amino acid (Asn1) appears to be important for the substrate recognition by the two cyclase thioesterases. It has also been shown that the N-terminal amino acid is important for substrate recognition for other thioesterases.<sup>4, 8</sup> It is possible that the presence of the hydroxyl group promotes a hydrogen bond with the ocfD cyclase thioesterase domain or more likely promotes an interaction within the T domain of the NRPS. Different bound orientations of the peptide to the T domain would establish a basis for the coordinated function of two cyclase thioesterases. It is also possible that the enzymatic conversion of one of the residues between L- and D-isomers is not completed by one of the epimerization domains. A combination of differences in the N-terminal amino acid and a possible difference in amino acid configuration (L or D), may contribute to the selective differences by the cyclase thioesterases that result in the formation of the observed configurational isomers.

The presence of the hydroxyl group on the beta carbon and the bound orientation of the peptide to the T domain may prevent the interaction of the OcfN cyclase, while enabling the continued substrate recognition by OcfD TE domain. There is evidence for the need of a bound orientation of the peptide to the T domain for the successful function of the cyclase thioesterase. Conformational diversity of the T domain has been shown to be important for the directed movement of the peptide substrate bound to the ppan cofactor and its interaction with externally acting enzymes.<sup>3</sup> More specifically, the active site serine of the cyclase thioesterase needs to attack the linear peptide attached by a thioester linkage to the ppan forming an acyl-O-TE inter-

mediate. The position of the peptide bound to the ppan in the T domain will be important for bringing the peptide substrate in proximity of the appropriate cyclase thioesterase.

Furthermore, some cyclase thioesterases are capable of transacylation of the peptide to the active site serine, when the peptide is bound to a biomimetic prosthetic group.<sup>4, 16</sup> However, there are several cyclase thioesterases that will not function when the product is bound to a biomimetic group. These data suggest that the interaction of the peptide with the T domain is important for the enzymatic activity of some thioesterases and this interaction cannot be mimicked using a prosthetic group. It is conceivable that the coordinated function of the two cyclase thioesterases, involved in the synthesis of occidiofungin, utilize differences in the interaction of the ppan bound peptide within the T domain.

Presumably, ocfN was integrated into the occidiofungin biosynthetic gene cluster to improve its spectrum of activity against fungi. Given the broad spectrum of antifungal activity associated with occidiofungin, the molecular target is likely to be highly conserved. However, there must be some variation among fungal species to account for the differences in biological activity. Increasing the conformational repertoire must be a selective advantage to the bacterium for it to maintain the two functional cyclase thioesterases. The microbial environment is considerably different than how we intend to apply the natural products produced by microorganisms. For instance, the bacterium *Streptomyces roseosporus* is a soil saprotroph responsible for the production of daptomycin.<sup>31, 32</sup> The microbial community that this bacterium encounters is far more diverse than the group of bacteria that cause human infection. Thus, evolutionary pressures that selected for the current conformers of daptomycin may not necessarily be the best conformers for treating a *Staphylococcus aureus* infection. It is very likely that the therapeutic application of daptomycin or other cyclic peptide drugs could be improved by engineering novel conformational or configurational isomers.

Creating novel diastereomers of other cyclic peptide drugs using new or engineered cyclase thioesterases may lead to improvements in their therapeutic activity against clinically relevant pathogens. This is true for occidiofungin produced by the bacterium *Burkholderia contaminans* MS14, which accomplishes this goal by the evolutionary integration of an additional cyclase thioesterase into the occidiofungin biosynthetic gene cluster.

All patents, patent applications, provisional applications, and publications referred to or cited herein are incorporated by reference in their entirety, including all figures and tables, to the extent they are not inconsistent with the explicit teachings of this specification.

Following are examples which illustrate procedures for practicing the invention. These examples should not be construed as limiting. All percentages are by weight and all solvent mixture proportions are by volume unless otherwise noted.

#### EXAMPLE 1

##### Experimental Section

Materials. Occidiofungin produced by both the wild type strain MS14 and the ocfN mutant MS14GG88 were purified as previously described for the wild-type sample. Chemicals were purchased from Sigma-Aldrich (St. Louis, Mo.) and were the highest grade, unless otherwise stated. Media were purchased from Fisher Scientific, enzymes were purchased from New England BioLabs, and primers were purchased from Integrated DNA Technologies (IDT) unless otherwise

stated. *Candida* strains used were purchased from the ATCC biological resource center and were a gift from Thomas Edlind (Drexel University College of Medicine).

Site Directed Mutagenesis. A nonpolar mutation was constructed in the open reading frame of wild-type *ocfN* by the insertion of a kanamycin resistance gene, *nptII*.<sup>33</sup> To mutate *ocfN*, a 1-kb fragment containing *ocfN* was obtained by PCR using primers *MocfNF* (5'-CGCCACCCGTTAC-GAGGATTC, SEQ ID NO: 1) and *MocfNR* (5'-ACGCGTC-CCCTCTCCTACG, SEQ ID NO: 2). The 1-kb PCR product was cloned into the pGEM-T Easy Vector System I (Promega Corporation, Madison, WI) resulting in plasmid pGG30. The *nptII* gene was inserted into the cloned *ocfN* at *SmaI*, generating plasmid pGG31. The ~2-kb *EcoRI* fragment of pGG31 harboring the *ocfN* gene disrupted by insertion of *nptII* was cloned into pBR325<sup>34</sup> at the *EcoRI* site to generate pGG32. Mutagenesis of the *ocfN* gene was conducted via a marker exchange procedure as described previously<sup>35</sup>, to generate the mutant MS 14GG88. PCR analysis and sequencing were used to verify the double crossover mutants. Production and purification of the antifungal were done as previously described.<sup>23</sup>

NMR Spectroscopy. A 2 mM sample of *ocfN* thioesterase mutant fraction of occidiofungin was prepared in dimethyl sulfoxide (DMSO-d<sub>6</sub>, Cambridge Isotopes) and data were collected as previously described for the wild-type fraction.<sup>22</sup> The NMR data were collected on a Bruker Advance DRX spectrometer, equipped with a CryoProbe, operating at a proton frequency of 600 MHz. The <sup>1</sup>H resonances were assigned according to standard methods<sup>36</sup> using COSY (correlation spectroscopy), TOCSY (total correlation spectroscopy), NOESY (nuclear overhauser effect spectroscopy) and <sup>13</sup>C-HSQC (heteronuclear single quantum coherence) experiments. NMR experiments were collected at 25° C. The carrier frequency was centered on the residual water resonance (3.333 ppm), which was suppressed minimally using standard presaturation methods. A 2.0 s relaxation delay was used between scans. The TOCSY experiment was acquired with a 60 ms mixing time using the Bruker DIPSI-2 spinlock sequence. The NOESY experiment was acquired with 400 ms mixing time. The parameters for collecting the HSQC spectrum were optimized to observe aliphatic and aromatic CH groups. The spectral sweep width for the TOCSY and NOESY was 11.35 ppm in both dimensions. The spectral sweep widths for HSQC were 11.35 ppm in the proton dimensions and 0 and 85 ppm for the carbon dimension. All 2D data were collected with 2048 complex points in the acquisition dimension and 256 complex points for the indirect dimensions, except for the HSQC which was collected with 2048 and 128 complex points in the direct and indirect dimension, respectively. Phase sensitive indirect detection for NOESY, TOCSY, and COSY experiments was achieved using the standard Bruker pulse sequences. <sup>1</sup>H chemical shifts were referenced to the residual water peak (3.33 ppm). Data were processed with *nmrPipe*<sup>37</sup> by first removing the residual water signal by deconvolution, multiplying the data in both dimensions by a squared sinebell function with 45 or 60 degree shifts (for the <sup>1</sup>H dimension of HSQC), zerofilling once, Fourier transformation, and baseline correction. Data were analyzed with the interactive computer program *NMRView*.<sup>38</sup> One-dimensional NMR temperature titrations were collected on the wild type and mutant peptides, using a Bruker AVANCE III HD 600 MHz spectrometer equipped with a cryoprobe. Eight scans were collected in each 1-D experiment, using 32K points, at a temperature of 298 K. The experiments were repeated using higher temperatures for both samples in 5 degrees K incre-

ments, up to a temperature of 323 K. 2-D TOCSY spectra were collected at a temperature of 323 K, using a mixing time of 60 milliseconds. Eight scans and 256 indirect points were used for both the wild type and mutant peptides. The 2-D spectra were processed using *NMRPipe*, with 45 degree sinebell squared shifts in both dimensions.

Mass Spectrometry. The wild-type occidiofungin and the *ocfN* mutant sample (10 μg) were evaporated to dryness in a Speed Vac Concentrator (ThermoScientific, San Jose, Calif.) and the residue was taken up in 50 μl methanol and analyzed by direct infusion at 3 μl/minutes into an LCQ DecaXP (ThermoScientific, San Jose, Calif.). Data were acquired over a mass range of m/z 200 to 2000.

In Vitro Susceptibility Testing. Microdilution broth susceptibility testing was performed in triplicate according to the CLSI M27-A3 method in RPMI (Roswell Park Memorial Institute) 1640 [buffered to a pH of 7.0 with MOPS (morpholinepropanesulfonic acid)] growth medium. 100x stock solutions of occidiofungin were prepared in dimethyl sulfoxide (DMSO). MIC endpoints for occidiofungin were determined by visual inspection and were based on the wells that had no visible growth (an optically clear well) after 24 hours of incubation. DMSO containing no antifungal agent was used as a negative control. Colony forming units (CFUs) were determined in triplicate by plating 100 μl from the MIC wells onto a Yeast Peptone Dextrose (YPD) plate as well as plating 100 μl from 10-fold serial dilutions of the cell suspension in Yeast Peptone Dextrose (YPD) Broth. Colony counts were performed and reported as CFUs/ml. Time-kill experiments were performed as previously reported.<sup>19</sup> *Candida glabrata* (ATCC 66032) colonies on 24-h-old YPD plates were suspended in 9 ml of sterile water. The density was adjusted to a 0.5 McFarland standard and was diluted 10-fold with RPMI 1640 medium to a final volume of 10 ml containing a final concentration of 2, 1, 0.5 and 0 μg/ml of occidiofungin from wild type strain MS14 and the *ocfN* mutant MS14GG88. The cultures were incubated at 35° C. with agitation. Samples were drawn, serially diluted, and plated on YPD medium for colony counts.

It should be understood that the examples and embodiments described herein are for illustrative purposes only and that various modifications or changes in light thereof will be suggested to persons skilled in the art and are to be included within the spirit and purview of this application and the scope of the appended claims. In addition, any elements or limitations of any invention or embodiment thereof disclosed herein can be combined with any and/or all other elements or limitations (individually or in any combination) or any other invention or embodiment thereof disclosed herein, and all such combinations are contemplated with the scope of the invention without limitation thereto.

TABLE 1

Chemical Shift Values for Occidiofungin derived from the <i>ocfN</i> mutant MS14GG88 <sup>a</sup>			
Unit	No.	δ <sub>C</sub>	δ <sub>H</sub>
Asn1	2	52.71, CH	4.59
	2-NH		7.75
	3	39.91, CH <sub>2</sub>	2.62, 2.41
	4	—	
BHN1	4-NH <sub>2</sub>		7.39, 6.93
	2	58.47, CH	4.66, 4.61
	2-NH		7.81, 7.9
	3	75.01, C	3.98, 4.02
	3-OH		4.66
4	4	—	
	4-NH <sub>2</sub>		7.24

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TABLE 1-continued

Chemical Shift Values for Occidiofungin derived from the oclN mutant MS14GG88 <sup>a</sup>			
Unit	No.	$\delta_C$	$\delta_H$
NAA2	2	43.88, CH2	2.34, 2.36
	3	47.25, CH	4.23
	3-NH		7.31, 7.34
	4	41.57, CH2	1.39, 1.76
	5	66.36, CH	3.50
	6	76.07, CH	3.08
	7	79.61, CH	3.72
	8	33.19, CH2	1.54
	9-17	25.14-28.02, CH2	1.27
	18	16.94, CH3	0.86
Ser3	2	58.59, CH	4.07, 4.15
	2-NH		8.11, 8.14
	3	70.23, 64.29	3.49, 3.45
3-OH			4.95
BHY4	2	58.71, CH	4.06, 4.15
	2-NH		7.83, 7.94
	3	73.75, CH	4.98, 5.08
	3-OH		5.66, 5.73

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TABLE 1-continued

Chemical Shift Values for Occidiofungin derived from the oclN mutant MS14GG88 <sup>a</sup>				
Unit	No.	$\delta_C$	$\delta_H$	
5	4	—		
	5, 6	—	7.15	
	8, 9	—	6.67	
	DABA5	2	53.49, CH	4.43
	10	2-NH		7.66
		3	32.68, CH2	1.88, 2.11
		4	39.17, CH2	2.92
		4, NH		7.71
	Gly6	2	44.76, CH2	3.87, 3.58, 3.84, 3.70
		2-NH		7.68, 7.85
15	2	53.25, CH	4.51, 4.58	
	2-NH		8.35, 8.41	
	3	40.03, CH2	2.61, 2.38	
	4	—		
Ser8	4-NH2		7.39, 6.93	
	2	58.11, CH	4.33, 4.32	
	2-NH		7.76, 7.78	
20	3	64.59	3.61, 3.62	
	3-OH		4.79	

<sup>a</sup>Proton chemical shift values are from a TOCSY and NOESY experiments. Chemical shifts in brackets are <sup>13</sup>C values from the HSQC experiment.

GenBank: EU938698.5

Go to:

LOCUS EU938698 58101 by DNA linear BCT 13-DEC-2010

DEFINITION *Burkholderia contaminans* strain MS14 putative FAD linked oxidase domain protein gene, partial cds; and putative LuxR-type regulator (ambR1), putative LuxR-type regulator (ambR2), putative cyclic peptide transporter, hypothetical protein, putative glycosyl transferase, putative nonribosomal peptide synthetases, putative beta-lactamase domain protein, putative beta-ketoacyl synthase nonribosomal peptide synthetase, putative short chain dehydrogenase/reductase SDR, putative beta-ketoacyl synthetase, putative taurine catabolism dioxygenase, putative transaminase, putative epimerase/dehydratase, putative thioesterase, and hypothetical protein genes, complete cds.

ACCESSION EU938698

VERSION EU938698.5 GI: 314950578

KEYWORDS .

SOURCE *Burkholderia contaminans*ORGANISM *Burkholderia contaminans*

Bacteria; Proteobacteria; Betaproteobacteria; Burkholderiales; Burkholderiaceae; Burkholderia; Burkholderia cepacia complex.

REFERENCE 1 (bases 1 to 58101)

AUTHORS Gu, G., Smith, L., Wang, N., Wang, H. and Lu, S. E.

TITLE Biosynthesis of an antifungal oligopeptide in *Burkholderia contaminans* strain MS14

JOURNAL Biochem. Biophys. Res. Commun. 380 (2), 328-332 (2009)

PUBMED 19167363

REFERENCE 2 (bases 1 to 58101)

AUTHORS Gu, G., Wang, N., Chaney, N., Smith, L. and Lu, S. E.

TITLE AmbR1 is a key transcriptional regulator for production of antifungal activity of *Burkholderia contaminans* strain MS14

JOURNAL FEMS Microbiol. Lett. 297 (1), 54-60 (2009)

PUBMED 19500142

REFERENCE 3 (bases 1 to 58101)

AUTHORS Gu, G., Smith, L., Wang, N., Wang, H. and Lu, S.

TITLE Direct Submission

JOURNAL Submitted (01-AUG-2008) Entomology and Plant Pathology, Mississippi State University, 32 Creelman St., Mississippi State, MS 39762, USA

REFERENCE 4 (bases 1 to 58101)

AUTHORS Gu, G., Smith, L., Wang, N., Wang, H. and Lu, S.

TITLE Direct Submission

JOURNAL Submitted (12-NOV-2008) Entomology and Plant Pathology, Mississippi State University, 32 Creelman St., Mississippi State, MS 39762, USA

REMARK Sequence update by submitter

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REFERENCE 5 (bases 1 to 58101)  
 AUTHORS Gu, G., Smith, L., Wang, N., Wang, H. and Lu, S.  
 TITLE Direct Submission  
 JOURNAL Submitted (15-JAN-2009) Entomology and Plant Pathology,  
 Mississippi  
 State University, 32 Creelman St., Mississippi State, MS 39762,  
 USA  
 REMARK Sequence update by submitter  
 REFERENCE 6 (bases 1 to 58101)  
 AUTHORS Gu, G., Smith, L., Wang, N., Wang, H. and Lu, S.  
 TITLE Direct Submission  
 JOURNAL Submitted (24-FEB-2009) Entomology and Plant Pathology,  
 Mississippi  
 State University, 32 Creelman St., Mississippi State, MS 39762,  
 USA  
 REMARK Sequence update by submitter  
 REFERENCE 7 (bases 1 to 58101)  
 AUTHORS Gu, G., Smith, L., Wang, N., Wang, H. and Lu, S.  
 TITLE Direct Submission  
 JOURNAL Submitted (13-DEC-2010) Entomology and Plant Pathology,  
 Mississippi  
 State University, 32 Creelman St., Mississippi State, MS 39762,  
 USA

REMARK Sequence update by submitter  
 COMMENT On Dec 13, 2010 this sequence version replaced gi: 224016442.

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CDS 2480 . . . 3301

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CDS complement (26061 . . . 29981)

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LIVEPASGCAEPVPLLDAGEPGPTVTRAYLNQLRDSVQRSYSYQDFPIAALAHKLHGER

RATNVGVRFDGLHEAWAAADYDLSIEIRHREYIEIVLTGRQTVFTLHYLQHVARHLRN

VVAGFGALDAPLDTVSLDDEERARLRSHAAFVAVQGTFLQFAQRVAAAPDSVAVVT

ADASLTYAELDDQASRLASFLLAEYAIERGDVVGVDADRSEWIVGMLGALKAGAVYL

PLDPEFPRERLRFMIEDAKVKALLTHSEHLPLLDWFVAIPMFALDQLDRTLAPASASA

QVEVRPDDAAYIIYTSGSTGVPKGVVLEHAGLLNMAQYHVDAFGFDSDRFRVQFYSYG

FDGIMEIVFTLLAGARLVLAKTAVIRDVPRFVDYIAQQGVTTVNATPAYLAALDWAH

LGAVKRVI SAGDSARVADLRELARTRTCHNSYGPTEATVCIADYVVDPAITYGARLPV

GRPIHNTHLVLLDEHGALAPEGCAGEICVSGIALARGYVGRDDLTAAAFVAHPFEAGE

RLYRTGDLGVWLPDGNLEVTGRRDTQVKIRGYRIEMGEIEAALRQHAGVADAI VVRE

DTPQHKQLVACVATATASVASLREHLKERLPEFMPASIVTLERLPLTPNGKPRKAL

AALELAFAPSETAYVAPANDVEARLGRICDVLGREPIGVHDNPFELGGDSILIIQVM

SLAQVGLKFTADQFFAHTPIAELAQVATEAPSIIRIAQEPVVGPAPLTPIQHWFFAQD

VADPHHYNQSTMIEVPASLRPDTIERALAAVATHHDALRLSFACVAGVWQQSHAAPPL

AIPLGVTSLADAAPAARQAAMLATATGMQESFTLSAPPLRAHLFQFGPDAPQRLAV

AHLVIDGVSWRILFEDLYTACRQLEAGDAVQLPARTTAWRDWSTRLSGLGATALDGL

GLDYWLQGNAGEPACFDDMPAGTVAEAGSTIVEFDAQQTALLQDVPRAFNTQINEVL

LTALLLAFGDWTGNASLVVDLEGHGREDIFDGVDTSRTIGWFTTHYPVCLNAGDATVA

VDALRHVKEQLRAVPMRGLGYIARYLGEDAGIAAALERQPPAPVRFNYLGQVDRVLP

DDTGWKPVLDFQSPHESPRARRGHLFEIDGMVFDGRLRLTWHYNREACAPGVIEQLTQ

CYRSRLLSIVAAGGDGPRALSPDFPAARISQEALDALVSRIS"

CDS complement (29969 . . . 31585)

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/note = "ORF8"
/codon_start = 1
/transl_table = 11
/product = "putative beta-lactamase domain protein"
/protein_id = "ACN32488.1"
/db_xref = "GI: 224016446"

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(SEQ ID NO: 14)

/translation = "MTISSAQVYLRQNIQFEPLINSWYAWYHTLPLPLTAALNVAERF

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AYKAFSTLLLERATGMASDPLYPEIPEVLKGYVEIYYDLNHNPSFRVFESLLYASPFY

ARDAQSIALSAIEEHTPRPFILSTPRLRDERTVFSNMAFDDRALDTLFRMRDTFGSYA

KIVDLMRVEEKDEPLFRSFFVEEAPAPKPDERSFDGDDIRIRIYGHACVLIQSRGVSIL

IDPVISYGYDTALPRYTFADLPDQIDYVLI THSHDHIVLETLQLRHVKVTVVVGRN

LDGFPQDPSMELALRKLGFDDVLEVRDAQEIKVPGGAI TAI PFMGEHNDLAIHSKQSF

MIRFGSRSVLCIADSCNLDPRLYEHVFRLAGKPDTLFVGMETEGAPPSWVYGPLFPKA

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LPRDIDQSRRRARGCQFGEAAALVDDFAFNAAVYVYAMGQEPWLNHLLDNTFDENSPSHI  
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CDS complement (31596 . . . 45005)  
/note = "ORF9"  
/codon\_start = 1  
/transl\_table = 11  
/product = "putative beta-ketoacyl synthase nonribosomal  
peptide synthetase"  
/protein\_id = "ACN32489.1"  
/dp\_xref = "GI: 224016447"

(SEQ ID NO: 15)

/translation = "MNAKATHALKAALDELRLRRAETIAALRSRNEPIAVIGMACRFP  
GRSDTPDAFWQLLDGAHDAVTEVPGERWDIDRYDDPDPSTPGKMATRHGAFLEVRDQF  
DAAFPGIAPREATYLDPPQRLLEVAWEALENAHLAPERFRQSATGVYVGI TCFDHAI  
QVSNASMPSSSYAGTGSALNMAAGRLSPVLGLTGPSMAIDTACSSSLVCLHLACESLR  
SRESNMALAGGVNMLSPVMSVFSQARMLSPDGRCKTFDAAADGYVRGEGCGMVVLLK  
RLDALADGDRVLGIVRGTAVDQGGAGGLTVPSRDSQERVIRRALNQAAGLAPGDVSY  
VEAHGTGTSGLDPI EVEALAGVYGPGRANEPLVIGSVKTNIGHLESASGIAGLIKVL  
LSFEHDRI PAHLHFTQPNPHTPWQDIP IRVAADPVAVRRGERRR IAGVSAPGFSGTNA  
HAIVEEPPVAPAHAAQRALLLSARSEALAALVPRYERAIAGATPQELAAICRAAAT  
GRSHYPFRAAYVSGARVASAAPRTGKALRMGFGVGPDTGVAHALHASEPLFRDAFA  
RCSVPLDALET DAGRFAIQFAWELWKGWGLRPAVVS GHGIGEYVAACVAGVVSVA DA  
LRLVAARSDAEALRAVLRDMP LARPSVRLISGYLGTDV TDEVTHPQYWLQLAGASDQA  
DASHPPEGLADGWLPPPCAGDALERALAALYVQGAQFDWRALFPAPAQPATTLPNYPF  
ERQRFSLKIPSPIVGMDAGS IDAALRHLKSSGKYPEDMLNAPDLLRTAFAAAETVA  
SNAHPLYHVWVEQQAAMPAPAAADAS PWLIFADASGVGERLAALLRARGASC SLVRP  
GIDYVTGAEAGWQVAPERPD FVRLLNETAASGQRIVFLWALDEAVGETRMSAALLHL  
VHALVGSEREWTPSTRPRI SVVTRDAVEAGEAPHVSGLAQAALSGLARGAMIEHPPEWF  
GTAIDLDPAAPENETQALLQEMLGESREEQVALRHGARHVARLSPLAPAETAALPVDP  
DAAYLITGGFGALGLHTARWLAARGAGTLILVGRQGAASDESQRAIAELRERNVTLRC  
ERLDIADPAAVAAFALRRDGVPLKGI VHAAGIVGYKPI MQVERDELDAVLQPKVAG  
AWLLHQQSEHFPLDFFLFFSSIASAWSREQAHYSAANRFLDALAHHRRGQGLPALS  
V  
NWGPWAEAGMTFPEAEALLRRV GIRS LAADRALDVLNRLPAVPQVAVVDIDLALFQGS  
YEARGPKPFLDHRVAKSAPSAPAMPALSDASPRERKRL LADS IDRAVAQVLGYDAGT  
LDRDLGFFEMGMDSLMALDVR THLENALGIPLSVALLFDHPTVNALADFLAEQASGTA  
QAQTVPQQQPRPIAPAI EARDAGTPEPIAIVGMSCRFPGAHDLDAYWNLNDGVDA  
ISEVPRERWDVDAYDPDPEAPGRMYSRFGGFLDDVDQFDPAPFRITPREAAAMPDQQ  
RLLLEVSHAELEHAGIPVDSLKGSRTGVFVGI TTNDYANLQLRNGGSGIDGYFFTGN  
PLNTAAGRISYGLGVQGPSMAIDTACSSSLTAIHTASQNLRSGECDLAIAGGVNLI LS  
PDNSIAVSRTRALAPDGRCKTFDAAADGFVRS EGCGALV LKRLSDALAAGDRVLA VLR  
GSAVNHDGASSGFTAPNGRAQEAVIRQALGGLPAASIDYVEAHGTGTPLGDPVELQAL  
ATVFGAGRDAGRRLRVG SVKTNIGHTESAAGIAGVIKVVLSLNHDRLPAHLHFRQPS  
LVQWDAMPVEICAEASAWPRGERPRRAGVSAPGASGTNAHLVLEEAPAPARQATPSRH  
KVHPLVLSAKTPAALRELAGRYQRRLEAEPGLDIAAVAFSAATGRSHFAHRLAWPVTS  
LDDAIDKLR AFHAKEPAGAAQPAPRVKMAFLFTGQGSQYAGMGRRLYDAYPVFRDAID

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RCRAVADPLLDKPLLEVLSAQGEDIHQGTGYSQPALFSLQYALTTLLASFGVVPDAVMG  
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 HEVAVAAVNGPASI VISGKRERIAMLVDFAAARDIRSVPLNTSHAFHSPLEPMLDSF  
 QLAAKTVPPARPAIFPYSNLTGAVMDEAPTDTYWRRHCREPVQFASSVERLAEAGFNV  
 LVEIGPKPVLVNLARACCAPDAGIQFLALQRPQVEQQALIE TLSSLYARGVDVDWAPT  
 ETPAPARIALPSYPQFSRTWFQKADTSMTQTSASPIAAAPTHNRSGETLEWLGRKIG  
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 TVERVLREQNQLLSHVMSQQMELLRTSLTGQPGVRPATAAVQAVASTASVAPKAASAA  
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 RTRKSKDSVQASRPVLADSRATVGFRRFSTKEMLYPIVGDRAAGSRLWDIDGNEYIDFT  
 MGFGVHLFGHTPDFIQQQVTREWQRPLEL GARSSLVGEVAARFARVTGLDRVAFSNTG  
 TEAVMTAMRLARAVTGRDKIVMFTHSYHGHADGTLAAAANAEGVTETIAPGVFPFSGVEN  
 MILLDYGSDAALEAIRGMAS TLAAMVVEPVQSRNPSLQPVAFKELRRI TEEAGVALI  
 FDEMITGFRVHPGGSQAMFGIRADLATYGKI IGGGLPLGVIAGTSRPFMDAIDGGMWY  
 GDHSFPAADR TAFGGTFCQYPLAMAAALAVLEKI EQEGPALQAALNERTAQIAGTLNA  
 FFAEAEAPIKVTFWFGSMFRFEFTENLDLFFYHMLEKGIYIWEWRT CFLSTAHTDADID  
 RFIRAVKDSVADLRRGGFIRPHSKHGTVAALSEAQRQLWVLEIDPEGSLAYNVNTTL  
 ELNGLRLEAAMRAAVQSLVDRHEALRTTVMADGSGQIVHPSLTLEIPLIDTDPNAWRE  
 QESRQFPDLVNGPLFRAALVRLGSRHLLVMTAHHIICDGSTFGVLEDLARAYAGAA  
 PADAPLQFRAYLQKLDGQRHS PETKANREYWLAQCARQAAPLNL PVDYPRPAVKTFHG  
 ERVSLHLDAATAATLRTAARQNGCTLYMVLLAGFNLFLHRVAGQQEIVTGI PVTGRSV  
 AGSDRLAGYCTHLLPLHSTLPEQATVASFLAGTRQNLLDALEHQDYFPAELVREIGAQ  
 RDLNAAPLVSAVFNLEPVSALELPGLTVGLVAPLIRHTAFDLNVNVL DQALLIDC  
 DYN TDLFDASTVQRFLDIYRLLTHLADDAASAARLP LSSDAERNLLTVEWNRDTD  
 FGEDAAQPLHRLFEQQVERTPDAVAIVFDDTALTYAELNLRANRLAHHLVALGVGPDS  
 LVGVAMERSLDM SVALLAI LKAGGAYVPVDPDYPAERVRFMIDHAQLRWLLTQQLHLD  
 ALPDTDAHVI VVDRDSLDDAAATSNPAPALNGDNLAYMI YTSGSTGRPKGALNTHRA  
 ITNRILWMQHAYALGADDAVLQKTPFSFDVSVWELFWPLVVTGARLVFARPGGQRETDY  
 LVELIERERITTIHFVPSMLRAFLDHPDLDAH CASLRRVVCSGEALPHDLQQRCLERL  
 DVELYNLYGPTAAAVDVTAWECRRDDPHRIVPIGRPIANTRLYIVDAQMQPTPIGVAG  
 ELLIGGTPVGRGYHGEPELSAEKFIADPFSADFLARLYRTGDLARYRDPGNI EFLGRI  
 DHQIKLRGLRIEPGEIEAALTSHP LVDAAVVALRGVDDGARLVGWLCS SHPEAELIEA  
 VRGHLRQLRPDYMVPSAFVVVSAFEHL PNGKLDRTLRLEPGDGLDHVAPVNALEAQLA  
 AIWQEVLGQARISTTGNFFDLGGNSLLATKVVVARIRRD LHVKLEIRSLFALPTISSLA  
 KRIADTQPIDYAPVTPLPAQASYALSPAQTRLWVQDR LHAAQAEGLPTSLLFEGVLD  
 VDALVRAFRLASERHEILRTRFVLEGNQPVQHVLPPGEAAPVEIVDLQDAEDRDAQA  
 AAIQASERLVPMDLATGPLFRVKLLRLESEVRHVCLCTMHHIVSDGWS TEVLLDDLSAL  
 YDAFVQRRDDPLPALPIQYKDYAGWLNRLLAGPDGARMKDYWLTKLGGGLRAL ELPGD  
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GTPVAGRELPELESQVGPYLNVLALRDRVAGDDRFDTLLTRVRDTTLEAFSHPLYPLD
RLLELHI KR VAGRNPLFDIGLTLQNQRHGPVDRYAGQVHIAELPDHDPQRADTEAAT
DFWFLAEPHAEGLAIRVVYHAGRFSEALVQGLANELTSVIGEVLANPGVRI RNLT LGQ
RALHAEARQPTVELSAF"

CDS complement (45002 . . . 48325)
/note = "ORF10"
/codon\_start = 1
/transl\_table = 11
/product = "putative short chain dehydrogenase/reductase
SDR"
/protein\_id = "ACT64845.1"
/db\_xref = "GI: 314954101"

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/translation = "MKFGLMFFASSEALSGNKYQLVMESARFADANGFSSVWVPERH
FTEFGSLYPNPAVLHAALAAATQRVKLVAGSVVAALHNPIRIAEEWSMVDNLSNGRVG
VSFASGWNPDFFVAPDKYATRQDDMLTTRAVQHLWRGGTLDATNGVGKPVRLRVYP
TPVQPELPVWVTAASNPTFVRAGEAGANLLTHVLDQDRDQLAHKIALYREARAKHGF
DPAAGTVSVMHLTFVGDAAQAREQARVFCNYIRSNIGLLNGLAQSRGQSVDRAMG
ARELDEFVEFLYERFAQSRGLIGTPETCVELVRDLESIGVDEVAQLLDFGPPVERILG
NLPQLRRLREMCAPRRSAAPTRFDAAEVQARCTETTSGADFNGEIRQHGVIQIDGVFDA
IRQIWRRTGEALGKISLPADALASSPYQVHPAFLDACSRVLA A AIDPALESGDLYLP
SSIGAVRVHQPPASTEAWSHATLRTPIGQGALEGDIVHDLA GRLLI EIDALRLQQVR
AARAVERHDF A ALLYQRVWRPSNVDAATGGS AHGEWLI LADRGVGAQLSALLEAHGD
TCTLRFADATPELPAADRPLKGV IHLWSDL LAPS DIAARRRASASV LHLVRLASRAP
SARQARLWLVTSGAMNVL DGESIAVAQAPLWGLGRAIAVEHAALWGGLVDLDPEQPSA
ADIMQAVQAGGREDMIAFRDQHYVARIARDNREYVSHRPIRFHG DATYLV TGGLGGL
GLRLASWLADNGAGKIVLLGRGEP SAAAGKILRTLDARFIRADLSRREDVGQALGEIA
HSMPPLKGI FHLGALDDALLTRQDDFFHRAGSGKADGAWYLHELTAGLP LDHFVLF
SSMAALITMFGQGN YAAANSFLDALAQHRAQ GK PGLSVNWGFWAEIGHAATDYGRRA
HEQLGALGVGTLPPELAIATLERLMASGVAQSGVARIDWPTLFRVDAPAAGSALFSEL
TQPAQAQPAQOETALLRQLHACAPRERVERITDTLAAMLAETLRLSGPDIAIPEQSLLD
LGLDSLVALELTDRLTKVFRPFRATLFFSYPNLQTLAQYVLNLSPLPAPVVDEAS
DDLDEDDLSELIAQEIGAQ"

CDS complement (48322 . . . 52749)
/note = "ORF11"
/codon\_start = 1
/transl\_table = 11
/product = "putative beta-ketoacyl synthetase"
/protein\_id = "ADT64846.1"
/db\_xref = "GI: 314954102"

(SEQ ID NO: 17)

/translation = "MLPDKERTVTEILLFRGKVEPEKTAFI FLENGEAELTRLTFGD
LDKRARGIAARLQAIAPGDRVLLVYPPGLEFICAWVGCLYAGLIGVPAYPPRRHRPA
DRLKAI VADATPVVALTDAATLDGIAH HADGYSDTLELKI LATDQRFDAPAEQWRAPD
ITPQTLALLQYTS GSTGTPKGMISHANILSNMAVIAEASDADASTV FVSWLPV FHD M
GFFGKVL LPIYLGVL SVLMAPAAFVQKPV RWLQAITKYRGTHCAAPDFAYDL CARKIA
DEARAQLDLS SWRVAFNGAEPVRAESVARF SRAFAACGFHAHTMRPVVGM AEATL F IS
GQPARS LPRVADYDADALAQGVATRNDSGKRHALVSCGRTWAEHRVRI VNPDTGERCA

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PGRIGEIWLTPGSVGVYWNRIIDETERTFRAKLDGDDARYLRTGDLGDFVDGEDLFVTG  
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 NTLDAEAVAAEIRHTLAEVHDVDLYAAVLLKPATILRRTSSGKIQRSRIRQAFLDEQGL  
 AIAGEWRRRAFSAPPAPPQTAEPRDTQALVQWCIERVSRLSGIASGKIDPDAPFSVHGL  
 DSKDAIMLSGELQDWLGRPVSPVVYDPPSISLLARHLSGTGSAMPDQAPGSAEARAD  
 IAIVGMGCRFPAGNPDADFQWLLLEGRDAVGAATQRAADLPLAGLLDQVDFDAAFG  
 ISAREAESMDPQORLLEVAWETLEHAGIAPRSLAGGR TAVIVGINSNDYIRLAQDEV  
 ADVGPYVATGNALSVAANRISYALDLRGPSSWAVDTACSSSLVAVHQACRALQRGESDA  
 ALAGGVNLLILAPQLSASFTQAGMLSPDGRCKAFDAAANGYVRGEGVGMVLLKRLDDAL  
 ENGDTVFAVIRGSAVNQDGRSNGLTAPNGPAQQAVIHGALRDAGVRAQDIGFVETHGT  
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 AIPPNLHFRSINPQIALDGTFFRIPRQVTPWNSEHGPRLAGVSSFGFGGTNAHLILSE  
 APGLPEIEAEPVAPAARVVTL SARTPDALQALAAASYAAYLDAHPEAGVRDVAF TANTG  
 RTHFTQRAAIVAPSRDSLRAQLDSVSSGEP AETPPAVTFHFCADDGASADAVRQLRAA  
 SPAPDALMQRQSDASGAPALAPDEAGFTRFQRALQWLWMSFGIAPDAVSS TGDGQRAA  
 AAWAGVPQAPDSGAAGHPGIVIDIGAHTAAWDAILHTLAALYVRGASIDWDAVEQGAP  
 HRRALPTYPFERRGFWRPHARRHPLLRRLMEQHAHAPATWIWQSRLDAPATNFLDG  
 HRVKGSPVLPYSAFVEMALSATSEIGAAGHTTLKDLALHAPLPLHPHESHTVQTVLSR  
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CDS complement (52936 . . . 53922)  
 /note = "ORF12"  
 /codon\_start = 1  
 /transl\_table = 11  
 /product = "putative taurine catabolism dioxygenase"  
 /protein\_id = "ADT64847.1"  
 /db\_xref = "GI: 314954103"

(SEQ ID NO: 18)

/translation = "MEGMTERKLLAEGSTPWLLEPVSNGRDLAQAVNDNRRALESRL  
 EHGVLFRGEDVSSVGGFEAFARAI SAHQSDYVYRSTPRTSISNGIFTATEYPPSETI  
 ALHCENAYQRSWPLRVAFCC LTPAATGGETPIADMREVSRRI GPRILDHFEARQVRYV  
 RHYRRHVDIPWETVFQTS DRNQVAAFCADNGIALEWLDLDDTLRTAQINQGVAYHPVTG  
 ERVFFNQAHLFHISNLEASLASSIVSLFGEDRIPRNACHGDGSPFDLADLEQIRHAFR  
 ECAITFPWQRGDVLLVDNMRFAHGRNPFEGERKVVVSLDPYTPDIEGIADR"

CDS complement (53999 . . . 55369)  
 /note = "ORF13"  
 /codon\_start = 1  
 /transl\_table = 11  
 /product = "putative transaminase"  
 /protein\_id = "ADT64848.1"  
 /db\_xref = "GI: 314954104"

(SEQ ID NO: 19)

/translation = "MKRFSCASVHQALQAGSARMEKLEYLKVESNARTYATSFPR  
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 AKHAFVEQLFSLLPKIAESGKI QFCSPSGADGVEAAI KLTRHYTGRPTIMAFHGAYH  
 GMTSGALAASGNLTPKSAGGNRVDVHFLPYPYAFRC PFGTDGSATDQLSINYIRTVLS  
 DPESGITKPAAIVEVVQEGGCIPAPD TWLIELRELT LRHEIPLIVDEVQTGLGR TG  
 ALFAIEHSGIRPDVLVLSKAFGGGYPLSVVYDERLDTWPPGAHAGTFRGNQIAMVAG  
 LSTMRIVEREDLSAHADRVGKLLVAGLEELAERFPCLGQIRGRGLMIGAEVVVPGTHG

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RAGPPHTERARAIAKQNCRLRNLIVETGGRNGAVLRFLLPPLIVSEADIHDILNRFEHAV

ETACRA"

CDS complement (55516 . . . 56466)

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/note = "ORF14"
/codon_start = 1
/transl_table = 11
/product = "putative epemerase/dehydratase"
/protein_id = "ADT64849.1"
/db_xref = "GI: 314954105"

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(SEQ ID NO: 20)

/translation = "MQRNRKRILVTGGAGFLGSHLCERLVELGHDVLCVDNYFTGTKQ

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DYHRQQNVRIKVVRIFNNTYGPRMHPNDGRVVSNFIVQALRGEDITLYGDGSGTRAFICY

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CDS complement (56622 . . . 57341)

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/note = "ORF15"
/codon_start = 1
/transl_table = 11
/product = "putative thioesterase"
/protein_id = "ADT64850.1"
/db_xref = "GI: 314954106"

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(SEQ ID NO: 21)

/translation = "MRLICFPYAGGSAAVYRTLQASLPGIEVCRHELAGRGSRLSEPA

VRDMATLVDTLLCDLDDCFDRPFALLGHSMGAAIAELALRLPAHARPNLRHLFVSAR

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CDS 57710 . . . 57997

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/note = "ORF16"
/codon_start = 1
/transl_table = 11
/product = "hypothetical protein"
/protein_id = "ADT64851.1"
/db_xref = "GI: 314954107"

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(SEQ ID NO: 22)

/translation = "MQHRQKAVPTQQVANERVIVTEWRFPAGAEWGWHVHRHDYVVVP  
QTDGQLLETAQGNRESQLHAGRSYAGLKGVEHNVVNATDHEVVVFVEVEIL"

ORIGIN

(SEQ ID NO: 23)

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1 aattcctgca gcacggtgcg cgaccagccc cagatgtccc cgctgagcgt gagtgcgaga
61 ccggccgctcg tgatggccag ctgctgtctgg ccgaacagcg gcgtcaatgc gccttcgccg
121 ccgatcacga tccgcttgac gagatccgag atggactgcg agatcgaatc ggagaacgga
181 tagttgtacg gctgctgac ggccgcccag aggaacggct tgctgggctg cggcgtccag
241 accttgagcc acggcttggt cgtgaacggg aaccagatgg cttccacccg gcccgagccg
301 tcgagaaacg atgcgatcgt gggcccctc gtgcccggcg cggcgaacag ttcggaggcc
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541 accacgagat tgctcagcga gccgtaggta tggcccgggt gcaaggtttc accggcccgcg
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 961 tgcacggcga tctcgcocgt ccagttcagg aacgcctgct tgtaaagctg gatgtcggcc  
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 1201 ttttgcatg tggaagacg acaaatgacg cgttgagact cgtgtggcaa ttcgagcagg  
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 Ala His Leu Ala Tyr Val Leu Tyr Thr Ser Gly Ser Thr Gly Lys Pro  
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 Lys Gly Cys Leu Leu Glu His Arg Asn Leu Ala His Tyr Ile Ala Trp  
 405 410 415  
 Ala Ala Gly Tyr Tyr Phe Pro Glu Ser Thr Thr Gly Ser Phe Gly Leu  
 420 425 430  
 Tyr Ser Ser Leu Cys Phe Asp Phe Thr Leu Thr Asn Ile Phe Cys Pro  
 435 440 445  
 Leu Val Arg Gly Lys Thr Leu Arg Ile Tyr Pro Gln Ser Glu Ser Ile  
 450 455 460  
 Asp Thr Ile Leu Ala Arg Met Phe Gln Pro Gly Ser Gly Val Asp Thr  
 465 470 475 480  
 Leu Lys Leu Thr Pro Thr His Ile His Leu Leu Glu Tyr Met Asn Leu  
 485 490 495  
 Ala Arg Ser Gly Val Arg Lys Val Ile Val Gly Gly Glu Glu Leu Thr  
 500 505 510  
 Pro Gln His Ile Ala Thr Leu Arg Lys Ile Asp Pro Ala Ile Glu Ile  
 515 520 525  
 Tyr Asn Glu Tyr Gly Pro Thr Glu Ala Thr Val Gly Cys Ile Val Glu  
 530 535 540  
 Arg Val Glu Asp Ala Pro Pro Thr Val Leu Ile Gly Arg Pro Ile Ala  
 545 550 555 560  
 Asp Thr Arg Val Tyr Met Leu Asp Asp Ala Leu Arg Pro Val Pro Leu  
 565 570 575  
 Gly Val Pro Gly Glu Ile Cys Leu Ala Gly Ala Gly Leu Ala Arg Gly  
 580 585 590  
 Tyr His Gln Arg Pro Asp Val Thr Ala Ala Lys Phe Val Glu His Pro  
 595 600 605  
 Phe Pro Gly Glu Ala Arg Ile Tyr Arg Thr Gly Asp Ile Gly Arg Trp  
 610 615 620  
 Leu Pro Asp Gly Arg Ile Gln Cys Tyr Gly Arg Val Asp His Gln Val  
 625 630 635 640  
 Lys Ile Arg Gly His Arg Val Glu Leu Gly Glu Ile Glu Ala Ala Ile  
 645 650 655  
 Ala Ala His Glu Asp Val Val Gly Ala Ala Val Met Leu Arg Glu Ser

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660					665					670					
Ala	His	Gly	Val	Arg	Lys	Leu	Ala	Ala	Tyr	Val	Lys	Gly	Ala	Ala	Ser
	675						680					685			
Leu	Ser	Val	Pro	Asn	Leu	Arg	Ala	Tyr	Leu	Ala	Gly	Lys	Leu	Pro	Asp
	690					695					700				
Tyr	Met	Val	Pro	Ser	Asp	Ile	Ile	Pro	Ile	Ala	Glu	Phe	Pro	Leu	Asn
	705					710					715				720
Ala	Asn	Gly	Lys	Leu	Asp	Arg	Pro	Ala	Leu	Leu	Ala	Leu	Glu	Pro	Ala
				725					730					735	
Ala	Ala	Pro	Glu	Glu	Ala	Pro	Leu	Asp	Ala	Thr	Pro	Ile	Gln	Arg	Glu
			740					745					750		
Leu	Val	Arg	Ile	Trp	Arg	Asp	Val	Leu	Asp	Asn	Pro	Ala	Val	Asp	Leu
		755					760						765		
Ala	Gly	Arg	Phe	Phe	Asp	Tyr	Gly	Gly	Asp	Ser	Leu	Gln	Ala	Met	Gln
	770					775					780				
Leu	Val	Ser	Arg	Ile	Trp	Ser	Ser	Phe	Ser	Val	Glu	Ile	Gly	Ile	Asp
	785					790					795				800
Ala	Ile	Phe	Glu	Leu	Gln	Thr	Ile	Ser	Ala	Val	Ser	Asp	Leu	Ile	Glu
				805					810					815	
Ala	Ser	Ser	Pro	His	Pro	Gly	Ser	Thr	Ala	Gly	Ala	Ile	Pro	Pro	Arg
			820					825					830		
Ser	Arg	Ala	Asn	Asp	Leu	Pro	Leu	Ser	Phe	Pro	Gln	Gln	Arg	Leu	Trp
		835					840						845		
Phe	Leu	Ala	Gln	Leu	Glu	Gly	Pro	Ser	Ala	Thr	Tyr	Asn	Ile	Ser	Ser
	850					855					860				
Ala	Leu	Arg	Phe	Glu	Gly	Glu	Leu	Asp	Val	Ala	Arg	Leu	Arg	Phe	Ala
	865					870					875				880
Val	Ser	Glu	Ile	Ser	Arg	Arg	His	Glu	Ile	Leu	Arg	Thr	Thr	Phe	Pro
				885					890					895	
Ala	Val	Asp	Gly	Arg	Gly	Val	Gln	Arg	Ile	Ala	Pro	Pro	Ala	Pro	Val
			900					905					910		
Ala	Leu	Asp	Val	Val	Asp	Val	Ala	Ser	Glu	Ser	Asp	Thr	Leu	Ala	Leu
		915					920						925		
Leu	Ala	Glu	Glu	Ala	Asp	Arg	Pro	Phe	Asp	Leu	Ala	Ala	Gly	Pro	Leu
	930					935							940		
Tyr	Arg	Val	Val	Leu	Tyr	Arg	Val	His	Glu	Arg	Leu	His	Val	Phe	Gly
	945					950					955				960
Ile	Val	Met	His	His	Ile	Val	Ser	Asp	Ala	Trp	Ser	Ser	Gly	Ile	Leu
				965					970					975	
Ile	Gly	Glu	Leu	Ala	Ala	Leu	Tyr	Ala	Gly	Glu	Ser	Leu	Pro	Glu	Leu
			980					985						990	
Ala	Val	Gln	Tyr	Ala	Asp	Tyr	Ala	Val	Trp	Gln	His	Glu	Arg	Leu	Ala
		995					1000						1005		
Ser	Ala	Asp	Thr	His	Arg	Glu	Leu	Ala	Leu	Leu	Ser	Ala	Ala	Leu	
	1010						1015						1020		
Ala	Asp	Ala	Pro	Asp	Leu	Ile	Glu	Leu	Pro	Thr	Asp	Arg	Pro	Arg	
	1025						1030						1035		
Pro	Ala	Val	Gln	Gln	Phe	Arg	Gly	Ala	Val	Leu	Pro	Phe	Gln	Leu	
	1040						1045						1050		
Ser	Ala	Glu	Arg	Ala	Asp	Gly	Leu	Arg	Ala	Ile	Ala	Arg	Ala	Ser	
	1055						1060						1065		
Gly	Thr	Ser	Thr	Phe	Met	Val	Val	Leu	Ala	Ala	Tyr	Ala	Leu	Leu	
	1070						1075						1080		

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Leu Ser Arg Tyr Ser Asn Gln Gln Asp Leu Val Ile Gly Ser Pro 1085 1090 1095
Ile Ala Asn Arg Arg Ser Ser Met Thr Glu Pro Leu Ile Gly Phe 1100 1105 1110
Phe Ala Asn Met Leu Ala Leu Arg Val Asp Leu Ser Gly Asn Pro 1115 1120 1125
Thr Phe Gly Asp Leu Leu Ala Arg Val Lys Arg Val Ala Leu Asp 1130 1135 1140
Gly Tyr Ser Arg Gln Glu Ile Pro Phe Glu Gln Val Val Asp Ser 1145 1150 1155
Leu Glu Leu Glu Arg Asn Leu Gly Arg Thr Pro Val Phe Gln Val 1160 1165 1170
Val Phe Ala Tyr Glu Lys Ala Gln Pro Arg Ala Val Ser Phe Pro 1175 1180 1185
Gly Leu Val Ala Thr Pro Val Ala Val Glu Thr His Thr Ala Lys 1190 1195 1200
Phe Asp Leu Thr Leu His Val Gln Asp Ala Asp Asp Gly Leu Ala 1205 1210 1215
Gly Ser Leu Glu Tyr Asn Leu Asp Leu Phe Asp Ala Ala Thr Ile 1220 1225 1230
Asp Arg Met Ala Glu His Phe Arg Thr Leu Val Asp Ala Val Ile 1235 1240 1245
Ala Asp Pro Asp Arg Pro Leu Gly Ala Leu Ser Leu Ser Asn Asp 1250 1255 1260
Ala Glu Arg Asn Leu Leu Thr Val Glu Trp Asn Arg Thr Asp Thr 1265 1270 1275
Asp Phe Gly Glu Asp Ala Ala Gln Pro Leu His Arg Leu Phe Glu 1280 1285 1290
Gln Gln Val Glu Arg Thr Pro Asp Ala Val Ala Ile Val Phe Asp 1295 1300 1305
Asp Thr Ala Leu Thr Tyr Ala Glu Leu Asn Leu Arg Ala Asn Arg 1310 1315 1320
Leu Ala His His Leu Val Ala Leu Gly Val Gly Pro Asp Ser Leu 1325 1330 1335
Val Gly Val Ala Met Glu Arg Ser Leu Asp Met Ser Val Ala Leu 1340 1345 1350
Leu Ala Ile Leu Lys Ala Gly Gly Ala Tyr Val Pro Val Asp Pro 1355 1360 1365
Asp Tyr Pro Ala Glu Arg Val Arg Phe Met Ile Asp His Ala Gln 1370 1375 1380
Leu Arg Trp Leu Leu Thr Gln Gln His Leu His Asp Ala Leu Pro 1385 1390 1395
Asp Thr Asp Ala His Val Ile Val Val Asp Arg Asp Ser Leu Asp 1400 1405 1410
Leu Asp Ala Ala Ala Thr Ser Asn Pro Ala Pro Ala Leu Asn Gly 1415 1420 1425
Asp Asn Leu Ala Tyr Met Ile Tyr Thr Ser Gly Ser Thr Gly Arg 1430 1435 1440
Pro Lys Gly Ala Leu Asn Thr His Arg Ala Ile Thr Asn Arg Ile 1445 1450 1455
Leu Trp Met Gln His Ala Tyr Ala Leu Asp Ala Asp Asp Ala Val 1460 1465 1470

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Leu	Gln	Lys	Thr	Pro	Phe	Ser	Phe	Asp	Val	Ser	Val	Trp	Glu	Leu
1475						1480						1485		
Phe	Trp	Pro	Leu	Val	Thr	Gly	Ala	Arg	Leu	Val	Phe	Ala	Arg	Pro
1490						1495					1500			
Gly	Gly	Gln	Arg	Glu	Thr	Asp	Tyr	Leu	Val	Glu	Leu	Ile	Glu	Arg
1505						1510					1515			
Glu	Arg	Ile	Thr	Thr	Ile	His	Phe	Val	Pro	Ser	Met	Leu	Arg	Ala
1520						1525					1530			
Phe	Leu	Asp	His	Pro	Asp	Leu	Asp	Ala	His	Cys	Ala	Ser	Leu	Arg
1535						1540					1545			
Arg	Val	Val	Cys	Ser	Gly	Glu	Ala	Leu	Pro	His	Asp	Leu	Gln	Gln
1550						1555					1560			
Arg	Cys	Leu	Glu	Arg	Leu	Asp	Val	Lys	Leu	Tyr	Asn	Leu	Tyr	Gly
1565						1570					1575			
Pro	Thr	Glu	Ala	Ala	Val	Asp	Val	Thr	Ala	Trp	Glu	Cys	Arg	Arg
1580						1585					1590			
Asp	Asp	Pro	His	Arg	Ile	Val	Pro	Ile	Gly	Arg	Pro	Ile	Ala	Asn
1595						1600					1605			
Thr	Arg	Leu	Tyr	Ile	Val	Asp	Ala	Gln	Met	Gln	Pro	Thr	Pro	Ile
1610						1615					1620			
Gly	Val	Ala	Gly	Glu	Leu	Leu	Ile	Gly	Gly	Thr	Pro	Val	Gly	Arg
1625						1630					1635			
Gly	Tyr	His	Gly	Glu	Pro	Glu	Leu	Ser	Ala	Glu	Lys	Phe	Ile	Ala
1640						1645					1650			
Asp	Pro	Phe	Ser	Ala	Asp	Pro	Leu	Ala	Arg	Leu	Tyr	Arg	Thr	Gly
1655						1660					1665			
Asp	Leu	Ala	Arg	Tyr	Arg	Pro	Asp	Gly	Asn	Ile	Glu	Phe	Leu	Gly
1670						1675					1680			
Arg	Ile	Asp	His	Gln	Ile	Lys	Leu	Arg	Gly	Leu	Arg	Ile	Glu	Pro
1685						1690					1695			
Gly	Glu	Ile	Glu	Ala	Ala	Leu	Arg	Ala	His	Pro	Ser	Val	Asp	Asp
1700						1705					1710			
Cys	Val	Val	Ile	Ala	Lys	Thr	Glu	Gly	Ala	Arg	Thr	Phe	Leu	Ile
1715						1720					1725			
Ala	Tyr	Val	Ala	Thr	Ala	Ala	Pro	Asp	Ile	Ala	Asp	Leu	Arg	Gly
1730						1735					1740			
Tyr	Leu	Gly	Gly	Lys	Leu	Ala	Asp	Tyr	Met	Val	Pro	Ser	Gln	Phe
1745						1750					1755			
Phe	Ala	Leu	Glu	Ser	Leu	Pro	Met	Leu	Pro	Asn	Gly	Lys	Ile	Asn
1760						1765					1770			
Arg	Lys	Ala	Leu	Pro	Leu	Pro	Ala	Asp	Arg	Gly	Asp	Ala	Ala	Gln
1775						1780					1785			
Pro	His	Ala	Pro	Ala	Val	Thr	Pro	Arg	Glu	Ile	Leu	Leu	Ala	Ser
1790						1795					1800			
Ile	Cys	Ile	Asp	Val	Leu	Gln	Leu	Pro	Ser	Val	Gly	Ile	His	Asp
1805						1810					1815			
Asn	Phe	Phe	Glu	Leu	Gly	Gly	Asp	Ser	Ile	Leu	Ser	Ile	Gln	Val
1820						1825					1830			
Ile	Ala	Arg	Ala	Asn	Gln	Ala	Gly	Leu	Arg	Val	Thr	Ala	Lys	Gln
1835						1840					1845			
Leu	Phe	Gln	Tyr	Gln	Thr	Ile	Ala	Gln	Leu	Ala	Ala	Ala	Pro	Glu
1850						1855					1860			
Glu	Arg	Ala	Ala	Cys	Ala	Pro	Thr	Leu	Ser	Pro	Leu	Gly	Asp	Ala

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1865	1870	1875
Pro Leu Thr Pro Val Gln His Trp Phe Phe Glu Gln Glu Ile Asp 1880	1885	1890
Ala Pro Ser His Tyr Asn Gln Thr Val Leu Ile Gln Val Pro Ala 1895	1900	1905
Asp Ile Asp Ala Ser Arg Leu Ala Asp Ala Phe Arg Gln Val Tyr 1910	1915	1920
Glu His His Asp Ala Leu Arg Leu Arg Phe Ser His Asp Ala Gly 1925	1930	1935
Arg Trp Thr Gln Gln Val Val Ala Gly Gly Glu Met Pro Ala Leu 1940	1945	1950
Phe Ala Lys Gln Val Ile Ala Asp Asp Ala Gly Glu Arg Leu Ala 1955	1960	1965
Ala Met Arg Ala Ala Ala Ala Asp Ala Glu Arg Gly Ile Asp Ile 1970	1975	1980
Thr His Gly Pro Leu Leu Ala Ala Arg Leu Phe Cys Leu Ala Asp 1985	1990	1995
Glu Pro Leu Ala Arg Leu Phe Val Ser Ile His His Leu Ala Val 2000	2005	2010
Asp Gly Val Ser Trp Arg Val Leu Leu Glu Asp Leu His Ala Ala 2015	2020	2025
Tyr His Gly Gln Pro Leu Pro Gly Lys Thr Thr Ser Phe Arg Glu 2030	2035	2040
Trp Ala Leu His Leu Gln Gln Leu Ala Arg Ser Pro Ala Ile Gly 2045	2050	2055
Asp Glu Ala Arg Leu Trp Gln Ala Leu Leu Ala Gln Pro Val Glu 2060	2065	2070
Pro Met Pro Val Asp Tyr Pro Gly Thr Gly Ala Ala Asn Asn Ala 2075	2080	2085
Val Asp Asp Ala Ser Ser Val Ser Phe Glu Leu Gly Glu Ala Asp 2090	2095	2100
Thr Thr Ala Leu Leu Arg Arg Leu Pro Arg Ala Tyr Asp Thr Arg 2105	2110	2115
Ile Asn Asp Val Leu Leu Val Ala Leu Ala Gln Ala Cys Ser Met 2120	2125	2130
Val Thr Gly Asn Thr Arg Thr Arg Ile Asp Leu Glu Ser His Gly 2135	2140	2145
Arg His Val Ser Asp Ala Pro Leu Asp Leu Thr Arg Thr Val Gly 2150	2155	2160
Trp Phe Thr Ser Ile Tyr Pro Val Val Leu Asp Ala Asp Ala Met 2165	2170	2175
His Ala Pro Glu Gln Ala Leu Arg Ala Ala Arg Gln Gln Leu Arg 2180	2185	2190
Arg Ile Pro Ala Asp Gly Leu Gly Tyr Ser Leu Leu Arg Tyr Gln 2195	2200	2205
Ser Pro Asp Ala Ala Val Arg Asp Ser Leu Ala Ala Leu Pro Lys 2210	2215	2220
Ala Asp Ile Leu Phe Asn Tyr His Gly Gln Leu Asp Thr Val Leu 2225	2230	2235
Arg Gln Ser Asp Gly Trp Arg Pro Ala Ala Glu Asp Leu Gly Ser 2240	2245	2250
Leu Arg Ala Gly Arg Ser Gln Arg Thr His Ala Phe Glu Ile Val 2255	2260	2265

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Ala	Ala	Val	Ala	Asp	Gly	Lys	Leu	Gln	Val	Asp	Trp	Arg	Tyr	Gly
2270						2275					2280			
Glu	Arg	Leu	His	Arg	Arg	Gln	Thr	Val	Glu	Asn	Leu	Ala	Ala	His
2285						2290					2295			
Phe	Arg	Asp	Arg	Leu	Leu	Asp	Phe	Ala	Ala	Ser	Val	Pro	Asp	Thr
2300						2305					2310			
Ala	Ala	Asp	Asp	Ile	Glu	Asp	Ser	Tyr	Pro	Leu	Ser	Ser	Leu	Gln
2315						2320					2325			
Gln	Gly	Ile	Leu	Phe	His	Ser	Leu	Tyr	Asp	Leu	Asp	Pro	Ala	Ala
2330						2335					2340			
Tyr	Phe	Gln	Gln	Phe	Ser	Phe	Val	Val	Ser	Gly	Pro	Leu	Gln	Val
2345						2350					2355			
Pro	Ala	Leu	Arg	Gln	Ala	Trp	Ala	Asn	Ala	Leu	Ala	Arg	His	Ala
2360						2365					2370			
Val	Leu	Arg	Thr	Ala	Phe	Ala	Trp	Ala	Asp	Arg	Asp	His	Pro	Val
2375						2380					2385			
Gln	Thr	Val	Arg	His	Thr	Val	Asp	Leu	Pro	Trp	Thr	Phe	Leu	Asp
2390						2395					2400			
Trp	Arg	His	Arg	Asp	Ala	Ser	Arg	Arg	Ala	Gln	Asp	Phe	Asp	Ala
2405						2410					2415			
Phe	Leu	Ala	Asp	Asp	Arg	Arg	Arg	Gly	Phe	Asp	Leu	Gln	Arg	Ala
2420						2425					2430			
Pro	Leu	Phe	Arg	Cys	Thr	Leu	Ile	Gln	Glu	Thr	Asp	Thr	Arg	His
2435						2440					2445			
Arg	Phe	Cys	Trp	Ser	Ala	His	His	Ile	Ile	Leu	Asp	Gly	Trp	Ser
2450						2455					2460			
Thr	Ala	Thr	Leu	Met	Lys	Glu	Val	Phe	Asp	Asp	Tyr	Leu	Ser	Leu
2465						2470					2475			
Ala	Arg	Thr	Gly	Met	Pro	Ala	Val	Ala	Ala	Ser	Ala	Pro	Gly	Tyr
2480						2485					2490			
Arg	Ala	Tyr	Ile	Asp	Trp	Leu	Ala	Arg	His	Pro	Arg	Ser	Ala	Asp
2495						2500					2505			
Glu	Thr	Trp	Trp	Arg	Ala	Glu	Leu	Ala	Gly	Phe	Lys	Ala	Ala	Thr
2510						2515					2520			
Pro	Val	Ala	Ala	Ser	Pro	Ala	Arg	Gln	Ala	Thr	Gly	Asp	Ala	Pro
2525						2530					2535			
Arg	Gln	Asp	Lys	Arg	Arg	Thr	Gln	Gln	Phe	Leu	Leu	Asp	Glu	Ala
2540						2545					2550			
Leu	Ala	Ala	Arg	Leu	Gln	Thr	Leu	Thr	Arg	Thr	His	Arg	Val	Thr
2555						2560					2565			
Leu	Asn	Val	Leu	Ile	Arg	Ala	Val	Trp	Ala	Leu	Val	Leu	Arg	Arg
2570						2575					2580			
His	Ala	Gly	Thr	Asp	Asp	Val	Val	Phe	Gly	Val	Thr	Val	Ser	Gly
2585						2590					2595			
Arg	Pro	Pro	Met	Leu	Asp	Gly	Val	Glu	Ser	Ile	Val	Gly	Leu	Phe
2600						2605					2610			
Ile	Asn	Thr	Leu	Pro	Leu	Arg	Leu	Arg	Ile	Ala	Pro	Glu	Arg	Pro
2615						2620					2625			
Phe	Ile	Glu	Trp	Leu	Ala	Glu	Val	His	Ala	Ala	Gln	Thr	Ala	Met
2630						2635					2640			
Glu	Pro	His	Ser	Tyr	Ser	Ser	Leu	Val	Asp	Ile	Gln	Ser	Trp	Ser
2645						2650					2655			

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Glu	Leu	Pro	Ala	Gly	Asp	Ser	Leu	Phe	Asp	Ser	Leu	Leu	Val	Phe
2660						2665					2670			
Glu	Asn	Phe	Pro	Val	Ala	Ala	Ala	Pro	Asp	Leu	Gly	Pro	Asp	Asp
2675						2680					2685			
Ile	Glu	Ile	Leu	Asp	Thr	Arg	Ala	Phe	Ala	Glu	Ser	Asn	Tyr	Pro
2690						2695					2700			
Leu	Thr	Leu	Thr	Val	His	Pro	Asn	Glu	Arg	Ile	Gly	Phe	His	Ile
2705						2710					2715			
Ser	His	Asp	Ala	His	Arg	Ile	Ala	Pro	Glu	Val	Val	Arg	Gln	Met
2720						2725					2730			
Leu	Asp	Thr	Leu	Arg	Thr	Leu	Leu	Glu	Arg	Phe	Ala	Glu	Asn	Pro
2735						2740					2745			
Gly	Gln	Leu	Thr	Gly	Gln	Leu	Ala	Asp	Pro	Pro	Ala	Ala	Asp	Gly
2750						2755					2760			
Arg	Pro	Ser	Ala	Pro	Arg	Ser	Gly	Ala	Gly	Pro	Ala	Ile	Glu	Ala
2765						2770					2775			
Ala	Ala	Gly	Ala	Ala	Ala	Ala	Ala	Arg	Ala	Val	Ala	His	Ala	Ala
2780						2785					2790			
Asp	Glu	Ser	Thr	Leu	Leu	Glu	Ile	Trp	Arg	Arg	Ile	Phe	Lys	Arg
2795						2800					2805			
Asp	Asp	Ile	Ala	Val	Ser	Asp	Asn	Tyr	Phe	Asp	Leu	Gly	Gly	His
2810						2815					2820			
Ser	Ile	Ile	Ala	Ile	Gln	Leu	Met	Ala	His	Val	Glu	Lys	Ala	Phe
2825						2830					2835			
Asp	Arg	Arg	Leu	Pro	Ile	Ser	Cys	Leu	Phe	Glu	Asn	Pro	Thr	Ile
2840						2845					2850			
Glu	Lys	Leu	Ala	Ala	Ala	Leu	Ala	Ala	Lys	Glu	Pro	Ser	Ala	Pro
2855						2860					2865			
Ala	Gly	Gly	Leu	Val	Pro	Ile	Arg	Asp	Gly	Gly	Pro	Ala	Ala	Pro
2870						2875					2880			
Leu	Phe	Leu	Leu	Pro	Gly	Ala	Gly	Gly	Asn	Val	Val	Tyr	Phe	Arg
2885						2890					2895			
Pro	Leu	Ala	Asn	His	Leu	Ser	Gly	Ala	His	Ala	Ile	His	Gly	Leu
2900						2905					2910			
Glu	Ala	Leu	Gly	Leu	Asp	Gly	Ala	Cys	Glu	Pro	Leu	Thr	Arg	Val
2915						2920					2925			
Glu	Asp	Ile	Ala	Ala	Arg	His	Ile	Glu	Arg	Ile	Trp	Pro	Leu	Val
2930						2935					2940			
Gly	Ala	Gly	Pro	Tyr	Tyr	Leu	Ala	Gly	His	Ser	Phe	Gly	Ala	His
2945						2950					2955			
Val	Ala	Leu	Glu	Met	Ser	Arg	Gln	Leu	Val	Ala	Lys	Gly	Ala	Asp
2960						2965					2970			
Val	Lys	Leu	Leu	Ala	Ile	Phe	Asp	Ala	Ser	Ala	Pro	Ile	Asp	Ser
2975						2980					2985			
Ser	Ala	Ala	Thr	Tyr	Trp	Gln	Asp	Trp	Asp	Asp	Thr	Glu	Trp	Leu
2990						2995					3000			
Val	Ala	Ile	Ala	His	Glu	Ile	Gly	Thr	Phe	Leu	Gly	Thr	Asp	Leu
3005						3010					3015			
Gln	Val	Thr	Arg	Ala	Asp	Leu	Val	His	Leu	Asp	Pro	Asp	Gly	Gln
3020						3025					3030			
Ala	Gly	Leu	Ile	Leu	Glu	Arg	Ile	Gly	Asp	Arg	Gly	Ser	Trp	Phe
3035						3040					3045			
Ala	Asp	Ala	Gly	Ser	Asp	Arg	Leu	Arg	Ala	Tyr	Leu	Arg	Val	Tyr



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3050	3055	3060
Gln Ala Asn Phe Lys Ser His	Tyr Ala Pro His Ala	Thr Pro Leu
3065	3070	3075
Pro Val Pro Ile Ala Leu Phe	Arg Ser Thr Glu Arg	Asp Pro Gly
3080	3085	3090
Asp Tyr Ala Pro Ser Ser Glu	Ile Ala Gln Leu Arg	Leu Asp Ala
3095	3100	3105
Thr Trp Gly Trp Ser Arg Phe	Ser Ala His Pro Val	Ala Val Thr
3110	3115	3120
Asp Val Pro Gly Asp His Leu	Thr Met Leu Leu Asp	Pro His Ala
3125	3130	3135
Gly Val Leu Ala Ala His Val	Asn Ser Phe Leu Glu	Lys Thr Pro
3140	3145	3150

Ser

<210> SEQ ID NO 5  
 <211> LENGTH: 391  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(391)  
 <223> OTHER INFORMATION: putative FAD linked oxidase domain protein

&lt;400&gt; SEQUENCE: 5

Met Ser His Asp Phe Arg Asp Glu Pro Ala Pro Arg Arg Ala Phe Leu	1	5	10	15
Ala Asp Met Ala Lys Leu Ala Ala Ala Gly Ile Val Thr Gly Trp Thr	20	25	30	
Pro Leu Tyr Gln Val Ala Ala His Ala Arg Thr Ala Gly Glu Thr Pro	35	40	45	
Pro Gly Phe Pro Ala Asp Ile Gln Leu Tyr Lys Gln Ala Phe Leu Asn	50	55	60	
Trp Ser Gly Glu Ile Ala Val Gln Asp Val Trp Thr Ala Ala Pro Arg	65	70	75	80
Ser Ala Asp Asp Val Val Ala Thr Val Asn Trp Ala Arg Ala Asn Gly	85	90	95	
Tyr Arg Ile Arg Pro Arg Gly Tyr Thr His Asn Trp Ser Pro Leu Thr	100	105	110	
Leu Asp Pro Gly Ala Gly Ala Ala Asn Leu Val Leu Leu Asp Thr Thr	115	120	125	
Lys Ser Leu Thr Ala Val Ser Val Asp Thr Ser Ala Arg Pro Ala Arg	130	135	140	
Val Thr Ala Gln Thr Gly Val Ser Leu Glu Ser Leu Leu Ala Thr Leu	145	150	155	160
Glu Gln Val Gly Leu Gly Val Ile Ala Ala Pro Ala Pro Gly Asp Ile	165	170	175	
Thr Leu Gly Gly Ala Leu Ala Ile Asp Ala His Gly Thr Ala Val Pro	180	185	190	
Ala Ala Gly Glu Thr Leu Gln Pro Gly His Thr Tyr Gly Ser Leu Ser	195	200	205	
Asn Leu Val Val Ala Leu Thr Ala Val Val Phe Asp Pro Ala Arg Gln	210	215	220	
Gln Tyr Val Leu Arg Arg Phe Glu Arg Ser Asp Pro Glu Ile Gly Ala	225	230	235	240

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Phe Leu Ala His Ile Gly Arg Ala Leu Val Val Glu Val Thr Leu Thr  
 245 250 255  
 Ala Gly Pro Asn Gln Arg Leu Arg Cys Gln Ser Tyr Val Asp Ile Pro  
 260 265 270  
 Ala Ser Glu Leu Phe Ala Ala Pro Gly Thr Thr Gly Arg Thr Ile Ala  
 275 280 285  
 Ser Phe Leu Asp Gly Ser Gly Arg Val Glu Ala Ile Trp Phe Pro Phe  
 290 295 300  
 Thr Thr Lys Pro Trp Leu Lys Val Trp Thr Pro Thr Pro Ser Lys Pro  
 305 310 315 320  
 Phe Leu Ser Arg Ala Val Thr Gln Pro Tyr Asn Tyr Pro Phe Ser Asp  
 325 330 335  
 Ser Ile Ser Gln Ser Ile Ser Asp Leu Val Lys Arg Ile Val Ile Gly  
 340 345 350  
 Gly Glu Gly Ala Leu Thr Pro Leu Phe Gly Gln Thr Gln Leu Ala Ile  
 355 360 365  
 Thr Thr Ala Gly Leu Ala Leu Thr Leu Ser Gly Asp Ile Trp Gly Trp  
 370 375 380  
 Ser Arg Thr Val Leu Gln Glu  
 385 390

<210> SEQ ID NO 6  
 <211> LENGTH: 273  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(273)  
 <223> OTHER INFORMATION: putative LuxR-type regulator

<400> SEQUENCE: 6

Met Phe Ala Lys Leu Gly Lys Val Ile Ser Ser Ala Gly Ser Glu Arg  
 1 5 10 15  
 Phe Ala Ser Asp Met His Ala Leu Leu Val Glu Ser Ile Pro Leu Thr  
 20 25 30  
 Ile Thr Arg Met Thr Glu Trp Thr Leu Asp Glu Pro Ala Gly Glu Val  
 35 40 45  
 Val Arg Val Gln Ser Leu Gly Ala Asp Gly Ala Pro Gly Asp Asp Gly  
 50 55 60  
 Arg Gly Ala Pro Ala Ala His Gly Glu Arg Glu Pro Ala Ala His Pro  
 65 70 75 80  
 Pro Leu Asn Arg Ile Leu Ala Ala Cys Asp Arg Gln Leu Ile His Ile  
 85 90 95  
 Asn Pro Leu Met Arg Arg Gly Asn Gly Gly Glu Val Ala Pro Ser Arg  
 100 105 110  
 Gly Pro Gly Gly Gly Phe Gln Cys His Leu Val Ser Gly Lys Ala Asn  
 115 120 125  
 Arg Arg Tyr Val Ile Ser Leu His Arg Thr Ala Ser His Arg Asp Phe  
 130 135 140  
 Ser Leu Arg Glu Met Ser Phe Leu Lys Asn Phe Ala Asp Thr Leu Leu  
 145 150 155 160  
 Pro Leu Val Glu Trp His Ala Ser Thr Cys Arg His Gly Glu Arg Glu  
 165 170 175  
 Gly Ala Thr Ala Pro Gly Ala Thr Ala Gly Met Pro Gly Val Glu Ala  
 180 185 190  
 Leu Arg His Glu Phe Glu Ser Arg Leu Ala Arg Ala Arg Val Val Leu

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195					200					205					
Ser	Ala	Arg	Glu	Asn	Glu	Val	Cys	Leu	Gly	Leu	Leu	Ala	Gly	Lys	Met
210					215					220					

Leu	Arg	Glu	Met	Ala	Gly	Glu	Leu	Gly	Val	Lys	Glu	Ser	Thr	Ile	Glu
225					230					235					240

Thr	Tyr	Ile	Lys	Arg	Ala	Ala	Val	Lys	Leu	Gly	Ile	Ser	Gly	Arg	His
				245					250					255	

Gly	Leu	Thr	Lys	Trp	Met	Ile	Asp	Asp	Ser	Val	Pro	Cys	Ala	Ser	Ala
			260					265					270		

Ala

<210> SEQ ID NO 7  
 <211> LENGTH: 296  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(296)  
 <223> OTHER INFORMATION: putative LuxR-type regulator

<400> SEQUENCE: 7

Met	Glu	Phe	Ser	Arg	Leu	Phe	Ala	His	Val	Gly	Glu	Ala	Ile	Ser	Ser
1				5					10					15	

Ser	Gly	Ser	Arg	Arg	Phe	Pro	Arg	Met	Met	Tyr	Asn	Leu	Ile	Ala	Ala
			20					25					30		

Ala	Val	Pro	Val	Asp	Glu	Ile	Arg	Ile	Ser	Glu	Leu	Ala	Ile	Asp	Asp
			35				40					45			

Val	Pro	Asp	Gly	Pro	Pro	Glu	Val	Arg	Ser	Leu	Gly	Ala	Val	Gly	Ala
			50			55					60				

Ala	Leu	Ala	Lys	Thr	Gly	Ala	Ala	Ala	Val	Cys	Cys	Gly	Pro	Gln	Met
65				70						75				80	

Pro	Pro	Arg	Pro	Gly	Thr	Ser	Pro	Leu	His	Val	Asp	Asp	Thr	Leu	Ala
				85					90					95	

Gly	His	Gly	Pro	Ile	His	Ala	Gln	Leu	Asp	Arg	Phe	Ile	Leu	Met	Gln
			100					105					110		

Ala	Ala	Ile	Val	Ser	Pro	Arg	Tyr	Ala	Gln	Phe	His	Leu	Val	Thr	Arg
			115				120					125			

Lys	Arg	Gly	His	Cys	Tyr	Val	Ile	Ser	Leu	Tyr	Arg	Thr	Cys	Thr	Phe
			130			135						140			

Asp	Asp	Phe	Ser	Pro	Gln	Glu	Arg	Thr	Phe	Leu	Lys	Glu	Leu	Ser	His
145				150					155					160	

Val	Leu	Phe	Pro	Ile	Val	Glu	Ser	His	Val	Ala	Ala	Leu	Asp	Ser	Ala
				165					170					175	

Pro	Pro	Ala	Ala	Arg	Val	Thr	Thr	Ala	Ala	Pro	Pro	Ala	Thr	Gln	Ser
				180				185					190		

Gly	Arg	Glu	Arg	Val	Ala	Arg	Arg	Phe	Ala	Asp	Arg	Leu	Gln	Gln	Ala
				195				200					205		

Gly	Val	Lys	Leu	Ser	Thr	Arg	Glu	Ile	Glu	Ala	Cys	Thr	Ala	Leu	Leu
	210					215					220				

Ala	Gly	Asp	Thr	Val	Pro	Ala	Ile	Ala	Met	Arg	Phe	Ala	Leu	Arg	Glu
225				230							235			240	

Ser	Thr	Val	Glu	Thr	Tyr	Leu	Lys	Arg	Ala	Ala	Val	Lys	Leu	Gly	Phe
				245					250					255	

Ser	Gly	Arg	His	Gly	Leu	Thr	Arg	Trp	Met	Leu	Asp	Glu	Thr	Ala	Gly
			260					265					270		

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Ala Ala Thr Glu Ala Ala Gly Gly Asp Met Arg Ser Met Arg Arg Asp  
 275 280 285

Tyr Ala Ser Pro Arg Leu Gly Thr  
 290 295

<210> SEQ ID NO 8  
 <211> LENGTH: 567  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(567)  
 <223> OTHER INFORMATION: putative cyclic peptide transporter

<400> SEQUENCE: 8

Met Asp Ser Ala Gln Ser Lys Ser Pro Pro Trp His Ser Ala Ala Thr  
 1 5 10 15

Leu Met Trp Arg Ser His Pro Trp Leu Thr Leu Gly Thr Val Val Thr  
 20 25 30

Gly Leu Val Ser Gly Ile Ala Ser Ile Ala Gly Val Gly Leu Ile Ser  
 35 40 45

Thr Val Leu His Asp Gln Asp Asp Arg Gln Thr Leu Leu Leu Phe  
 50 55 60

Ile Ala Val Asn Val Val Ala Val Val Cys Arg Ser Cys Ala Ala Val  
 65 70 75 80

Met Pro Ser Tyr Ala Cys Met Lys Val Met Thr Arg Leu Arg Val Asn  
 85 90 95

Leu Cys Lys Arg Ile Leu Ala Thr Pro Leu Asp Glu Ile Asp Arg Arg  
 100 105 110

Gly Ala Pro Asn Val Leu Thr Met Leu Thr Gln Asp Ile Pro Gln Leu  
 115 120 125

Ser Gln Thr Leu Leu Thr Ile Pro Thr Ile Ile Val Gln Ser Val Val  
 130 135 140

Leu Ile Cys Ser Ile Ala Tyr Leu Ala Tyr Leu Ser Trp Ile Val Phe  
 145 150 155 160

Ala Ser Thr Ile Ile Leu Thr Leu Val Gly Leu Val Leu Tyr Leu Phe  
 165 170 175

Phe Tyr Arg Lys Ala Val Asn Phe Thr Glu Arg Val Arg Asp Glu Phe  
 180 185 190

Val Gln Phe Asn Glu Tyr Thr His Gly Leu Val Phe Gly Ile Lys Glu  
 195 200 205

Leu Lys Leu Asn Arg Ala Arg Arg Arg Trp Phe Thr Arg Ala Ala Ile  
 210 215 220

Glu Leu Ser Ser Lys Arg Val Ala Gly Phe Asn Tyr Ile Glu Arg Phe  
 225 230 235 240

Trp Phe Met Ser Gly Asp Ser Ile Gly Gln Ile Thr Val Ala Val Leu  
 245 250 255

Leu Gly Cys Leu Leu Phe Gly Val Pro Ser Leu Gly Val Val Asp Pro  
 260 265 270

Ser Val Leu Thr Ala Ser Ile Leu Ala Val Leu Tyr Met Met Gly Pro  
 275 280 285

Leu Thr Met Leu Ile Asn Val Leu Pro Val Val Ala Glu Gly Lys Thr  
 290 295 300

Ala Leu Ala Arg Leu Ala Glu Phe Gly Phe Leu Ile Asp Asp Thr Gln  
 305 310 315 320

Ala Ser His Glu Glu Pro Arg Pro Ala Gly Asn Val Glu Thr Leu Ser

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325					330					335					
Ala	Lys	Ser	Trp	Lys	Val	Ile	Glu	Leu	Lys	Asp	Val	Thr	Met	Asn	Tyr
			340					345						350	
Arg	Asp	Asn	Glu	Ala	Ser	Val	Asp	Phe	Val	Leu	Gly	Pro	Ile	Asp	Met
		355					360					365			
Thr	Ile	His	Ala	Gly	Glu	Leu	Val	Tyr	Val	Ile	Gly	Gly	Asn	Gly	Ser
	370					375					380				
Gly	Lys	Ser	Thr	Leu	Gly	Lys	Val	Leu	Ser	Gly	Leu	Tyr	Ala	Pro	Thr
385					390					395					400
Gly	Gly	Thr	Ile	Ser	Leu	Asp	Gly	Lys	Val	Val	Asp	Asp	Ala	Ala	Arg
			405						410						415
Glu	Arg	Tyr	Arg	Asn	Leu	Phe	Ser	Ala	Val	Phe	Thr	Asp	Phe	His	Leu
			420					425						430	
Phe	Asn	Arg	Ile	Ile	Gly	Pro	Asp	Arg	Gly	Asn	Glu	Ser	Ile	Glu	Leu
		435					440						445		
Ala	Arg	Lys	Tyr	Leu	Ala	Thr	Leu	Lys	Leu	Ala	Asp	Lys	Ile	Glu	Ile
	450					455					460				
Ser	Gly	Arg	Thr	Tyr	Ser	Thr	Thr	Arg	Ala	Leu	Ser	Thr	Gly	Gln	Arg
465					470					475					480
Lys	Arg	Leu	Ala	Leu	Leu	Cys	Ala	Tyr	Ile	Glu	Asp	Arg	Pro	Ile	Tyr
			485						490						495
Ile	Leu	Asp	Glu	Trp	Ala	Ala	Asp	Gln	Asp	Pro	Val	Phe	Lys	Arg	Phe
			500					505						510	
Ser	Tyr	Glu	Val	Leu	Val	Pro	Asp	Leu	Lys	Ser	Arg	Gly	Lys	Cys	Val
		515					520						525		
Val	Ile	Ile	Thr	His	Asp	Asp	Gln	Tyr	Phe	Lys	Leu	Ala	Asp	Arg	Val
	530					535						540			
Ile	Arg	Leu	Asp	Ser	Gly	Arg	Ile	Phe	Ser	Asp	Thr	Ala	Met	Cys	Ala
545					550					555					560
Val	Arg	Ala	Glu	Ala	Ala	Gly									
					565										

&lt;210&gt; SEQ ID NO 9

&lt;211&gt; LENGTH: 160

&lt;212&gt; TYPE: PRT

&lt;213&gt; ORGANISM: Burkholderia contaminans

&lt;220&gt; FEATURE:

&lt;221&gt; NAME/KEY: MISC\_FEATURE

&lt;222&gt; LOCATION: (1)..(160)

&lt;223&gt; OTHER INFORMATION: hypothetical protein

&lt;400&gt; SEQUENCE: 9

Met	Gln	Leu	Thr	Thr	Val	Asp	Leu	Glu	Ala	Ala	Phe	Val	Lys	Ala	Ala
1				5					10					15	
Leu	Asp	Ala	Leu	His	Arg	Asp	Cys	Lys	Leu	Gly	Asp	Ala	Ile	Ser	Leu
		20					25						30		
Ala	Tyr	Gly	Lys	Cys	Glu	Ser	Thr	Ala	Gly	Val	Ile	Asp	Leu	Ile	Phe
		35					40					45			
Pro	Leu	Ile	Thr	Lys	Lys	Leu	Arg	Ile	Asp	Tyr	Ile	Leu	Met	Tyr	Ser
	50					55					60				
Ile	Glu	Ser	Asn	Pro	Arg	Thr	Leu	Leu	Gln	Phe	Leu	Arg	Gln	Ile	Glu
65					70					75					80
Ser	Gly	Leu	Ala	Arg	Ser	Glu	Asp	Trp	Thr	Ala	Ala	Ser	Val	Glu	Ala
			85						90					95	
Ala	Leu	Arg	Ser	Val	Ala	Asp	Ser	Pro	Asp	Gly	Val	Gly	Trp	Glu	Arg
			100					105						110	

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Ala Gln Arg Leu Leu Lys Cys Cys Ile Leu Phe Ser Asp Ser Pro Leu  
 115 120 125

Gly Ile Val Glu Ser Ile Thr Phe Leu Gly Lys His Glu Thr Ser Ser  
 130 135 140

Arg Leu Arg Ser Ala Ala Ser Asn Val Glu Leu Ser His Leu Ile Asn  
 145 150 155 160

<210> SEQ ID NO 10  
 <211> LENGTH: 218  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(218)  
 <223> OTHER INFORMATION: putative glycosyl transferase

<400> SEQUENCE: 10

Met Lys Ser Thr Pro Thr Ile Asp Asn Thr Phe Ala Arg Lys Val Cys  
 1 5 10 15

Ile Asn Leu Asp Arg Arg Pro Asp Arg Trp Glu Ala Met Gln Arg Lys  
 20 25 30

Phe Ala Glu Gln Asn Ile Leu Thr Val Glu Arg Leu Pro Ala Val Asp  
 35 40 45

Ala Arg Leu Val Ser Val Pro Glu Ser Leu Ser His Met Arg Ala Gln  
 50 55 60

Asp Tyr Gly Cys Thr Met Ser His Leu Ala Ala Val Lys Gln Ala Lys  
 65 70 75 80

Ala Ala Gly Ala Arg Glu Val Leu Ile Phe Glu Asp Asp Ala Phe Phe  
 85 90 95

Asp Ala Asp Phe Ala Ala Arg Phe Pro Glu Phe Ile Ala Gln Val Pro  
 100 105 110

Asp Asp Trp His Met Leu Phe Leu Gly Ala Tyr His Phe Thr Pro Pro  
 115 120 125

Ile Pro Val Ala Pro Asn Ile Val Lys Ala Val Glu Thr Leu Thr Ala  
 130 135 140

His Ala Tyr Val Val Arg Asn Ser Leu Tyr Asp Ala Phe Ile Ala Ile  
 145 150 155 160

Asn Glu Asn Pro Pro Ala Ile Asn Asp Arg Asn Asn Leu Val Leu Gln  
 165 170 175

Gln Thr Phe Asn Cys Tyr Cys Phe Glu Pro Asn Leu Val Gly Gln Glu  
 180 185 190

Ser Gly Tyr Ser Asp Ile Met Asp Glu Val Met Pro Glu Lys Pro Leu  
 195 200 205

Thr Tyr Ser Met Pro Ile Pro Asp Gly Trp  
 210 215

<210> SEQ ID NO 11  
 <211> LENGTH: 3164  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(3164)  
 <223> OTHER INFORMATION: putative nonribosomal peptide synthetase

<400> SEQUENCE: 11

Met Gln Asp Asn Asn Val Leu Val Thr Asp His Arg Tyr Ala Ala Thr  
 1 5 10 15

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Ala Arg Phe Trp Arg Glu Ser Leu Ser Arg Val Ala Gly Val Tyr Gly  
20 25 30

Ile Ala Ala Tyr Ala Pro Ser Gln Gln Pro Gly Arg Pro Leu Thr Arg  
35 40 45

Ser Val Arg Leu Thr Pro Ala Ser Leu Asp Leu Leu Arg Arg Ile Gly  
50 55 60

Asp Gly Glu Leu Ala Glu Phe Ala Val Ala Ala Ala Gly Ile Ala Phe  
65 70 75 80

Leu Leu Trp Lys Tyr Phe Arg Ile Pro Val Thr Val Leu Gly Thr Pro  
85 90 95

Gly Leu Ala Gly His Pro Ser Ala Arg Ala Ala Ile Val Pro Leu Ile  
100 105 110

Ile Glu Val Arg Pro Asp Glu Arg Ile Glu Asp Tyr Leu Ser Arg Val  
115 120 125

Ala Gly Ile Val Glu Asp Ser Tyr Ala Glu Pro Arg Phe Pro Leu Glu  
130 135 140

Thr Leu Val Arg Asn Glu Lys Asp Met Ala Leu Ala Gln Leu Thr Lys  
145 150 155 160

Val Ala Leu Ala Asp Asp Arg Val His His Ala Pro Thr Gly Arg Asp  
165 170 175

Asp Asp Leu Gln Leu His Leu Arg Leu Ala Arg Gly Glu Ile Glu Leu  
180 185 190

Arg Tyr Ser Gly Ala Ile Glu Pro Phe Ile Ile Asp Gly Phe Ala Gly  
195 200 205

Ser Leu Ala Ala Val Leu Glu Ala Phe Glu His Leu Asp Gly Ala Val  
210 215 220

Gly Asp Ile Glu Ala Ala Pro Pro Glu Gln Gly Pro Leu Leu Ala Ala  
225 230 235 240

Phe Asn Glu Thr Ala Thr Ala Gly Pro Ser His Pro Thr Val Val Ala  
245 250 255

Met Phe Glu Ala Gln Val Ala Arg Thr Pro Thr Ala Pro Ala Leu Val  
260 265 270

Thr Asp Ser Ser Leu Met Thr Tyr Ala Asp Leu Asn Ala Arg Ala Asn  
275 280 285

Ser Leu Ala His His Leu Arg Glu His His Gly Val Gly Pro Glu Ser  
290 295 300

Leu Val Gly Ile Met Leu Asp Arg Ser Glu Trp Met Ile Val Ala Ile  
305 310 315 320

Leu Gly Ile Leu Lys Ala Gly Ala Ala Phe Val Pro Leu Asp Pro Ala  
325 330 335

Tyr Pro Ala Glu Arg Ile Asn His Ile Leu Gly Asp Thr Gly Leu Ser  
340 345 350

Leu Leu Val Thr Gln Ser Ser Gln Leu Ala Gln Trp Tyr Glu Phe Ser  
355 360 365

Gly Val Thr Leu Leu Leu Asp Gln Glu Leu Pro Gly Trp Gln Pro Leu  
370 375 380

Pro Asp Asn Pro Pro His Arg Ala Glu Pro Ala His Leu Ala Tyr Val  
385 390 395 400

Leu Tyr Thr Ser Gly Ser Thr Gly Lys Pro Lys Gly Cys Leu Leu Glu  
405 410 415

His Arg Asn Leu Ala His Tyr Ile Ala Trp Ala Ala Gly Tyr Tyr Phe  
420 425 430

Pro Glu Ser Thr Thr Gly Ser Phe Gly Leu Tyr Ser Ser Leu Cys Phe

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435					440					445					
Asp	Phe	Thr	Leu	Thr	Asn	Ile	Phe	Cys	Pro	Leu	Val	Arg	Gly	Lys	Thr
450					455					460					
Leu	Arg	Ile	Tyr	Pro	Gln	Ser	Glu	Ser	Ile	Asp	Thr	Ile	Leu	Ala	Arg
465					470					475					480
Met	Phe	Gln	Pro	Gly	Ser	Gly	Val	Asp	Thr	Leu	Lys	Leu	Thr	Pro	Thr
				485					490					495	
His	Ile	His	Leu	Leu	Glu	Tyr	Met	Asn	Leu	Ala	Arg	Ser	Gly	Val	Arg
			500					505					510		
Lys	Val	Ile	Val	Gly	Gly	Glu	Glu	Leu	Thr	Pro	Gln	His	Ile	Ala	Thr
		515					520					525			
Leu	Arg	Lys	Ile	Asp	Pro	Ala	Ile	Glu	Ile	Tyr	Asn	Glu	Tyr	Gly	Pro
530					535					540					
Thr	Glu	Ala	Thr	Val	Gly	Cys	Ile	Val	Glu	Arg	Val	Glu	Asp	Ala	Pro
545					550					555					560
Pro	Thr	Val	Leu	Ile	Gly	Arg	Pro	Ile	Ala	Asp	Thr	Arg	Val	Tyr	Met
				565					570					575	
Leu	Asp	Asp	Ala	Leu	Arg	Pro	Val	Pro	Leu	Gly	Val	Pro	Gly	Glu	Ile
			580					585					590		
Cys	Leu	Ala	Gly	Ala	Gly	Leu	Ala	Arg	Gly	Tyr	His	Gln	Arg	Pro	Asp
		595					600					605			
Val	Thr	Ala	Ala	Lys	Phe	Val	Glu	His	Pro	Phe	Pro	Gly	Glu	Ala	Arg
	610					615					620				
Ile	Tyr	Arg	Thr	Gly	Asp	Ile	Gly	Arg	Trp	Leu	Pro	Asp	Gly	Arg	Ile
625					630					635					640
Gln	Cys	Tyr	Gly	Arg	Val	Asp	His	Gln	Val	Lys	Ile	Arg	Gly	His	Arg
				645					650					655	
Val	Glu	Leu	Gly	Glu	Ile	Glu	Ala	Ala	Ile	Ala	Ala	His	Glu	Asp	Val
			660					665					670		
Val	Gly	Ala	Ala	Val	Met	Leu	Arg	Glu	Ser	Ala	His	Gly	Val	Arg	Lys
		675					680					685			
Leu	Ala	Ala	Tyr	Val	Lys	Gly	Ala	Ala	Ser	Leu	Ser	Val	Pro	Asn	Leu
	690					695					700				
Arg	Ala	Tyr	Leu	Ala	Gly	Lys	Leu	Pro	Asp	Tyr	Met	Val	Pro	Ser	Asp
	705					710					715				720
Ile	Ile	Pro	Ile	Ala	Glu	Phe	Pro	Leu	Asn	Ala	Asn	Gly	Lys	Leu	Asp
				725					730					735	
Arg	Pro	Ala	Leu	Leu	Ala	Leu	Glu	Pro	Ala	Ala	Ala	Pro	Glu	Glu	Ala
			740					745					750		
Pro	Leu	Asp	Ala	Thr	Pro	Ile	Gln	Arg	Glu	Leu	Val	Arg	Ile	Trp	Arg
		755					760					765			
Asp	Val	Leu	Asp	Asn	Pro	Ala	Val	Asp	Leu	Ala	Gly	Arg	Phe	Phe	Asp
	770					775					780				
Tyr	Gly	Gly	Asp	Ser	Leu	Gln	Ala	Met	Gln	Leu	Val	Ser	Arg	Ile	Trp
	785					790					795				800
Ser	Ser	Phe	Ser	Val	Glu	Ile	Gly	Ile	Asp	Ala	Ile	Phe	Glu	Leu	Gln
				805					810					815	
Thr	Ile	Ser	Ala	Val	Ser	Asp	Leu	Ile	Glu	Ala	Ser	Ser	Pro	His	Pro
				820					825					830	
Gly	Ser	Thr	Ala	Gly	Ala	Ile	Pro	Pro	Arg	Ser	Arg	Ala	Asn	Asp	Leu
		835					840					845			
Pro	Leu	Ser	Phe	Pro	Gln	Gln	Arg	Leu	Trp	Phe	Leu	Ala	Gln	Leu	Glu
	850					855					860				



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Gly Pro Ser Ala Thr Tyr Asn Ile Ser Ser Ala Leu Arg Phe Glu Gly  
 865 870 875 880  
 Glu Leu Asp Val Ala Arg Leu Arg Phe Ala Val Ser Glu Ile Ser Arg  
 885 890 895  
 Arg His Glu Ile Leu Arg Thr Thr Phe Pro Ala Val Asp Gly Arg Gly  
 900 905 910  
 Val Gln Arg Ile Ala Pro Pro Ala Pro Val Ala Leu Asp Val Val Asp  
 915 920 925  
 Val Ala Ser Glu Ser Asp Thr Leu Ala Leu Leu Ala Glu Glu Ala Asp  
 930 935 940  
 Arg Pro Phe Asp Leu Ala Ala Gly Pro Leu Tyr Arg Val Val Leu Tyr  
 945 950 955 960  
 Arg Val His Glu Arg Leu His Val Phe Gly Ile Val Met His His Ile  
 965 970 975  
 Val Ser Asp Ala Trp Ser Ser Gly Ile Leu Ile Gly Glu Leu Ala Ala  
 980 985 990  
 Leu Tyr Ala Gly Glu Ser Leu Pro Glu Leu Ala Val Gln Tyr Ala Asp  
 995 1000 1005  
 Tyr Ala Val Trp Gln His Glu Arg Leu Ala Ser Ala Asp Thr His  
 1010 1015 1020  
 Arg Glu Leu Ala Leu Leu Ser Ala Ala Leu Ala Asp Ala Pro Asp  
 1025 1030 1035  
 Leu Ile Glu Leu Pro Thr Asp Arg Pro Arg Pro Ala Val Gln Gln  
 1040 1045 1050  
 Phe Arg Gly Ala Val Leu Pro Phe Gln Leu Ser Ala Glu Arg Ala  
 1055 1060 1065  
 Asp Gly Leu Arg Ala Ile Ala Arg Ala Ser Gly Thr Ser Thr Phe  
 1070 1075 1080  
 Met Val Val Leu Ala Ala Tyr Ala Leu Leu Leu Ser Arg Tyr Ser  
 1085 1090 1095  
 Asn Gln Gln Asp Leu Val Ile Gly Ser Pro Ile Ala Asn Arg Arg  
 1100 1105 1110  
 Ser Ser Met Thr Glu Pro Leu Ile Gly Phe Phe Ala Asn Met Leu  
 1115 1120 1125  
 Ala Leu Arg Val Asp Leu Ser Gly Asn Pro Thr Phe Gly Asp Leu  
 1130 1135 1140  
 Leu Ala Arg Val Lys Arg Val Ala Leu Asp Gly Tyr Ser Arg Gln  
 1145 1150 1155  
 Glu Ile Pro Phe Glu Gln Val Val Asp Ser Leu Glu Leu Glu Arg  
 1160 1165 1170  
 Asn Leu Gly Arg Thr Pro Val Phe Gln Val Val Phe Ala Tyr Glu  
 1175 1180 1185  
 Lys Ala Gln Pro Arg Ala Val Ser Phe Pro Gly Leu Val Ala Thr  
 1190 1195 1200  
 Pro Val Ala Val Glu Thr His Thr Ala Lys Phe Asp Leu Thr Leu  
 1205 1210 1215  
 His Val Gln Asp Ala Asp Asp Gly Leu Ala Gly Ser Leu Glu Tyr  
 1220 1225 1230  
 Asn Leu Asp Leu Phe Asp Ala Ala Thr Ile Asp Arg Met Ala Glu  
 1235 1240 1245  
 His Phe Arg Thr Leu Val Asp Ala Val Ile Ala Asp Pro Asp Arg  
 1250 1255 1260

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Pro 1265	Leu	Gly	Ala	Leu	Ser	Leu	Ser	Asn	Asp	Ala	Glu	Arg	Asn	Leu	1270	1275
Leu 1280	Thr	Val	Glu	Trp	Asn	Arg	Thr	Asp	Thr	Asp	Phe	Gly	Glu	Asp	1285	1290
Ala 1295	Ala	Gln	Pro	Leu	His	Arg	Leu	Phe	Glu	Gln	Gln	Val	Glu	Arg	1300	1305
Thr 1310	Pro	Asp	Ala	Val	Ala	Ile	Val	Phe	Asp	Asp	Thr	Ala	Leu	Thr	1315	1320
Tyr 1325	Ala	Glu	Leu	Asn	Leu	Arg	Ala	Asn	Arg	Leu	Ala	His	His	Leu	1330	1335
Val 1340	Ala	Leu	Gly	Val	Gly	Pro	Asp	Ser	Leu	Val	Gly	Val	Ala	Met	1345	1350
Glu 1355	Arg	Ser	Leu	Asp	Met	Ser	Val	Ala	Leu	Leu	Ala	Ile	Leu	Lys	1360	1365
Ala 1370	Gly	Gly	Ala	Tyr	Val	Pro	Val	Asp	Pro	Asp	Tyr	Pro	Ala	Glu	1375	1380
Arg 1385	Val	Arg	Phe	Met	Ile	Asp	His	Ala	Gln	Leu	Arg	Trp	Leu	Leu	1390	1395
Thr 1400	Gln	Gln	His	Leu	His	Asp	Ala	Leu	Pro	Asp	Thr	Asp	Ala	His	1405	1410
Val 1415	Ile	Val	Val	Asp	Arg	Asp	Ser	Leu	Asp	Leu	Asp	Ala	Ala	Ala	1420	1425
Thr 1430	Ser	Asn	Pro	Ala	Pro	Ala	Leu	Asn	Gly	Asp	Asn	Leu	Ala	Tyr	1435	1440
Met 1445	Ile	Tyr	Thr	Ser	Gly	Ser	Thr	Gly	Arg	Pro	Lys	Gly	Ala	Leu	1450	1455
Asn 1460	Thr	His	Arg	Ala	Ile	Thr	Asn	Arg	Ile	Leu	Trp	Met	Gln	His	1465	1470
Ala 1475	Tyr	Ala	Leu	Asp	Ala	Asp	Asp	Ala	Val	Leu	Gln	Lys	Thr	Pro	1480	1485
Phe 1490	Ser	Phe	Asp	Val	Ser	Val	Trp	Glu	Leu	Phe	Trp	Pro	Leu	Val	1495	1500
Thr 1505	Gly	Ala	Arg	Leu	Val	Phe	Ala	Arg	Pro	Gly	Gly	Gln	Arg	Glu	1510	1515
Thr 1520	Asp	Tyr	Leu	Val	Glu	Leu	Ile	Glu	Arg	Glu	Arg	Ile	Thr	Thr	1525	1530
Ile 1535	His	Phe	Val	Pro	Ser	Met	Leu	Arg	Ala	Phe	Leu	Asp	His	Pro	1540	1545
Asp 1550	Leu	Asp	Ala	His	Cys	Ala	Ser	Leu	Arg	Arg	Val	Val	Cys	Ser	1555	1560
Gly 1565	Glu	Ala	Leu	Pro	His	Asp	Leu	Gln	Gln	Arg	Cys	Leu	Glu	Arg	1570	1575
Leu 1580	Asp	Val	Lys	Leu	Tyr	Asn	Leu	Tyr	Gly	Pro	Thr	Glu	Ala	Ala	1585	1590
Val 1595	Asp	Val	Thr	Ala	Trp	Glu	Cys	Arg	Arg	Asp	Asp	Pro	His	Arg	1600	1605
Ile 1610	Val	Pro	Ile	Gly	Arg	Pro	Ile	Ala	Asn	Thr	Arg	Leu	Tyr	Ile	1615	1620
Val 1625	Asp	Ala	Gln	Met	Gln	Pro	Thr	Pro	Ile	Gly	Val	Ala	Gly	Glu	1630	1635
Leu 1640	Leu	Ile	Gly	Gly	Thr	Pro	Val	Gly	Arg	Gly	Tyr	His	Gly	Glu	1645	1650
Pro 1650	Glu	Leu	Ser	Ala	Glu	Lys	Phe	Ile	Ala	Asp	Pro	Phe	Ser	Ala	1655	1660

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1655	1660	1665
Asp Pro Leu Ala Arg Leu Tyr Arg Thr Gly Asp Leu Ala Arg Tyr 1670 1675 1680		
Arg Pro Asp Gly Asn Ile Glu Phe Leu Gly Arg Ile Asp His Gln 1685 1690 1695		
Ile Lys Leu Arg Gly Leu Arg Ile Glu Pro Gly Glu Ile Glu Ala 1700 1705 1710		
Ala Leu Arg Ala His Pro Ser Val Asp Asp Cys Val Val Ile Ala 1715 1720 1725		
Lys Thr Glu Gly Ala Arg Thr Phe Leu Ile Ala Tyr Val Ala Thr 1730 1735 1740		
Ala Ala Pro Asp Ile Ala Asp Leu Arg Gly Tyr Leu Gly Gly Lys 1745 1750 1755		
Leu Ala Asp Tyr Met Val Pro Ser Gln Phe Phe Ala Leu Glu Ser 1760 1765 1770		
Leu Pro Met Leu Pro Asn Gly Lys Ile Asn Arg Lys Ala Leu Pro 1775 1780 1785		
Leu Pro Ala Asp Arg Gly Asp Ala Ala Gln Pro His Ala Pro Ala 1790 1795 1800		
Val Thr Pro Arg Glu Ile Leu Leu Ala Ser Ile Cys Ile Asp Val 1805 1810 1815		
Leu Gln Leu Pro Ser Val Gly Ile His Asp Asn Phe Phe Glu Leu 1820 1825 1830		
Gly Gly Asp Ser Ile Leu Ser Ile Gln Val Ile Ala Arg Ala Asn 1835 1840 1845		
Gln Ala Gly Leu Arg Val Thr Ala Lys Gln Leu Phe Gln Tyr Gln 1850 1855 1860		
Thr Ile Ala Gln Leu Ala Ala Ala Pro Glu Glu Arg Ala Ala Cys 1865 1870 1875		
Ala Pro Thr Leu Ser Pro Leu Gly Asp Ala Pro Leu Thr Pro Val 1880 1885 1890		
Gln His Trp Phe Phe Glu Gln Glu Ile Asp Ala Pro Ser His Tyr 1895 1900 1905		
Asn Gln Thr Val Leu Ile Gln Val Pro Ala Asp Ile Asp Ala Ser 1910 1915 1920		
Arg Leu Ala Asp Ala Phe Arg Gln Val Tyr Glu His His Asp Ala 1925 1930 1935		
Leu Arg Leu Arg Phe Ser His Asp Ala Gly Arg Trp Thr Gln Gln 1940 1945 1950		
Val Val Ala Gly Gly Glu Met Pro Ala Leu Phe Ala Lys Gln Val 1955 1960 1965		
Ile Ala Asp Asp Ala Gly Glu Arg Leu Ala Ala Met Arg Ala Ala 1970 1975 1980		
Ala Ala Asp Ala Glu Arg Gly Ile Asp Ile Thr His Gly Pro Leu 1985 1990 1995		
Leu Ala Ala Arg Leu Phe Cys Leu Ala Asp Glu Pro Leu Ala Arg 2000 2005 2010		
Leu Phe Val Ser Ile His His Leu Ala Val Asp Gly Val Ser Trp 2015 2020 2025		
Arg Val Leu Leu Glu Asp Leu His Ala Ala Tyr His Gly Gln Pro 2030 2035 2040		
Leu Pro Gly Lys Thr Thr Ser Phe Arg Glu Trp Ala Leu His Leu 2045 2050 2055		

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Gln	Gln	Leu	Ala	Arg	Ser	Pro	Ala	Ile	Gly	Asp	Glu	Ala	Arg	Leu
	2060					2065					2070			
Trp	Gln	Ala	Leu	Leu	Ala	Gln	Pro	Val	Glu	Pro	Met	Pro	Val	Asp
	2075					2080					2085			
Tyr	Pro	Gly	Thr	Gly	Ala	Ala	Asn	Asn	Ala	Val	Asp	Asp	Ala	Ser
	2090					2095					2100			
Ser	Val	Ser	Phe	Glu	Leu	Gly	Glu	Ala	Asp	Thr	Thr	Ala	Leu	Leu
	2105					2110					2115			
Arg	Arg	Leu	Pro	Arg	Ala	Tyr	Asp	Thr	Arg	Ile	Asn	Asp	Val	Leu
	2120					2125					2130			
Leu	Val	Ala	Leu	Ala	Gln	Ala	Cys	Ser	Met	Val	Thr	Gly	Asn	Thr
	2135					2140					2145			
Arg	Thr	Arg	Ile	Asp	Leu	Glu	Ser	His	Gly	Arg	His	Val	Ser	Asp
	2150					2155					2160			
Ala	Pro	Leu	Asp	Leu	Thr	Arg	Thr	Val	Gly	Trp	Phe	Thr	Ser	Ile
	2165					2170					2175			
Tyr	Pro	Val	Val	Leu	Asp	Ala	Asp	Ala	Met	His	Ala	Pro	Glu	Gln
	2180					2185					2190			
Ala	Leu	Arg	Ala	Ala	Arg	Gln	Gln	Leu	Arg	Arg	Ile	Pro	Ala	Asp
	2195					2200					2205			
Gly	Leu	Gly	Tyr	Ser	Leu	Leu	Arg	Tyr	Gln	Ser	Pro	Asp	Ala	Ala
	2210					2215					2220			
Val	Arg	Asp	Ser	Leu	Ala	Ala	Leu	Pro	Lys	Ala	Asp	Ile	Leu	Phe
	2225					2230					2235			
Asn	Tyr	His	Gly	Gln	Leu	Asp	Thr	Val	Leu	Arg	Gln	Ser	Asp	Gly
	2240					2245					2250			
Trp	Arg	Pro	Ala	Ala	Glu	Asp	Leu	Gly	Ser	Leu	Arg	Ala	Gly	Arg
	2255					2260					2265			
Ser	Gln	Arg	Thr	His	Ala	Phe	Glu	Ile	Val	Ala	Ala	Val	Ala	Asp
	2270					2275					2280			
Gly	Lys	Leu	Gln	Val	Asp	Trp	Arg	Tyr	Gly	Glu	Arg	Leu	His	Arg
	2285					2290					2295			
Arg	Gln	Thr	Val	Glu	Asn	Leu	Ala	Ala	His	Phe	Arg	Asp	Arg	Leu
	2300					2305					2310			
Leu	Asp	Phe	Ala	Ala	Ser	Val	Pro	Asp	Thr	Ala	Ala	Asp	Asp	Ile
	2315					2320					2325			
Glu	Asp	Ser	Tyr	Pro	Leu	Ser	Ser	Leu	Gln	Gln	Gly	Ile	Leu	Phe
	2330					2335					2340			
His	Ser	Leu	Tyr	Asp	Leu	Asp	Pro	Ala	Ala	Tyr	Phe	Gln	Gln	Phe
	2345					2350					2355			
Ser	Phe	Val	Val	Ser	Gly	Pro	Leu	Gln	Val	Pro	Ala	Leu	Arg	Gln
	2360					2365					2370			
Ala	Trp	Ala	Asn	Ala	Leu	Ala	Arg	His	Ala	Val	Leu	Arg	Thr	Ala
	2375					2380					2385			
Phe	Ala	Trp	Ala	Asp	Arg	Asp	His	Pro	Val	Gln	Thr	Val	Arg	His
	2390					2395					2400			
Thr	Val	Asp	Leu	Pro	Trp	Thr	Phe	Leu	Asp	Trp	Arg	His	Arg	Asp
	2405					2410					2415			
Ala	Ser	Arg	Arg	Ala	Gln	Asp	Phe	Asp	Ala	Phe	Leu	Ala	Asp	Asp
	2420					2425					2430			
Arg	Arg	Arg	Gly	Phe	Asp	Leu	Gln	Arg	Ala	Pro	Leu	Phe	Arg	Cys
	2435					2440					2445			

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Thr 2450	Leu	Ile	Gln	Glu	Thr	Asp 2455	Thr	Arg	His	Arg	Phe 2460	Cys	Trp	Ser
Ala 2465	His	His	Ile	Ile	Leu	Asp 2470	Gly	Trp	Ser	Thr	Ala 2475	Thr	Leu	Met
Lys 2480	Glu	Val	Phe	Asp	Asp	Tyr 2485	Leu	Ser	Leu	Ala	Arg 2490	Thr	Gly	Met
Pro 2495	Ala	Val	Ala	Ala	Ser	Ala 2500	Pro	Gly	Tyr	Arg	Ala 2505	Tyr	Ile	Asp
Trp 2510	Leu	Ala	Arg	His	Pro	Arg 2515	Ser	Ala	Asp	Glu	Thr 2520	Trp	Trp	Arg
Ala 2525	Glu	Leu	Ala	Gly	Phe	Lys 2530	Ala	Ala	Thr	Pro	Val 2535	Ala	Ala	Ser
Pro 2540	Ala	Arg	Gln	Ala	Thr	Gly 2545	Asp	Ala	Pro	Arg	Gln 2550	Asp	Lys	Arg
Arg 2555	Thr	Gln	Gln	Phe	Leu	Leu 2560	Asp	Glu	Ala	Leu	Ala 2565	Ala	Arg	Leu
Gln 2570	Thr	Leu	Thr	Arg	Thr	His 2575	Arg	Val	Thr	Leu	Asn 2580	Val	Leu	Ile
Arg 2585	Ala	Val	Trp	Ala	Leu	Val 2590	Leu	Arg	Arg	His	Ala 2595	Gly	Thr	Asp
Asp 2600	Val	Val	Phe	Gly	Val	Thr 2605	Val	Ser	Gly	Arg	Pro 2610	Pro	Met	Leu
Asp 2615	Gly	Val	Glu	Ser	Ile	Val 2620	Gly	Leu	Phe	Ile	Asn 2625	Thr	Leu	Pro
Leu 2630	Arg	Leu	Arg	Ile	Ala	Pro 2635	Glu	Arg	Pro	Phe	Ile 2640	Glu	Trp	Leu
Ala 2645	Glu	Val	His	Ala	Ala	Gln 2650	Thr	Ala	Met	Glu	Pro 2655	His	Ser	Tyr
Ser 2660	Ser	Leu	Val	Asp	Ile	Gln 2665	Ser	Trp	Ser	Glu	Leu 2670	Pro	Ala	Gly
Asp 2675	Ser	Leu	Phe	Asp	Ser	Leu 2680	Leu	Val	Phe	Glu	Asn 2685	Phe	Pro	Val
Ala 2690	Ala	Ala	Pro	Asp	Leu	Gly 2695	Pro	Asp	Asp	Ile	Glu 2700	Ile	Leu	Asp
Thr 2705	Arg	Ala	Phe	Ala	Glu	Ser 2710	Asn	Tyr	Pro	Leu	Thr 2715	Leu	Thr	Val
His 2720	Pro	Asn	Glu	Arg	Ile	Gly 2725	Phe	His	Ile	Ser	His 2730	Asp	Ala	His
Arg 2735	Ile	Ala	Pro	Glu	Val	Val 2740	Arg	Gln	Met	Leu	Asp 2745	Thr	Leu	Arg
Thr 2750	Leu	Leu	Glu	Arg	Phe	Ala 2755	Glu	Asn	Pro	Gly	Gln 2760	Leu	Thr	Gly
Gln 2765	Leu	Ala	Asp	Pro	Pro	Ala 2770	Ala	Asp	Gly	Arg	Pro 2775	Ser	Ala	Pro
Arg 2780	Ser	Gly	Ala	Gly	Pro	Ala 2785	Ile	Glu	Ala	Ala	Ala 2790	Gly	Ala	Ala
Ala 2795	Ala	Ala	Arg	Ala	Val	Ala 2800	His	Ala	Ala	Asp	Glu 2805	Ser	Thr	Leu
Leu 2810	Glu	Ile	Trp	Arg	Arg	Ile 2815	Phe	Lys	Arg	Asp	Asp 2820	Ile	Ala	Val
Ser 2825	Asp	Asn	Tyr	Phe	Asp	Leu 2830	Gly	Gly	His	Ser	Ile 2835	Ile	Ala	Ile
Gln 2840	Leu	Met	Ala	His	Val	Glu 2845	Lys	Ala	Phe	Asp	Arg 2850	Arg	Leu	Pro

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2840	2845	2850
Ile Ser Cys Leu Phe Glu Asn Pro Thr Ile Glu Lys Leu Ala Ala 2855 2860 2865		
Ala Leu Ala Ala Lys Glu Pro Ser Ala Pro Ala Gly Gly Leu Val 2870 2875 2880		
Pro Ile Arg Asp Gly Gly Pro Ala Ala Pro Leu Phe Leu Leu Pro 2885 2890 2895		
Gly Ala Gly Gly Asn Val Val Tyr Phe Arg Pro Leu Ala Asn His 2900 2905 2910		
Leu Ser Gly Ala His Ala Ile His Gly Leu Glu Ala Leu Gly Leu 2915 2920 2925		
Asp Gly Ala Cys Glu Pro Leu Thr Arg Val Glu Asp Ile Ala Ala 2930 2935 2940		
Arg His Ile Glu Arg Ile Trp Pro Leu Val Gly Ala Gly Pro Tyr 2945 2950 2955		
Tyr Leu Ala Gly His Ser Phe Gly Ala His Val Ala Leu Glu Met 2960 2965 2970		
Ser Arg Gln Leu Val Ala Lys Gly Ala Asp Val Lys Leu Leu Ala 2975 2980 2985		
Ile Phe Asp Ala Ser Ala Pro Ile Asp Ser Ser Ala Ala Thr Tyr 2990 2995 3000		
Trp Gln Asp Trp Asp Asp Thr Glu Trp Leu Val Ala Ile Ala His 3005 3010 3015		
Glu Ile Gly Thr Phe Leu Gly Thr Asp Leu Gln Val Thr Arg Ala 3020 3025 3030		
Asp Leu Val His Leu Asp Pro Asp Gly Gln Ala Gly Leu Ile Leu 3035 3040 3045		
Glu Arg Ile Gly Asp Arg Gly Ser Trp Phe Ala Asp Ala Gly Ser 3050 3055 3060		
Asp Arg Leu Arg Ala Tyr Leu Arg Val Tyr Gln Ala Asn Phe Lys 3065 3070 3075		
Ser His Tyr Ala Pro His Ala Thr Pro Leu Pro Val Pro Ile Ala 3080 3085 3090		
Leu Phe Arg Ser Thr Glu Arg Asp Pro Gly Asp Tyr Ala Pro Ser 3095 3100 3105		
Ser Glu Ile Ala Gln Leu Arg Leu Asp Ala Thr Trp Gly Trp Ser 3110 3115 3120		
Arg Phe Ser Ala His Pro Val Ala Val Thr Asp Val Pro Gly Asp 3125 3130 3135		
His Leu Thr Met Leu Leu Asp Pro His Ala Gly Val Leu Ala Ala 3140 3145 3150		
His Val Asn Ser Phe Leu Glu Lys Thr Pro Ser 3155 3160		

&lt;210&gt; SEQ ID NO 12

&lt;211&gt; LENGTH: 3021

&lt;212&gt; TYPE: PRT

&lt;213&gt; ORGANISM: Burkholderia contaminans

&lt;220&gt; FEATURE:

&lt;221&gt; NAME/KEY: MISC\_FEATURE

&lt;222&gt; LOCATION: (1)..(3021)

&lt;223&gt; OTHER INFORMATION: putative nonribosomal peptide synthetase

&lt;400&gt; SEQUENCE: 12

Met Gln Glu Gly Met Leu Phe His Ala Val His Glu Pro Gly Ser Arg  
 1 5 10 15

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Ser Ser Phe Asn Gln Leu Ser Cys Arg Ile Thr Gly Ser Leu Asp Pro  
 20 25 30  
 Ala Leu Phe His Ala Ala Trp Gln Gln Leu Ile Asp Arg His Pro Val  
 35 40 45  
 Met Arg Thr Ser Phe His Trp Glu Glu Phe Asp Lys Pro Met Gln Val  
 50 55 60  
 Val His Ala Arg Ala Thr Leu Pro Trp Val Gln Asp Asp Trp Leu Asp  
 65 70 75 80  
 Leu Pro Glu His Glu Gln Arg Ser Arg Trp Arg Ala His Leu Asp Asn  
 85 90 95  
 Asp Leu Ala Glu Gly Phe Ala Leu Asp Arg Ala Pro Leu Val Arg Cys  
 100 105 110  
 Arg Leu Val Arg Val Ala Ala Asp Ala Tyr Leu Phe Ser Trp Ser His  
 115 120 125  
 His His Ile Leu Ala Asp Gly Trp Cys Leu Ser Leu Val Ile Glu Glu  
 130 135 140  
 Ile Phe Glu Val Tyr Gly Ala Leu Ala Arg Gly Val Ser Pro Ala Leu  
 145 150 155 160  
 Pro Pro Val Arg Pro Tyr Arg Asp Tyr Ile Gln Trp Leu Gln Gln His  
 165 170 175  
 Glu Pro Gln Ala Ala Gln Gln Tyr Trp Thr Arg Tyr Leu Glu Gly Phe  
 180 185 190  
 Arg Thr Pro Thr Pro Leu Pro Thr Ala Ala Arg Ala Gly Ala Asp Glu  
 195 200 205  
 Arg Phe Gly Gln Gly Leu Ala Gln Val Gln Ala Asp Leu Ser Ala Asp  
 210 215 220  
 Leu Ser Ala Arg Leu Arg Gln Phe Ala Ala Arg His His Val Thr Leu  
 225 230 235 240  
 Asn Thr Leu Ala Gln Ala Ala Trp Ala Leu Val Leu Ser Arg Tyr Ser  
 245 250 255  
 Gly Glu Thr Asp Val Val Phe Gly Ala Val Val Ser Gly Arg Gly Ala  
 260 265 270  
 Asn Leu Pro Gly Ile Glu Thr Met Leu Gly Leu Phe Ile Asn Thr Val  
 275 280 285  
 Pro Val Arg Val Arg Val Asp Pro Arg Gln Pro Leu Val Pro Trp Leu  
 290 295 300  
 Lys Met Ile Gln Ala Arg Val Ala Ala Arg Ala Pro Phe Glu His Thr  
 305 310 315 320  
 Pro Leu Pro Asp Ile Gln Arg Cys Ser Asp Val Pro Pro Thr Ala Pro  
 325 330 335  
 Leu Phe Glu Ser Asn Ile Thr Phe Met Asn Tyr Pro Leu Asp Ala Ser  
 340 345 350  
 Leu Thr His Gly Ala His Gly Leu Ala Val Asp Glu Val Gln Leu Tyr  
 355 360 365  
 Asn Arg Ala Asp Ile Pro Leu Glu Phe Val Val Thr Ala Arg Asp Asp  
 370 375 380  
 Trp Lys Met Glu Leu Ser Phe Asp Pro Arg Arg Phe Asp Glu Asp Thr  
 385 390 395 400  
 Met Gln Arg Met Leu Gly His Val Ala Ala Thr Leu Asp Ala Phe Ala  
 405 410 415  
 Ala Asp Pro Asn Arg Leu Leu Gly Arg Val Pro Ile Leu Pro Asp Ala  
 420 425 430

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Glu Arg Arg Gln Leu Leu Glu Thr Phe Asn Asp Thr Ala Val Pro Phe  
 435 440 445  
 Asp Ala Ala Leu Thr Val Val His Arg Leu Glu Gln Ala Ala Ala Asp  
 450 455 460  
 His Pro Glu Arg Pro Ala Val Glu Tyr Arg Asp Gly Val Leu Ser Ala  
 465 470 475 480  
 Gly Glu Leu Asn Ala Arg Ala Asn Arg Ile Ala His Arg Leu Leu Ala  
 485 490 495  
 Ala Ala Asp Leu Gly Pro Asp Ala Leu Val Ala Ile Cys Met His Arg  
 500 505 510  
 Ser Ala Gln Leu Met Glu Ala Ile Leu Ala Val Trp Lys Cys Gly Ala  
 515 520 525  
 Ala Tyr Ile Pro Val Asp Pro Asn Tyr Pro Val Ala Arg Ile Arg Thr  
 530 535 540  
 Ile Leu Glu Asp Ser Gly Ala Ala Leu Val Ile Thr Cys Asp Gly Leu  
 545 550 555 560  
 Leu Pro Pro Glu Leu Ala Gly Ile Ala Leu Val Val Ser Leu Asp Ala  
 565 570 575  
 Ala Thr Asp Ala Val Asp Asp Ser Asn Pro Gly Arg Pro Val Ser Pro  
 580 585 590  
 Asp Ser Leu Ala Tyr Val Ile Tyr Thr Ser Gly Ser Thr Gly Lys Pro  
 595 600 605  
 Lys Gly Ala Met Val Glu His Ala Gly Met Leu Asn His Met Leu Ala  
 610 615 620  
 Glu Ile Asp Glu Phe Ser Ile Ser Ala Ser Ser Val Ile Ala Gln Thr  
 625 630 635 640  
 Ala Pro His Cys Phe Asp Ile Ser Val Trp Gln Phe Phe Thr Ala Pro  
 645 650 655  
 Leu Val Gly Gly Lys Thr Val Ile Val Asp Asp Asp Cys Ile Arg Asp  
 660 665 670  
 Pro Ala Arg Phe Val Ala Tyr Leu Glu Thr Thr Arg Ile Ser Ile Leu  
 675 680 685  
 Glu Leu Val Pro Ser Tyr Leu Ser Ala Val Leu Asp Arg Ala Ser Glu  
 690 695 700  
 Arg Pro Ala Leu Met Arg His Leu Arg His Leu Leu Val Thr Gly Glu  
 705 710 715 720  
 Met Val Ser Pro Ala Leu Val Lys Gln Trp Phe Asp Val Phe Pro Ala  
 725 730 735  
 Ile Pro Leu Val Asn Ala Tyr Gly Pro Ala Glu Ala Ser Asp Asp Val  
 740 745 750  
 Ala Gln His Arg Met Thr Gly Ala Pro Ser Thr Pro Tyr Val Pro Val  
 755 760 765  
 Gly Lys Pro Ile Arg Asn Val Arg Leu Tyr Val Val Asp Pro Gln Met  
 770 775 780  
 Asn Leu Cys Pro Ile Gly Ile Pro Gly Glu Leu Cys Val Ser Gly Val  
 785 790 795 800  
 Ala Val Gly Arg Gly Tyr Leu Asn Asn Glu Ala Ala Thr Gln Asp Ala  
 805 810 815  
 Phe Val Glu Asp Pro Phe His Pro Gln Arg Gly Val Arg Leu Tyr Arg  
 820 825 830  
 Thr Arg Asp Ile Gly Cys Tyr Leu Pro Asp Gly Thr Ile Val Leu His  
 835 840 845  
 Gly Arg Lys Asp His Gln Leu Lys Ile Arg Gly Tyr Arg Ile Glu Leu



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850				855				860							
Gly	Glu	Ile	Asp	Gln	Arg	Arg	Leu	Ala	Asp	His	Ser	Arg	Lys	Leu	Arg
865					870					875				880	
Gln	Ala	Ala	Ala	Leu	Asp	Tyr	Arg	Asp	Glu	Ala	Gly	Arg	Ala	Ala	Leu
				885					890					895	
Cys	Ala	Tyr	Val	Ala	Phe	Arg	Asp	Gly	Ala	Ser	Leu	Ser	Asp	Ala	Gly
				900					905					910	
Ile	Ala	Ala	Ala	Leu	Ser	Ala	Thr	Leu	Pro	Asp	Tyr	Met	Val	Pro	Gly
				915					920					925	
Ile	Tyr	Val	Val	Leu	Asp	Ala	Leu	Pro	Leu	Ser	Gly	Asn	Gly	Lys	Ile
				930					935					940	
Asp	Arg	Asn	Ala	Leu	Pro	Pro	Leu	Asp	Arg	Ala	Arg	Leu	Ala	Ala	Thr
945					950					955				960	
Ala	His	Ala	Pro	Thr	Pro	Pro	Arg	Thr	Pro	Thr	Glu	Thr	Leu	Leu	Cys
				965						970				975	
Arg	Ile	Trp	Gly	Glu	Ala	Leu	Gly	Ile	Pro	Ser	Pro	Gly	Ile	His	Asp
				980					985					990	
Asn	Leu	Phe	Ala	Leu	Gly	Gly	Asp	Ser	Ile	Leu	Ser	Met	Arg	Ile	Val
				995					1000					1005	
Ser	Leu	Ala	Ala	Lys	Ala	Gly	Leu	Lys	Leu	Thr	Thr	Arg	Leu	Ile	
				1010					1015					1020	
Phe	Gln	His	Pro	Thr	Val	Ala	Glu	Leu	Ala	Ala	Val	Ala	Thr	Arg	
				1025					1030					1035	
Gly	Thr	Val	Gly	Ala	Ala	Ala	Phe	Val	Ala	Ser	Ser	Gly	Pro	Leu	
				1040					1045					1050	
Pro	Leu	Thr	Pro	Ile	Gln	Lys	Arg	Phe	Phe	Ala	Gln	Gly	Lys	His	
				1055					1060					1065	
Asp	Pro	Asp	Gln	Tyr	Asn	Gln	Ala	Val	Leu	Leu	Asp	Val	Pro	Ala	
				1070					1075					1080	
Asp	Leu	Asp	Pro	Val	Leu	Leu	Arg	Gln	Ala	Leu	Arg	His	Ala	Val	
				1085					1090					1095	
Lys	Trp	His	Asp	Ala	Leu	Arg	Leu	Arg	Phe	Arg	Glu	Gly	Glu	Ser	
				1100					1105					1110	
Gly	Trp	Thr	Gln	Glu	Val	Val	Asp	Asp	Pro	Glu	Ile	Pro	Val	Val	
				1115					1120					1125	
Val	Ser	Asp	Ile	Ala	Arg	Asp	Gln	Leu	Ala	Gln	Tyr	Val	Ala	Gln	
				1130					1135					1140	
Ser	His	Ala	Ser	Leu	Asn	Leu	Ala	Asp	Gly	Pro	Val	Val	Arg	Ala	
				1145					1150					1155	
Asp	Leu	Phe	Arg	Val	Asp	Glu	Gly	Arg	Ser	Leu	Arg	Leu	Leu	Leu	
				1160					1165					1170	
Val	Ala	His	His	Leu	Val	Val	Asp	Gly	Val	Ser	Trp	Gly	Ala	Leu	
				1175					1180					1185	
Leu	Glu	Thr	Val	Tyr	Asp	Ala	Tyr	Thr	Arg	Leu	Arg	Asn	Gly	Lys	
				1190					1195					1200	
Ala	Pro	Glu	Phe	Ala	Gly	Gly	Ser	Ala	Thr	Trp	Thr	Ala	Trp	Thr	
				1205					1210					1215	
Arg	Ala	Ile	Ser	Thr	Trp	Ala	Gly	Ser	Gly	Ala	Ala	Asp	Ala	Asp	
				1220					1225					1230	
Leu	Ala	His	Trp	Gln	Ala	Leu	Ala	Arg	Ala	Ala	Leu	Pro	Gly	Leu	
				1235					1240					1245	
Pro	Leu	Asp	Arg	Asp	Ala	Pro	Ala	Asp	Ala	Asn	Thr	Val	Ser	Ser	
				1250					1255					1260	

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Ala Asp	Thr Ile Val Val	Glu	Leu Gly Glu Ala	Ala	Thr Thr Ala
1265		1270		1275	
Leu Leu	Gly Ala Ala Pro	Arg	Ala Tyr Asp Ala	Gln	Val Asn Asp
1280		1285		1290	
Val Leu	Leu Ala Ala Leu	Ala	Arg Ala Val Ser	Glu	Trp Ser Gly
1295		1300		1305	
Cys Ala	Asp Val Leu Leu	Asp	Leu Glu Ala His	Gly	Arg Glu Glu
1310		1315		1320	
Leu Ile	Asp Ala Leu Asp	Ile	Ser Arg Thr Val	Gly	Trp Phe Thr
1325		1330		1335	
Ser Val	Phe Pro Val Leu	Leu	Thr Val Asp Ala	Gly	Ser His Asp
1340		1345		1350	
Pro Ala	Ser Leu Val Ala	Ser	Val Arg Thr Arg	Leu	Arg Ala Val
1355		1360		1365	
Pro Asn	Ala Gly Ile Thr	Tyr	Gly Leu Leu Leu	Asp	Arg Leu Asp
1370		1375		1380	
Gly Pro	Leu Pro Gln Pro	Arg	Leu Gln Phe Asn	Tyr	Leu Gly Gln
1385		1390		1395	
Thr Asp	Gln Leu Phe Thr	Ala	Ala Arg Asp Trp	Lys	Gln Ala Ala
1400		1405		1410	
Glu Pro	Ser Gly Asp Gly	Arg	Asn Ala Asn Gln	Leu	Arg Glu His
1415		1420		1425	
Leu Leu	Asp Ile Asn Ala	Tyr	Val Thr Gly Asn	Arg	Leu His Val
1430		1435		1440	
Ala Trp	Glu Phe Ser Arg	Ala	Cys His Asp Thr	Ala	Thr Ile Leu
1445		1450		1455	
Arg Val	Ala Gln Ala Tyr	Ile	Ala Ala Leu Glu	Thr	Leu Val Ala
1460		1465		1470	
Gly His	Ala Val Pro Ser	Ala	Ser Thr Arg Pro	Ala	Thr Ala Leu
1475		1480		1485	
Pro Gln	Ala Pro Ala Pro	Ala	Ser Val Ser Pro	Asp	Glu Ile Ala
1490		1495		1500	
Asp Val	Tyr Pro Leu Thr	Pro	Thr Gln Gln Gly	Met	Leu Phe His
1505		1510		1515	
Ser Leu	Tyr Glu Pro Ala	Ser	Asp Ala Tyr Phe	Ser	Ser Leu Asn
1520		1525		1530	
Phe Arg	Ile Asp Gly Ala	Leu	Asp Val Glu Arg	Phe	Arg Arg Ala
1535		1540		1545	
Trp Glu	Thr Val Ala His	Arg	His Asp Ile Leu	Arg	Thr Ser Phe
1550		1555		1560	
His Trp	Glu Asp Ile Glu	Ser	Pro Val Gln Val	Val	His Arg Arg
1565		1570		1575	
Ile Asp	Leu Pro Trp His	Asp	Glu Asp Leu Arg	Ala	Ala Ser Ala
1580		1585		1590	
Ala Glu	Ala Glu Gln Arg	Trp	Glu Ala Tyr Val	Ala	Gln Asp Arg
1595		1600		1605	
Ala Arg	Gly Phe Asp Phe	Thr	Arg Ala Pro Leu	Met	Arg Leu Ala
1610		1615		1620	
Leu Phe	Arg Val Gly Glu	His	Ala Trp Arg Phe	His	Trp Ser His
1625		1630		1635	
His His	Ile Leu Leu Asp	Gly	Trp Ser Ser Ala	Arg	Leu Leu Ser
1640		1645		1650	

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Asp	Val	Ala	Ala	Ala	Tyr	Gln	Ala	Pro	Pro	Ala	Glu	Gly	Ala	Pro	1655	1660	1665
Gln	Arg	Asp	Ala	Pro	Pro	Ala	Phe	Ala	Gly	Tyr	Val	Arg	Trp	Leu	1670	1675	1680
Ala	Arg	Gln	Asp	Ala	Ala	Ala	Ala	Gln	Arg	Phe	Trp	Lys	Thr	Lys	1685	1690	1695
Leu	Ala	Asp	Phe	Pro	Ala	Thr	Thr	Pro	Leu	Val	Leu	Gly	Arg	Pro	1700	1705	1710
Glu	Leu	Asp	Gly	Thr	Ala	Ala	Pro	Gly	Ala	Tyr	Val	Glu	Glu	Pro	1715	1720	1725
Leu	Leu	Leu	Ser	Glu	Ser	Asp	Thr	Gln	Arg	Leu	Val	Ala	Phe	Ala	1730	1735	1740
Gln	Ser	Arg	Arg	Leu	Thr	Leu	Asn	Thr	Leu	Ala	Gln	Gly	Ala	Trp	1745	1750	1755
Ala	Gln	Leu	Leu	Ser	Arg	Tyr	Ser	Gly	Glu	Ser	Asp	Val	Val	Phe	1760	1765	1770
Gly	Thr	Ile	Val	Ser	Gly	Arg	Pro	Ala	Ser	Leu	Pro	Ala	Ser	Asp	1775	1780	1785
Glu	Met	Val	Gly	Leu	Phe	Ile	Asn	Thr	Leu	Pro	Val	Arg	Val	Arg	1790	1795	1800
Ile	Asp	Ala	Arg	Pro	Thr	Ser	Ala	Trp	Leu	Ala	Gln	Leu	Gln	Met	1805	1810	1815
Asp	Leu	Ala	Gln	Gln	Glu	Asp	Tyr	Ala	His	Tyr	Pro	Leu	Ala	Asp	1820	1825	1830
Ile	Gln	Lys	Phe	Ala	Gly	Leu	Pro	Pro	Gly	Val	Pro	Leu	Phe	Glu	1835	1840	1845
Ser	Leu	Leu	Ile	Phe	Gln	Asn	Tyr	Pro	Val	Glu	Glu	Ala	Leu	Ala	1850	1855	1860
Asp	Ala	Leu	Pro	Gly	Leu	Arg	Ile	Gly	Ala	Phe	Glu	Val	Ser	Asp	1865	1870	1875
Pro	Asn	Asn	Tyr	Pro	Leu	Thr	Leu	Val	Val	Thr	Pro	Gly	Lys	Arg	1880	1885	1890
Leu	Ser	Leu	Gln	Val	Leu	Tyr	Asp	Asp	Gly	Arg	Phe	Asp	Arg	Asp	1895	1900	1905
Thr	Ile	Val	Arg	Leu	Leu	Arg	His	Val	Glu	Thr	Leu	Leu	Thr	Gly	1910	1915	1920
Leu	Ala	Gly	Ala	Glu	Asp	Arg	Pro	Asn	Arg	Ser	Val	Pro	Leu	Leu	1925	1930	1935
Ala	Ala	Ala	Glu	Arg	Asp	Ala	Ile	Leu	Leu	Gly	Trp	Asn	Asp	Thr	1940	1945	1950
Phe	Ala	Pro	Val	Pro	Ser	Asp	Arg	Thr	Leu	Pro	Glu	Leu	Ile	Glu	1955	1960	1965
Ala	Val	Ala	Ala	Ala	His	Pro	Glu	Arg	Val	Ala	Val	Arg	Cys	Gly	1970	1975	1980
Thr	Glu	Val	Arg	Thr	Tyr	Arg	Asp	Leu	Val	Glu	Gly	Ala	Asn	Arg	1985	1990	1995
Ile	Ala	Ala	His	Leu	Leu	Gln	Thr	Ala	Pro	Leu	Gln	Pro	Asp	Asp	2000	2005	2010
Arg	Ile	Ala	Val	Trp	Met	Pro	Arg	Ser	Pro	Leu	Met	Leu	Glu	Thr	2015	2020	2025
Ile	Leu	Ala	Ile	Trp	Lys	Cys	Gly	Ala	Ala	Tyr	Val	Pro	Val	Asp	2030	2035	2040
Pro	Ala	Tyr	Pro	Ala	Gln	Arg	Val	Glu	Thr	Ile	Leu	Thr	Leu	Ala			

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2045	2050	2055
Arg Pro Ala Val Ile Val Thr Thr Asp Cys Val Pro Pro Pro Ala 2060 2065 2070		
Leu Ala Ser Ile Pro Leu Val Asp Pro Ala Arg Leu Pro Asp Arg 2075 2080 2085		
Arg Gly Ala Glu Ala Pro Ala Pro Val Thr Pro Arg Cys Arg Pro 2090 2095 2100		
Ala Asp Leu Ala Tyr Val Ile Phe Thr Ser Gly Ser Thr Gly Gln 2105 2110 2115		
Pro Lys Gly Ala Met Val Glu His Arg Gly Met Leu Asn His Val 2120 2125 2130		
Leu Ala Met Ala Arg Arg Val Gly Leu Gly Ala Gln Ser Ala Val 2135 2140 2145		
Ala Gln Thr Ala Ser His Cys Ser Asp Ile Ser Val Trp Gln Cys 2150 2155 2160		
Phe Ala Ala Leu Ala Ser Gly Gly Thr Thr Val Ile Tyr Pro Asp 2165 2170 2175		
Ala Val Ile Leu Glu Pro Ala Arg Leu Ile Asp Ser Leu His Arg 2180 2185 2190		
Asp Arg Ile Thr Ala Met Gln Phe Val Pro Ser Tyr Leu Ala Thr 2195 2200 2205		
Phe Leu Gly Glu Leu Glu Arg His Ala Ala Pro Ala Phe Pro His 2210 2215 2220		
Leu Asp Thr Leu Leu Thr Ile Gly Glu Thr Leu Gln Pro Ala Thr 2225 2230 2235		
Ala Gln Ala Trp Phe Arg Leu Asn Pro Ala Val Arg Leu Ile Asn 2240 2245 2250		
Ala Tyr Gly Pro Thr Glu Ala Ser Asp Ser Val Ala His Tyr Cys 2255 2260 2265		
Leu Thr Arg Ala Pro Asp Gly Pro Ala Ile Pro Ile Gly Arg Pro 2270 2275 2280		
Ile Glu Asn Leu Arg Leu Tyr Val Val Asp Ala Asp Met Asn Pro 2285 2290 2295		
Cys Pro Ala Gly Val Lys Gly Glu Ile Cys Ile Gly Gly Val Gly 2300 2305 2310		
Val Gly Arg Gly Tyr Leu Phe Asp Glu Ala Arg Thr Arg Ala Val 2315 2320 2325		
Phe Arg Asp Asp Pro Phe Ser Pro Glu Pro Gly Ala Arg Leu Tyr 2330 2335 2340		
Arg Thr Gly Asp Ile Gly Cys Phe Gly Ala Asp Gly Asn Leu His 2345 2350 2355		
Phe Phe Gly Arg Arg Asp Phe Gln Val Lys Ile Arg Gly Tyr Arg 2360 2365 2370		
Ile Glu Leu Gly Glu Ile Glu Ala Ala Leu Thr Ser Leu Ala Gly 2375 2380 2385		
Ile Ser His Ala Val Val Val Ala Arg Glu Thr Ser Asp Ala Glu 2390 2395 2400		
Met Thr Leu Cys Gly Tyr Ala Ser Gly Thr Gly Trp Thr Pro Gln 2405 2410 2415		
Arg Val Arg Asp Ala Leu Arg Asp Thr Leu Pro Ala His Met Val 2420 2425 2430		
Pro Asp Thr Val Met Leu Leu Pro Ala Leu Pro Val Met Pro Asn 2435 2440 2445		

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Gly	Lys	Ile	Asn	Arg	Ala	Ala	Leu	Pro	Leu	Pro	Asp	Ala	Ala	Ser
2450						2455					2460			
Val	Pro	Asp	Gly	Val	Arg	Ala	Glu	Pro	Arg	Thr	Pro	Val	Glu	Ala
2465						2470					2475			
Ala	Leu	Leu	Arg	Leu	Phe	Ala	Glu	Val	Leu	Gly	Arg	Arg	Pro	Asn
2480						2485					2490			
Gly	Val	Asp	Asp	Asp	Phe	Phe	Glu	His	Gly	Gly	Gln	Ser	Leu	Lys
2495						2500					2505			
Ala	Ile	Gln	Met	Val	Ser	Arg	Ile	Pro	Arg	Ala	Ala	Leu	Asn	Val
2510						2515					2520			
Ala	Val	Ala	Asp	Ile	Phe	His	Ala	Pro	Thr	Pro	Arg	Ala	Leu	Ala
2525						2530					2535			
Gln	Arg	Leu	Ala	Ala	Met	Pro	Val	Asp	Gly	Ala	Ala	Asp	Asp	Asp
2540						2545					2550			
Ala	Ile	Ile	Pro	Ala	Leu	Ala	Ala	Gln	Pro	Ser	Tyr	Ala	Val	Ser
2555						2560					2565			
Arg	Ala	Gln	Lys	Arg	Ile	Trp	Leu	Ala	Ser	Arg	Gly	Ala	Asp	Pro
2570						2575					2580			
Ser	Thr	Tyr	Asn	Met	Ala	Gly	Ala	Leu	Gln	Leu	Asp	Gly	Ala	Val
2585						2590					2595			
Asp	Thr	Ala	Arg	Leu	Val	Arg	Ala	Phe	Asp	Thr	Leu	Val	Asp	Arg
2600						2605					2610			
His	Glu	Ser	Leu	Arg	Thr	Val	Phe	Ala	Met	Ile	Glu	Gly	Glu	Leu
2615						2620					2625			
Arg	Gln	Arg	Val	Leu	Ser	Arg	Glu	Ala	Ser	Gly	Phe	Arg	Val	Glu
2630						2635					2640			
Gln	Arg	Asp	Leu	Ala	Asp	Asp	Ala	Gly	Pro	Gln	Ala	Ile	Asp	Ala
2645						2650					2655			
Leu	Ile	Arg	Ala	Glu	Cys	Glu	Gln	Pro	Phe	Asp	Leu	Ala	Ser	Gly
2660						2665					2670			
Pro	Leu	Phe	Arg	Val	Lys	Leu	Val	Arg	Leu	Ser	Gln	Glu	Lys	His
2675						2680					2685			
Leu	Leu	Leu	Leu	Asn	Met	His	His	Val	Ile	Ser	Asp	Ala	Trp	Ser
2690						2695					2700			
Ile	Arg	Val	Leu	Thr	Asp	Asp	Leu	His	Ala	Leu	Tyr	Ala	Gly	Arg
2705						2710					2715			
Asp	Leu	Pro	Pro	Leu	Ser	Ile	Gln	Tyr	Arg	Asp	Tyr	Ala	Ala	Trp
2720						2725					2730			
His	Asn	Ala	Ser	Leu	Ala	Gly	Pro	Arg	Ala	Ala	Ala	His	Arg	Ala
2735						2740					2745			
Tyr	Trp	Leu	Glu	Gln	Leu	Ala	Pro	Pro	Leu	Pro	Arg	Leu	Gln	Leu
2750						2755					2760			
Ala	Ser	Asp	Phe	Pro	Arg	Pro	Glu	Arg	Leu	Gly	His	Ala	Gly	Gln
2765						2770					2775			
Thr	Leu	Glu	Val	Glu	Leu	Pro	Gln	Pro	His	Ala	Ala	Glu	Leu	Ala
2780						2785					2790			
Thr	Leu	Ala	Arg	Ala	His	His	Thr	Ser	Leu	His	Ala	Val	Leu	Leu
2795						2800					2805			
Ala	Ser	Phe	Cys	Val	Leu	Met	His	Arg	Tyr	Thr	Gly	Arg	Glu	Asp
2810						2815					2820			
Ile	Val	Ile	Gly	Ser	Val	Ser	Ala	Gly	Arg	Asp	Ser	Glu	Gln	Leu
2825						2830					2835			

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Glu Ser Gln Val Gly Val Tyr Leu Asn Thr Val Val Leu Arg Val  
 2840 2845 2850  
 Pro Val Arg Lys Ser Ala Thr Val Ala Glu Val Ile Asp Gly Val  
 2855 2860 2865  
 Ala Lys Ala Ser Ala Gln Ala Leu Glu His Ala Ser Tyr Pro Phe  
 2870 2875 2880  
 Asp Val Leu Leu Glu Asp Leu Lys Ile Arg Thr Pro Ala Asn His  
 2885 2890 2895  
 Phe Pro Ile Phe Asp Ile Gln Val Asn His Val Ser Met Pro Ala  
 2900 2905 2910  
 Pro Gln Pro Gly Leu Arg Ile Thr Asp Ile Ser Pro Ala Asp Thr  
 2915 2920 2925  
 Thr Ala Lys Phe Asp Leu Ser Phe Gln Val Val Glu Ser Glu Gly  
 2930 2935 2940  
 Arg His Leu Ile Gln Phe Ile Tyr Asn Thr His Leu Phe Arg Pro  
 2945 2950 2955  
 Ser Thr Ile Ala Ala Met Arg Asp Arg Leu Leu Ala Ile His Asp  
 2960 2965 2970  
 Val Phe Arg Arg Asp Pro Ala Thr Pro Val Asp Arg Ile Pro Leu  
 2975 2980 2985  
 Ser Asp Glu Ala Pro Ala Ala Gly Pro Arg Val Arg Val Gly Leu  
 2990 2995 3000  
 Arg Leu Lys Arg Ala Pro Ala Val Thr Ala Asp Asp Ala Leu Glu  
 3005 3010 3015  
 Glu Lys Thr  
 3020

<210> SEQ ID NO 13  
 <211> LENGTH: 1306  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(1306)  
 <223> OTHER INFORMATION: putative nonribosomal peptide synthetase

<400> SEQUENCE: 13

Met Ser Glu Leu Asn Leu Asn Ala Leu Ser Thr Ser Gly Gln Tyr Gln  
 1 5 10 15  
 Glu His Val Ala Phe Trp Asn Asp Ala Leu Gly Arg Ile Asp Glu Asp  
 20 25 30  
 Phe Arg Leu Gln Gln Ala Trp Gln Ala Tyr Ala Leu Pro Leu Gly Pro  
 35 40 45  
 Glu Pro Ala Leu Thr Phe Ala Leu Asp Gly Asp Ala Ala Gln Val Leu  
 50 55 60  
 Glu Arg Leu Ala Ala Gly Asn Glu Leu Gly Ala Phe Val Val Leu Leu  
 65 70 75 80  
 Ala Ala Leu Phe Arg Val Leu Gly Arg Tyr Asn Gly Ala Ala Gly Leu  
 85 90 95  
 Phe Val Ala Ser Pro Gln Leu Ile Val Glu Pro Ala Ser Gly Cys Ala  
 100 105 110  
 Glu Pro Val Pro Leu Leu Asp Ala Gly Glu Pro Gly Pro Thr Val Arg  
 115 120 125  
 Ala Tyr Leu Asn Gln Leu Arg Asp Ser Val Gln Arg Ser Tyr Ser Tyr  
 130 135 140  
 Gln Asp Phe Pro Ile Ala Ala Leu Ala His Lys Leu His Gly Glu Arg

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145	150	155	160
Arg Ala Thr Asn Val Gly Val Arg Phe Asp Gly Leu His Glu Ala Trp	165	170	175
Ala Ala Ala Asp Tyr Asp Leu Ser Ile Glu Ile Arg His Arg Glu Arg	180	185	190
Tyr Glu Ile Val Leu Thr Gly Arg Pro Thr Val Phe Thr Leu His Tyr	195	200	205
Leu Gln His Val Ala Arg His Leu Arg Asn Val Val Ala Gly Phe Gly	210	215	220
Ala Leu Asp Ala Pro Leu Asp Thr Val Ser Leu Leu Asp Asp Glu Glu	225	230	235
Arg Ala Arg Leu Arg Ser His Ala Ala Pro Val Ala Val Gln Gly Thr	245	250	255
Phe Leu Glu Gln Phe Ala Gln Arg Val Ala Ala Ala Pro Asp Ser Val	260	265	270
Ala Val Val Thr Ala Asp Ala Ser Leu Thr Tyr Ala Glu Leu Asp Asp	275	280	285
Gln Ala Ser Arg Leu Ala Ser Phe Leu Leu Ala Glu Tyr Ala Ile Glu	290	295	300
Arg Gly Asp Val Val Gly Val Val Ala Asp Arg Ser Glu Arg Trp Ile	305	310	315
Val Gly Met Leu Gly Ala Leu Lys Ala Gly Ala Val Tyr Leu Pro Leu	325	330	335
Asp Pro Glu Phe Pro Arg Glu Arg Leu Arg Phe Met Ile Glu Asp Ala	340	345	350
Lys Val Lys Ala Leu Leu Thr His Ser Glu His Leu Pro Leu Leu Ala	355	360	365
Asp Phe Trp Ala Ile Pro Met Phe Ala Leu Asp Phe Gln Leu Asp Thr	370	375	380
Leu Ala Pro Ala Ser Ala Ser Ala Gln Val Glu Val Arg Pro Asp Asp	385	390	395
Ala Ala Tyr Ile Ile Tyr Thr Ser Gly Ser Thr Gly Val Pro Lys Gly	405	410	415
Val Val Leu Glu His Ala Gly Leu Leu Asn Met Ala Gln Tyr His Val	420	425	430
Asp Ala Phe Gly Phe Asp Ser Ala Asp Arg Phe Val Gln Phe Tyr Ser	435	440	445
Pro Gly Phe Asp Gly Ser Ile Met Glu Ile Phe Val Thr Leu Leu Ala	450	455	460
Gly Ala Arg Leu Val Leu Ala Lys Thr Ala Val Ile Arg Asp Val Pro	465	470	475
Arg Phe Val Asp Tyr Ile Ala Gln Gln Gly Val Thr Thr Val Asn Ala	485	490	495
Thr Pro Ala Tyr Leu Ala Ala Leu Asp Trp His Ala Leu Gly Ala Val	500	505	510
Lys Arg Val Ile Ser Ala Gly Asp Ser Ala Arg Val Ala Asp Leu Arg	515	520	525
Glu Leu Ala Arg Thr Arg Thr Cys His Asn Ser Tyr Gly Pro Thr Glu	530	535	540
Ala Thr Val Cys Ile Ala Asp Tyr Val Val Asp Pro Ala Ile Thr Tyr	545	550	555
Gly Ala Arg Leu Pro Val Gly Arg Pro Ile His Asn Thr His Leu Tyr	565	570	575

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Leu	Leu	Asp	Glu	His	Gly	Ala	Leu	Ala	Pro	Glu	Gly	Cys	Ala	Gly	Glu
			580					585						590	
Ile	Cys	Val	Ser	Gly	Ile	Ala	Leu	Ala	Arg	Gly	Tyr	Val	Gly	Arg	Asp
		595					600						605		
Asp	Leu	Thr	Ala	Ala	Ala	Phe	Val	Ala	His	Pro	Phe	Glu	Ala	Gly	Glu
	610					615						620			
Arg	Leu	Tyr	Arg	Thr	Gly	Asp	Leu	Gly	Val	Trp	Leu	Pro	Asp	Gly	Asn
	625				630					635					640
Leu	Glu	Val	Thr	Gly	Arg	Arg	Asp	Thr	Gln	Val	Lys	Ile	Arg	Gly	Tyr
				645					650						655
Arg	Ile	Glu	Met	Gly	Glu	Ile	Glu	Ala	Ala	Leu	Arg	Gln	His	Ala	Gly
			660					665						670	
Val	Ala	Asp	Ala	Ile	Val	Phe	Val	Arg	Glu	Asp	Thr	Pro	Gln	His	Lys
		675					680						685		
Gln	Leu	Val	Ala	Cys	Val	Ala	Thr	Ala	Thr	Ala	Ser	Val	Ala	Ser	Leu
	690					695					700				
Arg	Glu	His	Leu	Lys	Glu	Arg	Leu	Pro	Glu	Phe	Met	Val	Pro	Ala	Ser
	705				710					715					720
Ile	Val	Thr	Leu	Glu	Arg	Leu	Pro	Leu	Thr	Pro	Asn	Gly	Lys	Pro	Asp
				725					730						735
Arg	Lys	Ala	Leu	Ala	Ala	Leu	Glu	Leu	Ala	Pro	Ala	Pro	Ser	Glu	Thr
			740					745						750	
Ala	Tyr	Val	Ala	Pro	Ala	Asn	Asp	Val	Glu	Ala	Arg	Leu	Gly	Arg	Ile
		755					760						765		
Trp	Cys	Asp	Val	Leu	Gly	Arg	Glu	Pro	Ile	Gly	Val	His	Asp	Asn	Phe
	770					775						780			
Phe	Glu	Leu	Gly	Gly	Asp	Ser	Ile	Leu	Ile	Ile	Gln	Val	Met	Ser	Leu
	785				790					795					800
Ala	Gln	Gln	Val	Gly	Leu	Lys	Phe	Thr	Ala	Asp	Gln	Phe	Phe	Ala	His
				805					810						815
Pro	Thr	Ile	Ala	Glu	Leu	Ala	Gln	Val	Ala	Thr	Glu	Ala	Pro	Ser	Ile
				820					825					830	
Arg	Ile	Ala	Gln	Glu	Pro	Val	Val	Gly	Pro	Ala	Pro	Leu	Thr	Pro	Ile
		835					840					845			
Gln	His	Trp	Phe	Phe	Ala	Gln	Asp	Val	Ala	Asp	Pro	His	His	Tyr	Asn
	850						855				860				
Gln	Ser	Thr	Met	Ile	Glu	Val	Pro	Ala	Ser	Leu	Arg	Pro	Asp	Thr	Ile
	865					870					875				880
Glu	Arg	Ala	Leu	Ala	Ala	Val	Ala	Thr	His	His	Asp	Ala	Leu	Arg	Leu
				885					890						895
Ser	Phe	Ala	Cys	Val	Ala	Gly	Val	Trp	Gln	Gln	Ser	His	Ala	Ala	Pro
			900					905						910	
Pro	Leu	Ala	Ile	Pro	Leu	Gly	Val	Thr	Ser	Leu	Ala	Asp	Ala	Ala	Pro
		915						920						925	
Ala	Ala	Arg	Gln	Ala	Ala	Met	Leu	Ala	Thr	Ala	Thr	Gly	Met	Gln	Glu
		930					935								940
Ser	Phe	Thr	Leu	Ser	Ala	Pro	Pro	Leu	Leu	Arg	Ala	His	Leu	Phe	Gln
	945					950					955				960
Phe	Gly	Pro	Asp	Ala	Pro	Gln	Arg	Leu	Leu	Ala	Val	Ala	His	His	Leu
				965					970						975
Val	Ile	Asp	Gly	Val	Ser	Trp	Arg	Ile	Leu	Phe	Glu	Asp	Leu	Tyr	Thr
			980						985						990



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Ala	Cys	Arg	Gln	Leu	Glu	Ala	Gly	Asp	Ala	Val	Gln	Leu	Pro	Ala	Arg
	995						1000					1005			
Thr	Thr	Ala	Trp	Arg	Asp	Trp	Ser	Thr	Arg	Leu	Ser	Gly	Leu	Gly	
	1010					1015					1020				
Ala	Thr	Ala	Leu	Asp	Gly	Leu	Gly	Leu	Asp	Tyr	Trp	Leu	Gln	Gly	
	1025					1030					1035				
Asn	Ala	Gly	Glu	Pro	Ala	Cys	Phe	Asp	Asp	Met	Pro	Ala	Gly	Thr	
	1040					1045					1050				
Val	Ala	Glu	Ala	Gly	Ser	Thr	Ile	Val	Glu	Phe	Asp	Ala	Gln	Gln	
	1055					1060					1065				
Thr	Leu	Ala	Leu	Leu	Gln	Asp	Val	Pro	Arg	Ala	Phe	Asn	Thr	Gln	
	1070					1075					1080				
Ile	Asn	Glu	Val	Leu	Leu	Thr	Ala	Leu	Leu	Leu	Ala	Phe	Gly	Asp	
	1085					1090					1095				
Trp	Thr	Gly	Asn	Ala	Ser	Leu	Val	Val	Asp	Leu	Glu	Gly	His	Gly	
	1100					1105					1110				
Arg	Glu	Asp	Ile	Phe	Asp	Gly	Val	Asp	Thr	Ser	Arg	Thr	Ile	Gly	
	1115					1120					1125				
Trp	Phe	Thr	Thr	His	Tyr	Pro	Val	Cys	Leu	Asn	Ala	Gly	Asp	Ala	
	1130					1135					1140				
Thr	Val	Ala	Val	Asp	Ala	Leu	Arg	His	Val	Lys	Glu	Gln	Leu	Arg	
	1145					1150					1155				
Ala	Val	Pro	Met	Arg	Gly	Leu	Gly	Tyr	Gly	Ile	Ala	Arg	Tyr	Leu	
	1160					1165					1170				
Gly	His	Asp	Ala	Gly	Ile	Ala	Ala	Ala	Leu	Glu	Arg	Gln	Pro	Pro	
	1175					1180					1185				
Ala	Pro	Val	Arg	Phe	Asn	Tyr	Leu	Gly	Gln	Val	Asp	Arg	Val	Leu	
	1190					1195					1200				
Pro	Asp	Asp	Thr	Gly	Trp	Lys	Pro	Val	Leu	Asp	Phe	Gln	Ser	Pro	
	1205					1210					1215				
Glu	His	Ser	Pro	Arg	Ala	Arg	Arg	Gly	His	Leu	Phe	Glu	Ile	Asp	
	1220					1225					1230				
Gly	Met	Val	Phe	Asp	Gly	Arg	Leu	Arg	Leu	Thr	Trp	His	Tyr	Asn	
	1235					1240					1245				
Arg	Glu	Ala	Cys	Ala	Pro	Gly	Val	Ile	Glu	Gln	Leu	Thr	Gln	Cys	
	1250					1255					1260				
Tyr	Arg	Ser	Arg	Leu	Leu	Ser	Ile	Val	Ala	Ala	Gly	Gly	Asp	Gly	
	1265					1270					1275				
Pro	Arg	Ala	Leu	Ser	Pro	Ser	Asp	Phe	Pro	Ala	Ala	Arg	Ile	Ser	
	1280					1285					1290				
Gln	Glu	Ala	Leu	Asp	Ala	Leu	Val	Ser	Arg	Ile	Lys	Ser			
	1295					1300					1305				

<210> SEQ ID NO 14  
 <211> LENGTH: 538  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(538)  
 <223> OTHER INFORMATION: putative beta-lactamase domain protein  
  
 <400> SEQUENCE: 14

Met	Thr	Ile	Ser	Ser	Ser	Ala	Gln	Val	Tyr	Leu	Arg	Gln	Asn	Ile	Gln
1			5					10					15		
Phe	Glu	Pro	Leu	Ile	Asn	Ser	Trp	Tyr	Ala	Trp	Tyr	His	Thr	Leu	Pro

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20				25				30							
Pro	Leu	Thr	Ala	Ala	Leu	Asn	Val	Ala	Glu	Arg	Phe	Leu	Pro	Leu	Leu
	35					40					45				
Lys	Ser	Tyr	Ala	Ala	Ser	Pro	Met	Met	His	Ala	Ala	Ala	Cys	Lys	Asp
	50				55						60				
Pro	Ala	Met	Arg	Gly	Gly	Pro	Phe	Leu	Asp	Leu	Gly	Gly	Gln	Arg	Val
	65				70					75					80
Asp	Glu	Ile	Arg	Thr	Leu	Ile	Glu	Gln	Thr	Thr	Gln	Arg	Ala	Thr	Arg
			85						90					95	
Gln	Leu	Glu	Leu	Ala	Lys	Ala	Tyr	Lys	Ala	Phe	Ser	Thr	Leu	Leu	Leu
			100						105					110	
Glu	Arg	Ala	Thr	Gly	Met	Ala	Ser	Asp	Pro	Leu	Tyr	Pro	Glu	Ile	Pro
			115				120						125		
Glu	Val	Leu	Lys	Gly	Tyr	Val	Glu	Ile	Tyr	Tyr	Asp	Leu	Asn	His	Asn
	130					135					140				
Pro	Ser	Phe	Arg	Val	Phe	Glu	Ser	Leu	Leu	Tyr	Ala	Ser	Pro	Phe	Tyr
	145				150						155				160
Ala	Arg	Asp	Ala	Gln	Ser	Ile	Ala	Leu	Ser	Ala	Ile	Glu	Glu	His	Thr
			165							170					175
Pro	Arg	Pro	Phe	Ile	Leu	Ser	Thr	Pro	Arg	Leu	Arg	Asp	Glu	Arg	Thr
			180						185					190	
Val	Phe	Ser	Asn	Met	Ala	Phe	Asp	Asp	Arg	Ala	Leu	Asp	Thr	Leu	Phe
		195					200						205		
Arg	Met	Arg	Asp	Thr	Pro	Gly	Ser	Tyr	Ala	Lys	Ile	Val	Asp	Leu	Met
	210					215					220				
Arg	Val	Glu	Glu	Lys	Asp	Glu	Pro	Leu	Phe	Arg	Ser	Phe	Phe	Val	Glu
	225				230					235					240
Glu	Ala	Pro	Ala	Pro	Lys	Pro	Asp	Arg	Ser	Phe	Asp	Gly	Asp	Asp	Ile
			245							250				255	
Arg	Ile	Arg	Tyr	Tyr	Gly	His	Ala	Cys	Val	Leu	Ile	Gln	Ser	Arg	Gly
		260							265					270	
Val	Ser	Ile	Leu	Ile	Asp	Pro	Val	Ile	Ser	Tyr	Gly	Tyr	Asp	Thr	Ala
		275					280						285		
Leu	Pro	Arg	Tyr	Thr	Phe	Ala	Asp	Leu	Pro	Asp	Gln	Ile	Asp	Tyr	Val
	290					295					300				
Leu	Ile	Thr	His	Ser	His	His	Asp	His	Ile	Val	Leu	Glu	Thr	Leu	Leu
	305				310					315					320
Gln	Leu	Arg	His	Lys	Val	Lys	Thr	Val	Val	Val	Gly	Arg	Asn	Leu	Asp
			325							330				335	
Gly	Phe	Pro	Gln	Asp	Pro	Ser	Met	Glu	Leu	Ala	Leu	Arg	Lys	Leu	Gly
			340						345					350	
Phe	Asp	Asp	Val	Leu	Glu	Val	Arg	Asp	Ala	Gln	Glu	Ile	Lys	Val	Pro
		355					360						365		
Gly	Gly	Ala	Ile	Thr	Ala	Ile	Pro	Phe	Met	Gly	Glu	His	Asn	Asp	Leu
	370					375					380				
Ala	Ile	His	Ser	Lys	Gln	Ser	Phe	Met	Ile	Arg	Phe	Gly	Ser	Arg	Ser
	385				390					395					400
Val	Leu	Cys	Ile	Ala	Asp	Ser	Cys	Asn	Leu	Asp	Pro	Arg	Leu	Tyr	Glu
			405							410				415	
His	Val	Phe	Arg	Leu	Ala	Gly	Lys	Pro	Asp	Thr	Leu	Phe	Val	Gly	Met
			420						425				430		
Glu	Thr	Glu	Gly	Ala	Pro	Pro	Ser	Trp	Val	Tyr	Gly	Pro	Leu	Phe	Pro
		435					440						445		

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Lys Ala Leu Pro Arg Asp Ile Asp Gln Ser Arg Arg Ala Arg Gly Cys  
 450 455 460

Gln Phe Gly Glu Ala Ala Ala Leu Val Asp Asp Phe Ala Phe Asn Ala  
 465 470 475 480

Ala Tyr Val Tyr Ala Met Gly Gln Glu Pro Trp Leu Asn His Leu Leu  
 485 490 495

Asp Asn Thr Phe Asp Glu Asn Ser Pro Ser His Ile Gln Ser Thr Gln  
 500 505 510

Phe Val Ala His Cys Lys Ala Lys Gly Ile Ala Ser Glu Ile Leu Tyr  
 515 520 525

Ala Thr Arg Glu Ile Val Leu Cys Gln Asn  
 530 535

<210> SEQ ID NO 15  
 <211> LENGTH: 4469  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(4469)  
 <223> OTHER INFORMATION: putative beta-ketoacyl synthase nonribosomal  
 peptide synthetase

<400> SEQUENCE: 15

Met Asn Ala Lys Ala Thr His Ala Leu Lys Ala Ala Leu Asp Glu Leu  
 1 5 10 15

Arg Leu Arg Arg Ala Glu Ile Ala Ala Leu Arg Ser Asp Arg Asn Glu  
 20 25 30

Pro Ile Ala Val Ile Gly Met Ala Cys Arg Phe Pro Gly Arg Ser Asp  
 35 40 45

Thr Pro Asp Ala Phe Trp Gln Leu Leu Asp Gly Ala His Asp Ala Val  
 50 55 60

Thr Glu Val Pro Gly Glu Arg Trp Asp Ile Asp Arg Tyr Tyr Asp Pro  
 65 70 75 80

Asp Pro Ser Thr Pro Gly Lys Met Ala Thr Arg His Gly Ala Phe Leu  
 85 90 95

Glu Arg Val Asp Gln Phe Asp Ala Ala Phe Phe Gly Ile Ala Pro Arg  
 100 105 110

Glu Ala Thr Tyr Leu Asp Pro Gln Gln Arg Leu Leu Leu Glu Val Ala  
 115 120 125

Trp Glu Ala Leu Glu Asn Ala His Leu Ala Pro Glu Arg Phe Arg Gln  
 130 135 140

Ser Ala Thr Gly Val Tyr Val Gly Ile Thr Cys Phe Asp His Ala Ile  
 145 150 155 160

Gln Val Ser Asn Ala Ser Met Pro Ser Ser Tyr Ala Gly Thr Gly  
 165 170 175

Ser Ala Leu Asn Met Ala Ala Gly Arg Leu Ser Phe Val Leu Gly Leu  
 180 185 190

Thr Gly Pro Ser Met Ala Ile Asp Thr Ala Cys Ser Ser Ser Leu Val  
 195 200 205

Cys Leu His Leu Ala Cys Glu Ser Leu Arg Ser Arg Glu Ser Asn Met  
 210 215 220

Ala Leu Ala Gly Gly Val Asn Leu Met Leu Ser Pro Glu Val Met Val  
 225 230 235 240

Ser Phe Ser Gln Ala Arg Met Leu Ser Pro Asp Gly Arg Cys Lys Thr  
 245 250 255

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Phe Asp Ala Ala Ala Asp Gly Tyr Val Arg Gly Glu Gly Cys Gly Met  
 260 265 270  
 Val Val Leu Lys Arg Leu Ala Asp Ala Leu Ala Asp Gly Asp Arg Val  
 275 280 285  
 Leu Gly Ile Val Arg Gly Thr Ala Val Asp Gln Gly Gly Ala Gly Gly  
 290 295 300  
 Gly Leu Thr Val Pro Ser Arg Asp Ser Gln Glu Arg Val Ile Arg Arg  
 305 310 315 320  
 Ala Leu Asn Gln Ala Gly Leu Ala Pro Gly Asp Val Ser Tyr Val Glu  
 325 330 335  
 Ala His Gly Thr Gly Thr Ser Leu Gly Asp Pro Ile Glu Val Glu Ala  
 340 345 350  
 Leu Ala Gly Val Tyr Gly Pro Gly Arg Ala Ala Asn Glu Pro Leu Val  
 355 360 365  
 Ile Gly Ser Val Lys Thr Asn Ile Gly His Leu Glu Ser Ala Ser Gly  
 370 375 380  
 Ile Ala Gly Leu Ile Lys Val Leu Leu Ser Phe Glu His Asp Arg Ile  
 385 390 395 400  
 Pro Ala His Leu His Phe Thr Gln Pro Asn Pro His Thr Pro Trp Gln  
 405 410 415  
 Asp Ile Pro Ile Arg Val Ala Ala Asp Pro Val Ala Trp Arg Arg Gly  
 420 425 430  
 Glu Arg Arg Arg Ile Ala Gly Val Ser Ala Phe Gly Phe Ser Gly Thr  
 435 440 445  
 Asn Ala His Ala Ile Val Glu Glu Pro Pro Val Ala Pro Ala His Ala  
 450 455 460  
 Ala Gln Arg Ala Leu Leu Leu Leu Ser Ala Arg Ser Glu Ala Ala Leu  
 465 470 475 480  
 Ala Ala Leu Val Pro Arg Tyr Glu Arg Ala Ile Ala Gly Ala Thr Pro  
 485 490 495  
 Gln Glu Leu Ala Ala Ile Cys Arg Ala Ala Ala Thr Gly Arg Ser His  
 500 505 510  
 Tyr Pro Phe Arg Ala Ala Tyr Val Ser Gly Ala Arg Val Ala Ser Ala  
 515 520 525  
 Ala Ala Pro Arg Thr Gly Lys Ala Leu Arg Met Gly Phe Gly Phe Gly  
 530 535 540  
 Val Pro Asp Thr Gly Val Ala His Ala Leu His Ala Ser Glu Pro Leu  
 545 550 555 560  
 Phe Arg Asp Ala Phe Ala Arg Cys Ser Val Pro Leu Asp Ala Leu Glu  
 565 570 575  
 Thr Asp Ala Gly Arg Phe Ala Ile Gln Phe Ala Trp Ala Glu Leu Trp  
 580 585 590  
 Lys Gly Trp Gly Leu Arg Pro Ala Val Val Ser Gly His Gly Ile Gly  
 595 600 605  
 Glu Tyr Val Ala Ala Cys Val Ala Gly Val Val Ser Val Ala Asp Ala  
 610 615 620  
 Leu Arg Leu Val Ala Ala Arg Ser Asp Ala Glu Ala Leu Arg Ala Val  
 625 630 635 640  
 Leu Arg Asp Met Pro Leu Ala Arg Pro Ser Val Arg Leu Ile Ser Gly  
 645 650 655  
 Tyr Leu Gly Thr Asp Val Thr Asp Glu Val Thr His Pro Gln Tyr Trp  
 660 665 670

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Leu Gln Leu Ala Gly Ala Ser Asp Gln Ala Asp Ala Ser His Pro Pro  
           675                                  680                                  685

Glu Gly Leu Ala Asp Gly Trp Leu Pro Pro Pro Cys Ala Gly Asp Ala  
   690                                  695                                  700

Leu Glu Arg Ala Leu Ala Ala Leu Tyr Val Gln Gly Ala Gln Phe Asp  
   705                                  710                                  715                                  720

Trp Arg Ala Leu Phe Pro Ala Pro Ala Gln Pro Ala Thr Thr Leu Pro  
                                   725                                  730                                  735

Asn Tyr Pro Phe Glu Arg Gln Arg Phe Ser Leu Glu Lys Ile Pro Ser  
                   740                                  745                                  750

Pro Ile Val Gly Met Asp Ala Gly Ser Ile Asp Ala Ala Leu Arg His  
                   755                                  760                                  765

Leu Lys Ser Ser Gly Lys Tyr Pro Glu Asp Met Leu Asn Ala Phe Pro  
   770                                  775                                  780

Asp Leu Leu Arg Thr Ala Phe Ala Ala Ala Glu Thr Val Ala Ser Asn  
   785                                  790                                  795                                  800

Ala His Pro Leu Tyr His Val Val Trp Glu Gln Gln Ala Ala Met Pro  
                                   805                                  810                                  815

Ala Ala Pro Ala Ala Ala Asp Ala Ser Pro Trp Leu Ile Phe Ala Asp  
                                   820                                  825                                  830

Ala Ser Gly Val Gly Glu Arg Leu Ala Ala Leu Leu Arg Ala Arg Gly  
                   835                                  840                                  845

Ala Ser Cys Ser Leu Val Arg Pro Gly Ile Asp Tyr Val Thr Gly Ala  
   850                                  855                                  860

Glu Ala Gly Trp Gln Val Ala Pro Glu Arg Pro Asp Asp Phe Val Arg  
   865                                  870                                  875                                  880

Leu Leu Asn Glu Thr Ala Ala Ser Gly Gln Arg Ile Val Phe Leu Trp  
                                   885                                  890                                  895

Ala Leu Asp Glu Ala Val Gly Glu Thr Arg Met Ser Ala Ala Leu Leu  
                   900                                  905                                  910

His Leu Val His Ala Leu Val Gly Ser Glu Arg Glu Trp Thr Pro Ser  
                   915                                  920                                  925

Thr Arg Pro Arg Ile Ser Val Val Thr Arg Asp Ala Val Glu Ala Gly  
   930                                  935                                  940

Glu Ala Pro His Val Ser Gly Leu Ala Gln Ala Ala Leu Ser Gly Leu  
   945                                  950                                  955                                  960

Ala Arg Gly Ala Met Ile Glu His Pro Glu Trp Phe Gly Thr Ala Ile  
                                   965                                  970                                  975

Asp Leu Asp Pro Ala Ala Pro Glu Asn Glu Thr Gln Ala Leu Leu Gln  
                   980                                  985                                  990

Glu Met Leu Gly Glu Ser Arg Glu Glu Gln Val Ala Leu Arg His Gly  
                   995                                  1000                                  1005

Ala Arg His Val Ala Arg Leu Ser Pro Leu Ala Pro Ala Glu Thr  
   1010                                  1015                                  1020

Ala Ala Leu Pro Val Asp Pro Asp Ala Ala Tyr Leu Ile Thr Gly  
   1025                                  1030                                  1035

Gly Phe Gly Ala Leu Gly Leu His Thr Ala Arg Trp Leu Ala Ala  
   1040                                  1045                                  1050

Arg Gly Ala Gly Thr Leu Ile Leu Val Gly Arg Gln Gly Ala Ala  
   1055                                  1060                                  1065

Ser Asp Glu Ser Gln Arg Ala Ile Ala Glu Leu Arg Glu Arg Asn  
   1070                                  1075                                  1080

Val Thr Leu Arg Cys Glu Arg Leu Asp Ile Ala Asp Pro Ala Ala

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1085	1090	1095
Val Ala Ala Phe Phe Ala Ala 1100	Leu Arg Arg Asp 1105	Gly Val Pro Leu 1110
Lys Gly Ile Val His Ala Ala 1115	Gly Ile Val Gly Tyr 1120	Lys Pro Ile 1125
Met Gln Val Glu Arg Asp Glu 1130	Leu Asp Ala Val 1135	Leu Gln Pro Lys 1140
Val Ala Gly Ala Trp Leu Leu 1145	His Gln Gln Ser 1150	Glu His Phe Pro 1155
Leu Asp Phe Phe Leu Leu Phe 1160	Ser Ser Ile Ala 1165	Ser Ala Trp Gly 1170
Ser Arg Glu Gln Ala His Tyr 1175	Ser Ala Ala Asn 1180	Arg Phe Leu Asp 1185
Ala Leu Ala His His Arg Arg 1190	Gly Gln Gly Leu 1195	Pro Ala Leu Ser 1200
Val Asn Trp Gly Pro Trp Ala 1205	Glu Gly Gly Met 1210	Thr Phe Pro Glu 1215
Ala Glu Ala Leu Leu Arg Arg 1220	Val Gly Ile Arg 1225	Ser Leu Ala Ala 1230
Asp Arg Ala Leu Asp Val Leu 1235	Asn Arg Leu Pro 1240	Ala Val Pro Gln 1245
Val Ala Val Val Asp Ile Asp 1250	Leu Ala Leu Phe 1255	Gln Gly Ser Tyr 1260
Glu Ala Arg Gly Pro Lys Pro 1265	Phe Leu Asp His 1270	Val Arg Val Ala 1275
Lys Ser Ala Pro Ser Ala Pro 1280	Ala Met Pro Ala 1285	Leu Ser Asp Ala 1290
Ser Pro Arg Glu Arg Lys Arg 1295	Leu Leu Ala Asp 1300	Ser Ile Asp Arg 1305
Ala Val Ala Gln Val Leu Gly 1310	Tyr Asp Ala Gly 1315	Thr Leu Asp Arg 1320
Asp Leu Gly Phe Phe Glu Met 1325	Gly Met Asp Ser 1330	Leu Met Ala Leu 1335
Asp Val Arg Thr His Leu Glu 1340	Asn Ala Leu Gly 1345	Ile Pro Leu Ser 1350
Val Ala Leu Leu Phe Asp His 1355	Pro Thr Val Asn 1360	Ala Leu Ala Asp 1365
Phe Leu Ala Glu Gln Ala Ser 1370	Gly Thr Ala Gln 1375	Ala Gln Thr Val 1380
Pro Pro Gln Gln Gln Pro Arg 1385	Pro Ile Ala Pro 1390	Ala Ile Glu Ala 1395
Arg Asp Ala Gly Thr Pro Glu 1400	Pro Ile Ala Ile 1405	Val Gly Met Ser 1410
Cys Arg Phe Pro Gly Ala Ala 1415	His Asp Leu Asp 1420	Ala Tyr Trp Asn 1425
Leu Leu Asn Asp Gly Val Asp 1430	Ala Ile Ser Glu 1435	Val Pro Arg Glu 1440
Arg Trp Asp Val Asp Ala Tyr 1445	Tyr Asp Pro Asp 1450	Pro Glu Ala Pro 1455
Gly Arg Met Tyr Ser Arg Phe 1460	Gly Gly Phe Leu 1465	Asp Asp Val Asp 1470
Gln Phe Asp Pro Ala Phe Phe 1475	Arg Ile Thr Pro 1480	Arg Glu Ala Ala 1485

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Ala Met Asp Pro Gln Gln Arg Leu Leu Leu Glu Val Ser His Glu  
1490 1495 1500

Ala Leu Glu His Ala Gly Ile Pro Val Asp Ser Leu Lys Gly Ser  
1505 1510 1515

Arg Thr Gly Val Phe Val Gly Ile Thr Thr Asn Asp Tyr Ala Asn  
1520 1525 1530

Leu Gln Leu Arg Asn Gly Gly Gly Ser Gly Ile Asp Gly Tyr Phe  
1535 1540 1545

Phe Thr Gly Asn Pro Leu Asn Thr Ala Ala Gly Arg Ile Ser Tyr  
1550 1555 1560

Gly Leu Gly Val Gln Gly Pro Ser Met Ala Ile Asp Thr Ala Cys  
1565 1570 1575

Ser Ser Ser Leu Thr Ala Ile His Thr Ala Ser Gln Asn Leu Arg  
1580 1585 1590

Ser Gly Glu Cys Asp Leu Ala Ile Ala Gly Gly Val Asn Leu Ile  
1595 1600 1605

Leu Ser Pro Asp Asn Ser Ile Ala Val Ser Arg Thr Arg Ala Leu  
1610 1615 1620

Ala Pro Asp Gly Arg Cys Lys Thr Phe Asp Ala Ala Ala Asp Gly  
1625 1630 1635

Phe Val Arg Ser Glu Gly Cys Gly Ala Leu Val Leu Lys Arg Leu  
1640 1645 1650

Ser Asp Ala Leu Ala Ala Gly Asp Arg Val Leu Ala Val Leu Arg  
1655 1660 1665

Gly Ser Ala Val Asn His Asp Gly Ala Ser Ser Gly Phe Thr Ala  
1670 1675 1680

Pro Asn Gly Arg Ala Gln Glu Ala Val Ile Arg Gln Ala Leu Gly  
1685 1690 1695

Gly Leu Pro Ala Ala Ser Ile Asp Tyr Val Glu Ala His Gly Thr  
1700 1705 1710

Gly Thr Pro Leu Gly Asp Pro Val Glu Leu Gln Ala Leu Ala Thr  
1715 1720 1725

Val Phe Gly Ala Gly Arg Asp Ala Gly Arg Arg Leu Arg Val Gly  
1730 1735 1740

Ser Val Lys Thr Asn Ile Gly His Thr Glu Ser Ala Ala Gly Ile  
1745 1750 1755

Ala Gly Val Ile Lys Val Val Leu Ser Leu Asn His Asp Arg Leu  
1760 1765 1770

Pro Ala His Leu His Phe Arg Gln Pro Ser Pro Leu Val Gln Trp  
1775 1780 1785

Asp Ala Met Pro Val Glu Ile Cys Ala Glu Ala Ser Ala Trp Pro  
1790 1795 1800

Arg Gly Glu Arg Pro Arg Arg Ala Gly Val Ser Ala Phe Gly Ala  
1805 1810 1815

Ser Gly Thr Asn Ala His Leu Val Leu Glu Glu Ala Pro Ala Pro  
1820 1825 1830

Ala Arg Gln Ala Thr Pro Ser Arg His Lys Val His Pro Leu Val  
1835 1840 1845

Leu Ser Ala Lys Thr Pro Ala Ala Leu Arg Glu Leu Ala Gly Arg  
1850 1855 1860

Tyr Gln Arg Arg Leu Glu Ala Glu Pro Gly Leu Asp Ile Ala Ala  
1865 1870 1875

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Val	Ala	Phe	Ser	Ala	Ala	Thr	Gly	Arg	Ser	His	Phe	Ala	His	Arg
1880						1885					1890			
Leu	Ala	Trp	Pro	Val	Thr	Ser	Leu	Asp	Asp	Ala	Ile	Asp	Lys	Leu
1895						1900					1905			
Arg	Ala	Phe	His	Ala	Lys	Glu	Pro	Ala	Gly	Ala	Ala	Gln	Pro	Ala
1910						1915					1920			
Pro	Arg	Val	Lys	Met	Ala	Phe	Leu	Phe	Thr	Gly	Gln	Gly	Ser	Gln
1925						1930					1935			
Tyr	Ala	Gly	Met	Gly	Arg	Arg	Leu	Tyr	Asp	Ala	Tyr	Pro	Val	Phe
1940						1945					1950			
Arg	Asp	Ala	Ile	Asp	Arg	Cys	Arg	Ala	Val	Ala	Asp	Pro	Leu	Leu
1955						1960					1965			
Asp	Lys	Pro	Leu	Leu	Glu	Val	Leu	Ser	Ala	Gln	Gly	Glu	Asp	Ile
1970						1975					1980			
His	Gln	Thr	Gly	Tyr	Ser	Gln	Pro	Ala	Leu	Phe	Ser	Leu	Gln	Tyr
1985						1990					1995			
Ala	Leu	Thr	Thr	Leu	Leu	Ala	Ser	Phe	Gly	Val	Val	Pro	Asp	Ala
2000						2005					2010			
Val	Met	Gly	His	Ser	Val	Gly	Glu	Tyr	Ala	Ala	Ala	Cys	Ala	Ala
2015						2020					2025			
Gly	Val	Phe	Ser	Pro	Glu	Asp	Gly	Leu	Arg	Leu	Ile	Ala	Glu	Arg
2030						2035					2040			
Gly	Arg	Leu	Met	Gln	Ala	Leu	Pro	Arg	Asp	Gly	Glu	Met	Ala	Ala
2045						2050					2055			
Ile	Phe	Thr	Asp	Leu	Ala	Thr	Val	Glu	Arg	Ala	Ile	Asp	Ala	Trp
2060						2065					2070			
Pro	His	Glu	Val	Ala	Val	Ala	Ala	Val	Asn	Gly	Pro	Ala	Ser	Ile
2075						2080					2085			
Val	Ile	Ser	Gly	Lys	Arg	Glu	Arg	Ile	Ala	Met	Leu	Val	Asp	Ala
2090						2095					2100			
Phe	Ala	Ala	Arg	Asp	Ile	Arg	Ser	Val	Pro	Leu	Asn	Thr	Ser	His
2105						2110					2115			
Ala	Phe	His	Ser	Pro	Leu	Leu	Glu	Pro	Met	Leu	Asp	Ser	Phe	Gln
2120						2125					2130			
Leu	Ala	Ala	Lys	Thr	Val	Pro	Val	Ala	Arg	Pro	Ala	Ile	Pro	Phe
2135						2140					2145			
Tyr	Ser	Asn	Leu	Thr	Gly	Ala	Val	Met	Asp	Glu	Ala	Pro	Thr	Asp
2150						2155					2160			
Thr	Tyr	Trp	Arg	Arg	His	Cys	Arg	Glu	Pro	Val	Gln	Phe	Ala	Ser
2165						2170					2175			
Ser	Val	Glu	Arg	Leu	Ala	Glu	Ala	Gly	Phe	Asn	Val	Leu	Val	Glu
2180						2185					2190			
Ile	Gly	Pro	Lys	Pro	Val	Leu	Val	Asn	Leu	Ala	Arg	Ala	Cys	Cys
2195						2200					2205			
Ala	Pro	Asp	Ala	Gly	Ile	Gln	Phe	Leu	Ala	Leu	Gln	Arg	Pro	Gln
2210						2215					2220			
Val	Glu	Gln	Gln	Ala	Leu	Ile	Glu	Thr	Leu	Ser	Ser	Leu	Tyr	Ala
2225						2230					2235			
Arg	Gly	Val	Asp	Val	Asp	Trp	Ala	Pro	Thr	Glu	Thr	Pro	Ala	Pro
2240						2245					2250			
Ala	Arg	Ile	Ala	Leu	Pro	Ser	Tyr	Pro	Phe	Gln	Arg	Ser	Arg	Thr
2255						2260					2265			
Trp	Phe	Gln	Lys	Ala	Asp	Thr	Ser	Met	Thr	Gln	Thr	Ser	Ala	Ser



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2270	2275	2280
Pro Ile Ala Ala Ala Pro Thr His Asn Arg Ser Gly Glu Ile Leu 2285	2290	2295
Glu Trp Leu Arg Gly Lys Ile Gly Glu Leu Ile Gln Ala Asp Pro 2300	2305	2310
Ala Thr Ile Asn Ile Glu Leu Pro Phe Leu Glu Met Gly Ala Asp 2315	2320	2325
Ser Ile Val Leu Ile Glu Ala Ile Arg His Ile Glu Ala Glu Tyr 2330	2335	2340
Gly Val Lys Leu Ala Met Arg Arg Phe Phe Glu Asp Leu Ala Thr 2345	2350	2355
Val Gln Ala Leu Ala Glu Tyr Val Ala Asp Asn Leu Pro Ala Ala 2360	2365	2370
Ala Ala Pro Ser Gly Ala Glu Ala Val Ala Val Ala Val Ala Ala 2375	2380	2385
Ala Glu Pro Ser Thr Pro Ala Val Ala Val Thr Pro Ser Ala Ala 2390	2395	2400
Gly Leu Ala Pro Leu Ala Ala Ala Pro Ala Glu Trp Val Ala Ala 2405	2410	2415
Glu Gly Gly Ser Thr Val Glu Arg Val Leu Arg Glu Gln Asn Gln 2420	2425	2430
Leu Leu Ser His Val Met Ser Gln Gln Met Glu Leu Leu Arg Thr 2435	2440	2445
Ser Leu Thr Gly Gln Pro Gly Val Arg Pro Ala Thr Ala Ala Val 2450	2455	2460
Gln Ala Val Ala Ser Thr Ala Ser Val Ala Pro Lys Ala Ala Ser 2465	2470	2475
Ala Ala Pro Ala Ala Ala Pro Ala Ala Lys Pro Ala Pro Ala Ala 2480	2485	2490
Ala Ala Ala Pro Ala Ala Asp Asn Pro Pro Pro Lys Pro Met Met 2495	2500	2505
Pro Trp Gly Ser Pro Val Gln Gln Arg Ala Arg Gly Leu Ser Ala 2510	2515	2520
Ala Gln Gln Glu His Leu Glu Ala Leu Ile Val Arg Tyr Thr Thr 2525	2530	2535
Arg Thr Arg Lys Ser Lys Asp Ser Val Gln Ala Ser Arg Pro Val 2540	2545	2550
Leu Ala Asp Ser Arg Ala Thr Val Gly Phe Arg Phe Ser Thr Lys 2555	2560	2565
Glu Met Leu Tyr Pro Ile Val Gly Asp Arg Ala Ala Gly Ser Arg 2570	2575	2580
Leu Trp Asp Ile Asp Gly Asn Glu Tyr Ile Asp Phe Thr Met Gly 2585	2590	2595
Phe Gly Val His Leu Phe Gly His Thr Pro Asp Phe Ile Gln Gln 2600	2605	2610
Gln Val Thr Arg Glu Trp Gln Arg Pro Leu Glu Leu Gly Ala Arg 2615	2620	2625
Ser Ser Leu Val Gly Glu Val Ala Ala Arg Phe Ala Arg Val Thr 2630	2635	2640
Gly Leu Asp Arg Val Ala Phe Ser Asn Thr Gly Thr Glu Ala Val 2645	2650	2655
Met Thr Ala Met Arg Leu Ala Arg Ala Val Thr Gly Arg Asp Lys 2660	2665	2670

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Ile Val Met Phe Thr His Ser Tyr His Gly His Ala Asp Gly Thr 2675 2680 2685
Leu Ala Ala Ala Asn Ala Glu Gly Val Thr Glu Thr Ile Ala Pro 2690 2695 2700
Gly Val Pro Phe Gly Ser Val Glu Asn Met Ile Leu Leu Asp Tyr 2705 2710 2715
Gly Ser Asp Ala Ala Leu Glu Ala Ile Arg Gly Met Ala Ser Thr 2720 2725 2730
Leu Ala Ala Val Met Val Glu Pro Val Gln Ser Arg Asn Pro Ser 2735 2740 2745
Leu Gln Pro Val Ala Phe Leu Lys Glu Leu Arg Arg Ile Thr Glu 2750 2755 2760
Glu Ala Gly Val Ala Leu Ile Phe Asp Glu Met Ile Thr Gly Phe 2765 2770 2775
Arg Val His Pro Gly Gly Ser Gln Ala Met Phe Gly Ile Arg Ala 2780 2785 2790
Asp Leu Ala Thr Tyr Gly Lys Ile Ile Gly Gly Gly Leu Pro Leu 2795 2800 2805
Gly Val Ile Ala Gly Thr Ser Arg Phe Met Asp Ala Ile Asp Gly 2810 2815 2820
Gly Met Trp Thr Tyr Gly Asp His Ser Phe Pro Ala Ala Asp Arg 2825 2830 2835
Thr Ala Phe Gly Gly Thr Phe Cys Gln Tyr Pro Leu Ala Met Ala 2840 2845 2850
Ala Ala Leu Ala Val Leu Glu Lys Ile Glu Gln Glu Gly Pro Ala 2855 2860 2865
Leu Gln Ala Ala Leu Asn Glu Arg Thr Ala Gln Ile Ala Gly Thr 2870 2875 2880
Leu Asn Ala Phe Phe Ala Glu Ala Glu Ala Pro Ile Lys Val Thr 2885 2890 2895
Trp Phe Gly Ser Met Phe Arg Phe Glu Phe Thr Glu Asn Leu Asp 2900 2905 2910
Leu Phe Phe Tyr His Met Leu Glu Lys Gly Ile Tyr Ile Trp Glu 2915 2920 2925
Trp Arg Thr Cys Phe Leu Ser Thr Ala His Thr Asp Ala Asp Ile 2930 2935 2940
Asp Arg Phe Ile Arg Ala Val Lys Asp Ser Val Ala Asp Leu Arg 2945 2950 2955
Arg Gly Gly Phe Ile Arg Pro His Ser Lys His Gly Thr Val Ala 2960 2965 2970
Ala Leu Ser Glu Ala Gln Arg Gln Leu Trp Val Leu Ser Glu Ile 2975 2980 2985
Asp Pro Glu Gly Ser Leu Ala Tyr Asn Val Asn Thr Thr Leu Glu 2990 2995 3000
Leu Asn Gly Arg Leu Asp Glu Ala Ala Met Arg Ala Ala Val Gln 3005 3010 3015
Ser Leu Val Asp Arg His Glu Ala Leu Arg Thr Thr Val Met Ala 3020 3025 3030
Asp Gly Ser Gly Gln Ile Val His Pro Ser Leu Thr Leu Glu Ile 3035 3040 3045
Pro Leu Ile Asp Thr Asp Pro Asn Ala Trp Arg Glu Gln Glu Ser 3050 3055 3060

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Arg	Gln	Pro	Phe	Asp	Leu	Val	Asn	Gly	Pro	Leu	Phe	Arg	Ala	Ala
3065						3070					3075			
Leu	Val	Arg	Leu	Gly	Ser	Glu	Arg	His	Leu	Leu	Val	Met	Thr	Ala
3080						3085					3090			
His	His	Ile	Ile	Cys	Asp	Gly	Ser	Thr	Phe	Gly	Val	Leu	Leu	Glu
3095						3100					3105			
Asp	Leu	Ala	Arg	Ala	Tyr	Ala	Gly	Ala	Ala	Pro	Ala	Asp	Ala	Pro
3110						3115					3120			
Leu	Gln	Phe	Arg	Ala	Tyr	Leu	Lys	Gln	Leu	Asp	Gly	Gln	Arg	His
3125						3130					3135			
Ser	Pro	Glu	Thr	Lys	Ala	Asn	Arg	Glu	Tyr	Trp	Leu	Ala	Gln	Cys
3140						3145					3150			
Ala	Arg	Gln	Ala	Ala	Pro	Leu	Asn	Leu	Pro	Val	Asp	Tyr	Pro	Arg
3155						3160					3165			
Pro	Ala	Val	Lys	Thr	Phe	His	Gly	Glu	Arg	Val	Ser	Leu	His	Leu
3170						3175					3180			
Asp	Ala	Ala	Thr	Ala	Ala	Thr	Leu	Arg	Thr	Ala	Ala	Arg	Gln	Asn
3185						3190					3195			
Gly	Cys	Thr	Leu	Tyr	Met	Val	Leu	Leu	Ala	Gly	Phe	Asn	Leu	Phe
3200						3205					3210			
Leu	His	Arg	Val	Ala	Gly	Gln	Gln	Glu	Ile	Val	Thr	Gly	Ile	Pro
3215						3220					3225			
Val	Thr	Gly	Arg	Ser	Val	Ala	Gly	Ser	Asp	Arg	Leu	Ala	Gly	Tyr
3230						3235					3240			
Cys	Thr	His	Leu	Leu	Pro	Leu	His	Ser	Thr	Leu	Pro	Glu	Gln	Ala
3245						3250					3255			
Thr	Val	Ala	Ser	Phe	Leu	Ala	Gly	Thr	Arg	Gln	Asn	Leu	Leu	Asp
3260						3265					3270			
Ala	Leu	Glu	His	Gln	Asp	Tyr	Pro	Phe	Ala	Glu	Leu	Val	Arg	Glu
3275						3280					3285			
Ile	Gly	Ala	Gln	Arg	Asp	Leu	Asn	Ala	Ala	Pro	Leu	Val	Ser	Ala
3290						3295					3300			
Val	Phe	Asn	Leu	Glu	Pro	Val	Ser	Ala	Leu	Pro	Glu	Leu	Pro	Gly
3305						3310					3315			
Leu	Thr	Val	Gly	Leu	Val	Ala	Pro	Leu	Ile	Arg	His	Thr	Ala	Phe
3320						3325					3330			
Asp	Leu	Asn	Val	Asn	Val	Leu	Asp	Ala	Gly	Gln	Ala	Leu	Leu	Ile
3335						3340					3345			
Asp	Cys	Asp	Tyr	Asn	Thr	Asp	Leu	Phe	Asp	Ala	Ser	Thr	Val	Gln
3350						3355					3360			
Arg	Phe	Leu	Asp	Ile	Tyr	Arg	Thr	Leu	Leu	Thr	His	Leu	Ala	Asp
3365						3370					3375			
Asp	Ala	Ser	Ala	Ala	Val	Ala	Arg	Leu	Pro	Leu	Ser	Ser	Asp	Ala
3380						3385					3390			
Glu	Arg	Asn	Leu	Leu	Thr	Val	Glu	Trp	Asn	Arg	Thr	Asp	Thr	Asp
3395						3400					3405			
Phe	Gly	Glu	Asp	Ala	Ala	Gln	Pro	Leu	His	Arg	Leu	Phe	Glu	Gln
3410						3415					3420			
Gln	Val	Glu	Arg	Thr	Pro	Asp	Ala	Val	Ala	Ile	Val	Phe	Asp	Asp
3425						3430					3435			
Thr	Ala	Leu	Thr	Tyr	Ala	Glu	Leu	Asn	Leu	Arg	Ala	Asn	Arg	Leu
3440						3445					3450			
Ala	His	His	Leu	Val	Ala	Leu	Gly	Val	Gly	Pro	Asp	Ser	Leu	Val

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3455	3460	3465
Gly Val Ala Met Glu Arg Ser Leu Asp Met Ser Val Ala Leu Leu 3470 3475 3480		
Ala Ile Leu Lys Ala Gly Gly Ala Tyr Val Pro Val Asp Pro Asp 3485 3490 3495		
Tyr Pro Ala Glu Arg Val Arg Phe Met Ile Asp His Ala Gln Leu 3500 3505 3510		
Arg Trp Leu Leu Thr Gln Gln His Leu His Asp Ala Leu Pro Asp 3515 3520 3525		
Thr Asp Ala His Val Ile Val Val Asp Arg Asp Ser Leu Asp Leu 3530 3535 3540		
Asp Ala Ala Ala Thr Ser Asn Pro Ala Pro Ala Leu Asn Gly Asp 3545 3550 3555		
Asn Leu Ala Tyr Met Ile Tyr Thr Ser Gly Ser Thr Gly Arg Pro 3560 3565 3570		
Lys Gly Ala Leu Asn Thr His Arg Ala Ile Thr Asn Arg Ile Leu 3575 3580 3585		
Trp Met Gln His Ala Tyr Ala Leu Gly Ala Asp Asp Ala Val Leu 3590 3595 3600		
Gln Lys Thr Pro Phe Ser Phe Asp Val Ser Val Trp Glu Leu Phe 3605 3610 3615		
Trp Pro Leu Val Thr Gly Ala Arg Leu Val Phe Ala Arg Pro Gly 3620 3625 3630		
Gly Gln Arg Glu Thr Asp Tyr Leu Val Glu Leu Ile Glu Arg Glu 3635 3640 3645		
Arg Ile Thr Thr Ile His Phe Val Pro Ser Met Leu Arg Ala Phe 3650 3655 3660		
Leu Asp His Pro Asp Leu Asp Ala His Cys Ala Ser Leu Arg Arg 3665 3670 3675		
Val Val Cys Ser Gly Glu Ala Leu Pro His Asp Leu Gln Gln Arg 3680 3685 3690		
Cys Leu Glu Arg Leu Asp Val Glu Leu Tyr Asn Leu Tyr Gly Pro 3695 3700 3705		
Thr Glu Ala Ala Val Asp Val Thr Ala Trp Glu Cys Arg Arg Asp 3710 3715 3720		
Asp Pro His Arg Ile Val Pro Ile Gly Arg Pro Ile Ala Asn Thr 3725 3730 3735		
Arg Leu Tyr Ile Val Asp Ala Gln Met Gln Pro Thr Pro Ile Gly 3740 3745 3750		
Val Ala Gly Glu Leu Leu Ile Gly Gly Thr Pro Val Gly Arg Gly 3755 3760 3765		
Tyr His Gly Glu Pro Glu Leu Ser Ala Glu Lys Phe Ile Ala Asp 3770 3775 3780		
Pro Phe Ser Ala Asp Pro Leu Ala Arg Leu Tyr Arg Thr Gly Asp 3785 3790 3795		
Leu Ala Arg Tyr Arg Pro Asp Gly Asn Ile Glu Phe Leu Gly Arg 3800 3805 3810		
Ile Asp His Gln Ile Lys Leu Arg Gly Leu Arg Ile Glu Pro Gly 3815 3820 3825		
Glu Ile Glu Ala Ala Leu Thr Ser His Pro Leu Val Asp Ala Ala 3830 3835 3840		
Val Val Ala Leu Arg Gly Val Asp Asp Gly Ala Arg Leu Val Gly 3845 3850 3855		

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Trp	Leu	Cys	Ser	Ser	His	Pro	Glu	Ala	Glu	Leu	Ile	Glu	Ala	Val
3860						3865					3870			
Arg	Gly	His	Leu	Arg	Gln	Arg	Leu	Pro	Asp	Tyr	Met	Val	Pro	Ser
3875						3880					3885			
Ala	Phe	Val	Val	Val	Ser	Ala	Phe	Glu	His	Leu	Pro	Asn	Gly	Lys
3890						3895					3900			
Leu	Asp	Arg	Thr	Arg	Leu	Pro	Glu	Pro	Gly	Asp	Gly	Leu	Asp	His
3905						3910					3915			
Val	Ala	Pro	Val	Asn	Ala	Leu	Glu	Ala	Gln	Leu	Ala	Ala	Ile	Trp
3920						3925					3930			
Gln	Glu	Val	Leu	Gly	Gln	Ala	Arg	Ile	Ser	Thr	Thr	Gly	Asn	Phe
3935						3940					3945			
Phe	Asp	Leu	Gly	Gly	Asn	Ser	Leu	Leu	Ala	Thr	Lys	Val	Val	Ala
3950						3955					3960			
Arg	Ile	Arg	Arg	Asp	Leu	His	Val	Lys	Leu	Glu	Ile	Arg	Ser	Leu
3965						3970					3975			
Phe	Ala	Leu	Pro	Thr	Ile	Ser	Ser	Leu	Ala	Lys	Arg	Ile	Ala	Asp
3980						3985					3990			
Thr	Gln	Pro	Ile	Asp	Tyr	Ala	Pro	Val	Thr	Pro	Leu	Pro	Ala	Gln
3995						4000					4005			
Ala	Ser	Tyr	Ala	Leu	Ser	Pro	Ala	Gln	Thr	Arg	Leu	Trp	Val	Gln
4010						4015					4020			
Asp	Arg	Leu	His	Ala	Ala	Gln	Ala	Glu	Gly	Pro	Leu	Pro	Thr	Ser
4025						4030					4035			
Leu	Leu	Phe	Glu	Gly	Val	Leu	Asp	Val	Asp	Ala	Leu	Val	Arg	Ala
4040						4045					4050			
Phe	Arg	Ala	Leu	Ser	Glu	Arg	His	Glu	Ile	Leu	Arg	Thr	Arg	Phe
4055						4060					4065			
Val	Leu	Glu	Gly	Asn	Gln	Pro	Val	Gln	His	Val	Leu	Pro	Pro	Gly
4070						4075					4080			
Glu	Ala	Ala	Phe	Pro	Val	Glu	Ile	Val	Asp	Leu	Gln	Asp	Ala	Glu
4085						4090					4095			
Asp	Arg	Asp	Ala	Gln	Ala	Ala	Ala	Ile	Gln	Ala	Ser	Glu	Arg	Leu
4100						4105					4110			
Val	Pro	Met	Asp	Leu	Ala	Thr	Gly	Pro	Leu	Phe	Arg	Val	Lys	Leu
4115						4120					4125			
Leu	Arg	Leu	Ser	Glu	Val	Arg	His	Val	Cys	Leu	Cys	Thr	Met	His
4130						4135					4140			
His	Ile	Val	Ser	Asp	Gly	Trp	Ser	Thr	Glu	Val	Leu	Leu	Asp	Asp
4145						4150					4155			
Leu	Ser	Ala	Leu	Tyr	Asp	Ala	Phe	Val	Gln	Arg	Arg	Asp	Asp	Pro
4160						4165					4170			
Leu	Pro	Ala	Leu	Pro	Ile	Gln	Tyr	Lys	Asp	Tyr	Ala	Gly	Trp	Leu
4175						4180					4185			
Asn	Arg	Leu	Leu	Ala	Gly	Pro	Asp	Gly	Ala	Arg	Met	Lys	Asp	Tyr
4190						4195					4200			
Trp	Leu	Thr	Lys	Leu	Gly	Gly	Gly	Leu	Arg	Ala	Leu	Glu	Leu	Pro
4205						4210					4215			
Gly	Asp	Val	Glu	Gln	Pro	Ala	Ala	Pro	Ser	Trp	Lys	Ser	Trp	Arg
4220						4225					4230			
Phe	Asp	Leu	Pro	Ala	Ala	Glu	Thr	Ala	Ala	Leu	Glu	Ser	Leu	Gly
4235						4240					4245			

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Lys Arg His Gly Ala Thr Leu Phe Ile Ala Leu Leu Ser Ala Ile  
 4250 4255 4260  
 Lys Ala Leu Phe Tyr Arg Arg Ser Gly Gln Glu Asp Ile Val Val  
 4265 4270 4275  
 Gly Thr Pro Val Ala Gly Arg Glu Leu Pro Glu Leu Glu Ser Gln  
 4280 4285 4290  
 Val Gly Pro Tyr Leu Asn Val Leu Ala Leu Arg Asp Arg Val Ala  
 4295 4300 4305  
 Gly Asp Asp Arg Phe Asp Thr Leu Leu Thr Arg Val Arg Asp Thr  
 4310 4315 4320  
 Thr Leu Glu Ala Phe Ser His Pro Leu Tyr Pro Leu Asp Arg Leu  
 4325 4330 4335  
 Leu Asp Glu Leu His Ile Lys Arg Val Ala Gly Arg Asn Pro Leu  
 4340 4345 4350  
 Phe Asp Ile Gly Leu Thr Leu Gln Asn Gln Arg His Gly Pro Val  
 4355 4360 4365  
 Asp Arg Tyr Ala Gly Gln Val His Ile Ala Glu Leu Pro Asp His  
 4370 4375 4380  
 Asp Pro Gln Arg Ala Asp Thr Glu Ala Ala Thr Asp Phe Trp Phe  
 4385 4390 4395  
 Leu Ala Glu Pro His Ala Glu Gly Leu Ala Ile Arg Val Val Tyr  
 4400 4405 4410  
 His Ala Gly Arg Phe Ser Glu Ala Leu Val Gln Gly Leu Ala Asn  
 4415 4420 4425  
 Glu Leu Thr Ser Val Ile Gly Glu Val Leu Ala Asn Pro Gly Val  
 4430 4435 4440  
 Arg Ile Arg Asn Leu Thr Leu Gly Gln Arg Ala Leu His Ala Glu  
 4445 4450 4455  
 Ala Arg Gln Pro Thr Val Glu Leu Ser Ala Phe  
 4460 4465

&lt;210&gt; SEQ ID NO 16

&lt;211&gt; LENGTH: 1107

&lt;212&gt; TYPE: PRT

&lt;213&gt; ORGANISM: Burkholderia contaminans

&lt;220&gt; FEATURE:

&lt;221&gt; NAME/KEY: MISC\_FEATURE

&lt;222&gt; LOCATION: (1)..(1107)

<223> OTHER INFORMATION: putative short chain dehydrogenase/reductase  
SDR

&lt;400&gt; SEQUENCE: 16

Met Lys Phe Gly Leu Met Phe Phe Ala Ser Ser Glu Glu Ala Leu Ser  
 1 5 10 15  
 Gly Asn Lys Tyr Gln Leu Val Met Glu Ser Ala Arg Phe Ala Asp Ala  
 20 25 30  
 Asn Gly Phe Ser Ser Val Trp Val Pro Glu Arg His Phe Thr Glu Phe  
 35 40 45  
 Gly Ser Leu Tyr Pro Asn Pro Ala Val Leu His Ala Ala Leu Ala Ala  
 50 55 60  
 Ala Thr Gln Arg Val Lys Leu Val Ala Gly Ser Val Val Ala Ala Leu  
 65 70 75 80  
 His Asn Pro Ile Arg Ile Ala Glu Glu Trp Ser Met Val Asp Asn Leu  
 85 90 95  
 Ser Asn Gly Arg Val Gly Val Ser Phe Ala Ser Gly Trp Asn Pro Asp  
 100 105 110

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Asp Phe Val Phe Ala Pro Asp Lys Tyr Ala Thr Arg Gln Asp Asp Met  
 115 120 125  
 Leu Thr Thr Met Arg Ala Val Gln His Leu Trp Arg Gly Gly Thr Leu  
 130 135 140  
 Asp Ala Thr Asn Gly Val Gly Lys Pro Val Arg Leu Arg Val Tyr Pro  
 145 150 155 160  
 Thr Pro Val Gln Pro Glu Leu Pro Val Trp Val Thr Ala Ala Ser Asn  
 165 170 175  
 Pro Gln Thr Phe Val Arg Ala Gly Glu Ala Gly Ala Asn Leu Leu Thr  
 180 185 190  
 His Val Leu Asp Gln Asp Arg Asp Gln Leu Ala His Lys Ile Ala Leu  
 195 200 205  
 Tyr Arg Glu Ala Arg Ala Lys His Gly Phe Asp Pro Ala Ala Gly Thr  
 210 215 220  
 Val Ser Val Met Leu His Thr Phe Val Gly Asp Asp Ala Ala Gln Ala  
 225 230 235 240  
 Arg Glu Gln Ala Arg Val Pro Phe Cys Asn Tyr Ile Arg Ser Asn Ile  
 245 250 255  
 Gly Leu Leu Asn Gly Leu Ala Gln Ser Arg Gly Gln Ser Val Asp Val  
 260 265 270  
 Arg Ala Met Gly Ala Arg Glu Leu Asp Glu Phe Val Glu Phe Leu Tyr  
 275 280 285  
 Glu Arg Phe Ala Gln Ser Arg Gly Leu Ile Gly Thr Pro Glu Thr Cys  
 290 295 300  
 Val Glu Leu Val Arg Asp Leu Glu Ser Ile Gly Val Asp Glu Val Ala  
 305 310 315 320  
 Cys Leu Leu Asp Phe Gly Pro Pro Val Glu Arg Ile Leu Gly Asn Leu  
 325 330 335  
 Pro Gln Leu Arg Arg Leu Arg Glu Met Cys Ala Pro Arg Arg Ser Ala  
 340 345 350  
 Ala Pro Thr Arg Phe Asp Ala Ala Glu Val Gln Ala Arg Cys Thr Glu  
 355 360 365  
 Thr Thr Ser Gly Ala Asp Phe Asn Gly Glu Ile Arg Gln His Gly Val  
 370 375 380  
 Gln Ile Asp Gly Val Phe Asp Ala Ile Arg Gln Ile Trp Arg Thr Thr  
 385 390 395 400  
 Gly Glu Ala Leu Gly Lys Ile Ser Leu Pro Ala Asp Ala Leu Ala Ser  
 405 410 415  
 Ser Pro Tyr Gln Val His Pro Ala Phe Leu Asp Ala Cys Ser Arg Val  
 420 425 430  
 Leu Ala Ala Ala Ile Asp Pro Asp Ala Leu Glu Ser Gly Asp Leu Tyr  
 435 440 445  
 Leu Pro Ser Ser Ile Gly Ala Val Arg Val His Gln Pro Pro Ala Ser  
 450 455 460  
 Thr Glu Ala Trp Ser His Ala Thr Leu Arg Thr Pro Ile Gly Gln Gly  
 465 470 475 480  
 Ala Leu Glu Gly Asp Ile Arg Val His Asp Leu Ala Gly Arg Leu Leu  
 485 490 495  
 Ile Glu Ile Asp Ala Leu Arg Leu Gln Gln Val Arg Ala Ala Arg Ala  
 500 505 510  
 Val Glu Arg His Asp Phe Ala Ala Leu Leu Tyr Gln Arg Val Trp Arg  
 515 520 525  
 Pro Ser Asn Val Asp Ala Ala Thr Gly Gly Ser Ala His Gly Glu Trp

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530					535					540					
Leu	Ile	Leu	Ala	Asp	Arg	Gly	Gly	Val	Gly	Ala	Gln	Leu	Ser	Ala	Leu
545					550					555					560
Leu	Glu	Ala	His	Gly	Asp	Thr	Cys	Thr	Leu	Arg	Phe	Ala	Asp	Ala	Thr
				565					570						575
Pro	Glu	Leu	Pro	Ala	Ala	Asp	Arg	Pro	Leu	Lys	Gly	Val	Ile	His	Leu
			580					585						590	
Trp	Ser	Leu	Asp	Leu	Ala	Pro	Ser	Asp	Ile	Ala	Ala	Arg	Arg	Arg	Ala
		595					600						605		
Ser	Ala	Ser	Val	Leu	His	Leu	Val	Arg	Ala	Leu	Ala	Ser	Arg	Ala	Pro
610						615							620		
Ser	Ala	Arg	Gln	Ala	Arg	Leu	Trp	Leu	Val	Thr	Ser	Gly	Ala	Met	Asn
625					630					635					640
Val	Leu	Asp	Gly	Glu	Ser	Ile	Ala	Val	Ala	Gln	Ala	Pro	Leu	Trp	Gly
				645					650						655
Leu	Gly	Arg	Ala	Ile	Ala	Val	Glu	His	Ala	Ala	Leu	Trp	Gly	Gly	Leu
			660						665						670
Val	Asp	Leu	Asp	Pro	Glu	Gln	Pro	Ser	Ala	Ala	Asp	Ile	Met	Gln	Ala
		675					680							685	
Val	Gln	Ala	Gly	Gly	Arg	Glu	Asp	Met	Ile	Ala	Phe	Arg	Arg	Asp	Gln
690						695					700				
Arg	Tyr	Val	Ala	Arg	Ile	Ala	Arg	Asp	Asn	Arg	Glu	Tyr	Val	Ser	His
705					710					715					720
Arg	Pro	Ile	Arg	Phe	His	Gly	Asp	Ala	Thr	Tyr	Leu	Val	Thr	Gly	Gly
				725					730						735
Leu	Gly	Gly	Leu	Gly	Leu	Arg	Leu	Ala	Ser	Trp	Leu	Ala	Asp	Asn	Gly
			740					745							750
Ala	Gly	Lys	Ile	Val	Leu	Leu	Gly	Arg	Gly	Glu	Pro	Ser	Ala	Ala	Ala
		755						760						765	
Gly	Lys	Ile	Leu	Arg	Thr	Leu	Asp	Ala	Arg	Phe	Ile	Arg	Ala	Asp	Leu
770						775					780				
Ser	Arg	Arg	Glu	Asp	Val	Gly	Gln	Ala	Leu	Gly	Glu	Ile	Ala	His	Ser
785					790					795					800
Met	Pro	Pro	Leu	Lys	Gly	Ile	Phe	His	Leu	Ala	Gly	Ala	Leu	Asp	Asp
				805					810						815
Ala	Leu	Leu	Thr	Arg	Gln	Asp	Asp	Asp	Phe	Phe	His	Arg	Ala	Gly	Ser
			820						825						830
Gly	Lys	Ala	Asp	Gly	Ala	Trp	Tyr	Leu	His	Glu	Leu	Thr	Ala	Gly	Leu
		835						840							845
Pro	Leu	Asp	His	Phe	Val	Leu	Phe	Ser	Ser	Met	Ala	Ala	Leu	Ile	Thr
850						855									860
Met	Pro	Gly	Gln	Gly	Asn	Tyr	Ala	Ala	Ala	Asn	Ser	Phe	Leu	Asp	Ala
865					870						875				880
Leu	Ala	Gln	His	Arg	Arg	Ala	Gln	Gly	Lys	Pro	Gly	Leu	Ser	Val	Asn
				885					890						895
Trp	Gly	Pro	Trp	Ala	Glu	Ile	Gly	His	Ala	Ala	Thr	Asp	Tyr	Gly	Arg
			900						905						910
Arg	Ala	His	Glu	Gln	Leu	Gly	Ala	Leu	Gly	Val	Gly	Thr	Leu	Pro	Pro
			915						920						925
Glu	Leu	Ala	Ile	Ala	Thr	Leu	Glu	Arg	Leu	Met	Ala	Ser	Gly	Val	Ala
			930						935						940
Gln	Ser	Gly	Val	Ala	Arg	Ile	Asp	Trp	Pro	Thr	Leu	Phe	Arg	Val	Asp
945					950						955				960



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Ala Pro Ala Ala Gly Ser Ala Leu Phe Ser Glu Leu Thr Gln Pro Ala  
                   965                                  970                                  975

Ala Gln Pro Ala Gln Gln Glu Thr Ala Leu Leu Arg Gln Leu His Ala  
                   980                                  985                                  990

Cys Ala Pro Arg Glu Arg Val Glu Arg Ile Thr Asp Thr Leu Ala Ala  
                   995                                  1000                                  1005

Met Leu Ala Glu Thr Leu Arg Leu Ser Gly Pro Asp Ala Ile Ala  
                   1010                                  1015                                  1020

Pro Glu Gln Ser Leu Leu Asp Leu Gly Leu Asp Ser Leu Val Ala  
                   1025                                  1030                                  1035

Leu Glu Leu Thr Asp Arg Leu Thr Lys Val Phe Gly Arg Pro Phe  
                   1040                                  1045                                  1050

Arg Ala Thr Leu Phe Phe Ser Tyr Pro Asn Leu Gln Thr Leu Ala  
                   1055                                  1060                                  1065

Gln Tyr Val Leu Asn Glu Leu Ser Pro Ser Leu Pro Ala Pro Val  
                   1070                                  1075                                  1080

Val Asp Glu Ala Ser Asp Asp Leu Asp Glu Asp Asp Leu Ser Glu  
                   1085                                  1090                                  1095

Leu Ile Ala Gln Glu Ile Gly Ala Gln  
                   1100                                  1105

<210> SEQ ID NO 17  
 <211> LENGTH: 1475  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(1475)  
 <223> OTHER INFORMATION: putative beta-ketoacyl synthetase

<400> SEQUENCE: 17

Met Leu Pro Asp Thr Lys Phe Arg Thr Val Thr Glu Ile Leu Leu Phe  
 1                  5                                  10                                  15

Arg Gly Lys Val Glu Pro Glu Lys Thr Ala Phe Ile Phe Leu Glu Asn  
                   20                                  25                                  30

Gly Glu Ala Glu Leu Thr Arg Leu Thr Phe Gly Asp Leu Asp Lys Arg  
                   35                                  40                                  45

Ala Arg Gly Ile Ala Ala Arg Leu Gln Ala Ile Ala Gln Pro Gly Asp  
                   50                                  55                                  60

Arg Val Leu Leu Val Tyr Pro Pro Gly Leu Glu Phe Ile Cys Ala Trp  
 65                  70                                  75                                  80

Val Gly Cys Leu Tyr Ala Gly Leu Ile Gly Val Pro Ala Tyr Pro Pro  
                   85                                  90                                  95

Arg Arg His Arg Pro Ala Asp Arg Leu Lys Ala Ile Val Ala Asp Ala  
                   100                                  105                                  110

Thr Pro Val Val Ala Leu Thr Asp Ala Ala Thr Leu Asp Gly Ile Ala  
                   115                                  120                                  125

His His Ala Asp Gly Tyr Ser Asp Thr Leu Glu Leu Lys Ile Leu Ala  
                   130                                  135                                  140

Thr Asp Gln Arg Phe Asp Ala Pro Ala Glu Gln Trp Arg Ala Pro Asp  
 145                  150                                  155                                  160

Ile Thr Pro Gln Thr Leu Ala Leu Leu Gln Tyr Thr Ser Gly Ser Thr  
                   165                                  170                                  175

Gly Thr Pro Lys Gly Val Met Ile Ser His Ala Asn Ile Leu Ser Asn  
                   180                                  185                                  190

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Met Ala Val Ile Ala Glu Ala Ser Asp Ala Asp Ala Ser Thr Val Phe  
 195 200 205  
 Val Ser Trp Leu Pro Val Phe His Asp Met Gly Phe Phe Gly Lys Val  
 210 215 220  
 Leu Leu Pro Ile Tyr Leu Gly Val Leu Ser Val Leu Met Ala Pro Ala  
 225 230 235 240  
 Ala Phe Val Gln Lys Pro Val Arg Trp Leu Gln Ala Ile Thr Lys Tyr  
 245 250 255  
 Arg Gly Thr His Cys Ala Ala Pro Asp Phe Ala Tyr Asp Leu Cys Ala  
 260 265 270  
 Arg Lys Ile Ala Asp Glu Ala Arg Ala Gln Leu Asp Leu Ser Ser Trp  
 275 280 285  
 Arg Val Ala Phe Asn Gly Ala Glu Pro Val Arg Ala Glu Ser Val Ala  
 290 295 300  
 Arg Phe Ser Arg Ala Phe Ala Ala Cys Gly Phe His Ala His Thr Met  
 305 310 315 320  
 Arg Pro Val Tyr Gly Met Ala Glu Ala Thr Leu Phe Ile Ser Gly Gln  
 325 330 335  
 Pro Ala Arg Ser Leu Pro Arg Val Ala Asp Tyr Asp Ala Asp Ala Leu  
 340 345 350  
 Ala Gln Gly Val Ala Thr Arg Asn Asp Ser Gly Lys Arg His Ala Leu  
 355 360 365  
 Val Ser Cys Gly Arg Thr Trp Ala Glu His Arg Val Arg Ile Val Asn  
 370 375 380  
 Pro Asp Thr Gly Glu Arg Cys Ala Pro Gly Arg Ile Gly Glu Ile Trp  
 385 390 395 400  
 Leu Thr Gly Pro Ser Val Gly Val Gly Tyr Trp Asn Arg Ile Asp Glu  
 405 410 415  
 Thr Glu Arg Thr Phe Arg Ala Lys Leu Asp Gly Asp Asp Ala Arg Tyr  
 420 425 430  
 Leu Arg Thr Gly Asp Leu Gly Phe Val Asp Gly Glu Asp Leu Phe Val  
 435 440 445  
 Thr Gly Arg Leu Lys Asp Leu Ile Ile Val Ala Gly Arg Asn His Tyr  
 450 455 460  
 Pro Gln Asp Leu Glu Gln Ser Ala Glu Gly Ser His Pro Ala Leu Ala  
 465 470 475 480  
 Pro Asn Ala Ser Ala Ala Phe Ser Ile His Val Asp Asn Val Glu Arg  
 485 490 495  
 Val Val Val Ala Cys Glu Val Arg Arg Glu Ala Leu Asn Thr Leu Asp  
 500 505 510  
 Ala Glu Ala Val Ala Ala Glu Ile Arg His Thr Leu Ala Glu Val His  
 515 520 525  
 Asp Val Asp Leu Tyr Ala Ala Val Leu Leu Lys Pro Ala Thr Ile Leu  
 530 535 540  
 Arg Thr Ser Ser Gly Lys Ile Gln Arg Ser Arg Ile Arg Gln Ala Phe  
 545 550 555 560  
 Leu Asp Glu Gln Gly Leu Ala Ile Ala Gly Glu Trp Arg Arg Ala Phe  
 565 570 575  
 Ser Ala Pro Pro Ala Pro Pro Gln Thr Ala Glu Pro Arg Asp Thr Gln  
 580 585 590  
 Ala Leu Val Gln Trp Cys Ile Glu Arg Val Ser Arg Leu Ser Gly Ile  
 595 600 605  
 Ala Ser Gly Lys Ile Asp Pro Asp Ala Pro Phe Ser Val His Gly Leu

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610			615			620									
Asp	Ser	Lys	Asp	Ala	Ile	Met	Leu	Ser	Gly	Glu	Leu	Gln	Asp	Trp	Leu
625				630						635					640
Gly	Arg	Pro	Val	Ser	Pro	Thr	Val	Val	Tyr	Asp	Phe	Pro	Ser	Ile	Ser
			645						650						655
Leu	Leu	Ala	Arg	His	Leu	Ser	Gly	Thr	Gly	Ser	Ala	Met	Pro	Asp	Gln
			660						665				670		
Ala	Pro	Gly	Ser	Ala	Glu	Ala	Arg	Ala	Asp	Ile	Ala	Ile	Val	Gly	Met
		675					680						685		
Gly	Cys	Arg	Phe	Pro	Gly	Ala	Gly	Asn	Pro	Asp	Ala	Phe	Trp	Gln	Leu
690						695					700				
Leu	Leu	Glu	Gly	Arg	Asp	Ala	Val	Gly	Ala	Ala	Thr	Gln	Arg	Ala	Ala
705				710						715					720
Asp	Leu	Pro	Leu	Ala	Gly	Leu	Leu	Asp	Gln	Val	Asp	Gln	Phe	Asp	Ala
			725						730						735
Ala	Phe	Phe	Gly	Ile	Ser	Ala	Arg	Glu	Ala	Glu	Ser	Met	Asp	Pro	Gln
			740						745				750		
Gln	Arg	Leu	Leu	Leu	Glu	Val	Ala	Trp	Glu	Thr	Leu	Glu	His	Ala	Gly
		755					760						765		
Ile	Ala	Pro	Arg	Ser	Leu	Ala	Gly	Gly	Arg	Thr	Ala	Val	Ile	Val	Gly
770					775					780					
Ile	Ser	Asn	Ser	Asp	Tyr	Ile	Arg	Leu	Ala	Gln	Asp	Glu	Val	Ala	Asp
785				790						795					800
Val	Gly	Pro	Tyr	Val	Ala	Thr	Gly	Asn	Ala	Leu	Ser	Val	Ala	Ala	Asn
			805						810						815
Arg	Ile	Ser	Tyr	Ala	Leu	Asp	Leu	Arg	Gly	Pro	Ser	Trp	Ala	Val	Asp
			820						825				830		
Thr	Ala	Cys	Ser	Ser	Ser	Leu	Val	Ala	Val	His	Gln	Ala	Cys	Arg	Ala
		835					840						845		
Leu	Gln	Arg	Gly	Glu	Ser	Asp	Ala	Ala	Leu	Ala	Gly	Gly	Val	Asn	Leu
850						855					860				
Ile	Leu	Ala	Pro	Gln	Leu	Ser	Ala	Ser	Phe	Thr	Gln	Ala	Gly	Met	Leu
865				870						875					880
Ser	Pro	Asp	Gly	Arg	Cys	Lys	Ala	Phe	Asp	Ala	Ala	Ala	Asn	Gly	Tyr
			885						890						895
Val	Arg	Gly	Glu	Gly	Val	Gly	Met	Val	Leu	Leu	Lys	Arg	Leu	Asp	Asp
			900						905				910		
Ala	Leu	Glu	Asn	Gly	Asp	Thr	Val	Phe	Ala	Val	Ile	Arg	Gly	Ser	Ala
		915					920						925		
Val	Asn	Gln	Asp	Gly	Arg	Ser	Asn	Gly	Leu	Thr	Ala	Pro	Asn	Gly	Pro
930						935					940				
Ala	Gln	Gln	Ala	Val	Ile	His	Gly	Ala	Leu	Arg	Asp	Ala	Gly	Val	Arg
945				950						955					960
Ala	Gln	Asp	Ile	Gly	Phe	Val	Glu	Thr	His	Gly	Thr	Gly	Thr	Pro	Leu
			965						970						975
Gly	Asp	Pro	Ile	Glu	Leu	Asn	Ser	Leu	Ala	Ala	Val	Leu	Asn	Glu	Ser
			980						985				990		
Arg	Arg	Pro	Asp	Asp	Leu	Cys	Trp	Ile	Gly	Ser	Val	Lys	Thr	Asn	Ile
			995				1000						1005		
Gly	His	Leu	Glu	Ser	Ala	Ala	Gly	Ile	Ala	Ser	Leu	Ile	Lys	Thr	
	1010						1015						1020		
Ala	Leu	Ala	Leu	His	His	Arg	Ala	Ile	Pro	Pro	Asn	Leu	His	Phe	
	1025						1030						1035		

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Arg	Ser	Ile	Asn	Pro	Gln	Ile	Ala	Leu	Asp	Gly	Thr	Pro	Phe	Arg
1040						1045					1050			
Ile	Pro	Arg	Gln	Val	Thr	Pro	Trp	His	Ser	Glu	His	Gly	Pro	Arg
1055						1060					1065			
Leu	Ala	Gly	Val	Ser	Ser	Phe	Gly	Phe	Gly	Gly	Thr	Asn	Ala	His
1070						1075					1080			
Leu	Ile	Leu	Ser	Glu	Ala	Pro	Gly	Leu	Pro	Glu	Ile	Glu	Ala	Glu
1085						1090					1095			
Pro	Val	Ala	Pro	Ala	Ala	Arg	Val	Val	Thr	Leu	Ser	Ala	Arg	Thr
1100						1105					1110			
Pro	Asp	Ala	Leu	Gln	Ala	Leu	Ala	Ala	Ser	Tyr	Ala	Ala	Tyr	Leu
1115						1120					1125			
Asp	Ala	His	Pro	Glu	Ala	Gly	Val	Arg	Asp	Val	Ala	Phe	Thr	Ala
1130						1135					1140			
Asn	Thr	Gly	Arg	Thr	His	Phe	Thr	Gln	Arg	Ala	Ala	Ile	Val	Ala
1145						1150					1155			
Pro	Ser	Arg	Asp	Ser	Leu	Arg	Ala	Gln	Leu	Asp	Ser	Val	Ser	Ser
1160						1165					1170			
Gly	Glu	Pro	Ala	Glu	Thr	Pro	Pro	Ala	Val	Thr	Phe	His	Phe	Cys
1175						1180					1185			
Ala	Asp	Asp	Gly	Ala	Ser	Ala	Asp	Ala	Val	Arg	Gln	Leu	Arg	Ala
1190						1195					1200			
Ala	Ser	Pro	Ala	Phe	Asp	Ala	Leu	Met	Gln	Arg	Gln	Ser	Asp	Ala
1205						1210					1215			
Ser	Gly	Ala	Pro	Ala	Leu	Ala	Pro	Asp	Glu	Ala	Gly	Phe	Thr	Arg
1220						1225					1230			
Phe	Gln	Arg	Ala	Leu	Ala	Gln	Leu	Trp	Met	Ser	Phe	Gly	Ile	Ala
1235						1240					1245			
Pro	Asp	Ala	Val	Ser	Ser	Thr	Gly	Asp	Gly	Gln	Arg	Ala	Ala	Ala
1250						1255					1260			
Ala	Trp	Ala	Gly	Val	Pro	Gln	Ala	Pro	Asp	Ser	Gly	Ala	Ala	Gly
1265						1270					1275			
His	Pro	Gly	Ile	Val	Ile	Asp	Ile	Gly	Ala	His	Thr	Ala	Ala	Trp
1280						1285					1290			
Asp	Ala	Ile	Leu	His	Thr	Leu	Ala	Ala	Leu	Tyr	Val	Arg	Gly	Ala
1295						1300					1305			
Ser	Ile	Asp	Trp	Asp	Ala	Val	Glu	Gln	Gly	Ala	Pro	His	Arg	Arg
1310						1315					1320			
Leu	Ala	Leu	Pro	Thr	Tyr	Pro	Phe	Glu	Arg	Arg	Gly	Phe	Trp	Ile
1325						1330					1335			
Arg	Pro	His	Ala	Arg	Arg	His	Pro	Leu	Leu	Gly	Arg	Leu	Met	Glu
1340						1345					1350			
Gln	His	Ala	His	Ala	Pro	Ala	Thr	Trp	Ile	Trp	Gln	Ser	Arg	Leu
1355						1360					1365			
Asp	Ala	Pro	Ala	Thr	Asn	Phe	Leu	Asp	Gly	His	Arg	Val	Lys	Gly
1370						1375					1380			
Ser	Pro	Val	Leu	Pro	Tyr	Ser	Ala	Phe	Val	Glu	Met	Ala	Leu	Ser
1385						1390					1395			
Ala	Thr	Ser	Glu	Ile	Gly	Ala	Ala	Gly	His	Thr	Thr	Leu	Lys	Asp
1400						1405					1410			
Leu	Ala	Leu	His	Ala	Pro	Leu	Pro	Leu	His	Pro	His	Glu	Ser	His
1415						1420					1425			

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Thr Val Gln Thr Val Leu Ser Arg Arg Ser Trp Gly Pro Phe Ser  
1430 1435 1440

Phe Ala Val Tyr His Arg Ile Asp Asp Thr Arg Ala Ala Ala Thr  
1445 1450 1455

Trp Gln Met Cys Ala Ser Ala Glu Ile His Glu Ser Asp Arg Ser  
1460 1465 1470

His Ala  
1475

<210> SEQ ID NO 18  
 <211> LENGTH: 328  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(328)  
 <223> OTHER INFORMATION: putative taurine catabolism dioxygenase

<400> SEQUENCE: 18

Met Leu Gly Met Thr Glu Arg Lys Leu Leu Ala Glu Gly Ser Thr Pro  
1 5 10 15

Trp Leu Leu Glu Pro Val Ser Asn Gly Arg Asp Leu Ala Gln Ala Val  
20 25 30

Asn Asp Asn Arg Ala Ala Leu Glu Ser Arg Leu Leu Glu His Gly Val  
35 40 45

Leu Leu Phe Arg Gly Phe Asp Val Ser Ser Val Gly Gly Phe Glu Ala  
50 55 60

Phe Ala Arg Ala Ile Ser Ala His Gln Ser Asp Tyr Val Tyr Arg Ser  
65 70 75 80

Thr Pro Arg Thr Ser Ile Gly Asn Gly Ile Phe Thr Ala Thr Glu Tyr  
85 90 95

Pro Pro Ser Glu Thr Ile Ala Leu His Cys Glu Asn Ala Tyr Gln Arg  
100 105 110

Ser Trp Pro Leu Arg Val Ala Phe Cys Cys Leu Thr Pro Ala Ala Thr  
115 120 125

Gly Gly Glu Thr Pro Ile Ala Asp Met Arg Glu Val Ser Arg Arg Ile  
130 135 140

Gly Pro Arg Ile Leu Asp His Phe Glu Ala Arg Gln Val Arg Tyr Val  
145 150 155 160

Arg His Tyr Arg Arg His Val Asp Ile Pro Trp Glu Thr Val Phe Gln  
165 170 175

Thr Ser Asp Arg Asn Gln Val Ala Ala Phe Cys Ala Asp Asn Gly Ile  
180 185 190

Ala Leu Glu Trp Leu Asp Asp Thr Leu Arg Thr Ala Gln Ile Asn  
195 200 205

Gln Gly Val Ala Tyr His Pro Val Thr Gly Glu Arg Val Phe Phe Asn  
210 215 220

Gln Ala His Leu Phe His Ile Ser Asn Leu Glu Ala Ser Leu Ala Ser  
225 230 235 240

Ser Ile Val Ser Leu Phe Gly Glu Asp Arg Ile Pro Arg Asn Ala Cys  
245 250 255

His Gly Asp Gly Ser Pro Phe Asp Leu Ala Asp Leu Glu Gln Ile Arg  
260 265 270

His Ala Phe Arg Glu Cys Ala Ile Thr Phe Pro Trp Gln Arg Gly Asp  
275 280 285

Val Leu Leu Val Asp Asn Met Arg Phe Ala His Gly Arg Asn Pro Phe



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Ala Gly Thr Phe Arg Gly Asn Gln Ile Ala Met Val Ala Gly Leu Ser  
 325 330 335  
 Thr Met Arg Ile Val Glu Arg Glu Asp Leu Ser Ala His Ala Asp Arg  
 340 345 350  
 Val Gly Lys Leu Leu Val Ala Gly Leu Glu Glu Leu Ala Glu Arg Phe  
 355 360 365  
 Pro Cys Leu Gly Gln Ile Arg Gly Arg Gly Leu Met Ile Gly Ala Glu  
 370 375 380  
 Val Val Val Pro Gly Thr His Gly Arg Ala Gly Pro Pro His Thr Glu  
 385 390 395 400  
 Arg Ala Arg Ala Ile Lys Gln Asn Cys Leu Arg Asn Gly Leu Ile Val  
 405 410 415  
 Glu Thr Gly Gly Arg Asn Gly Ala Val Leu Arg Phe Leu Pro Pro Leu  
 420 425 430  
 Ile Val Ser Glu Ala Asp Ile His Asp Ile Leu Asn Arg Phe Glu His  
 435 440 445  
 Ala Val Glu Thr Ala Cys Arg Ala  
 450 455

<210> SEQ ID NO 20  
 <211> LENGTH: 316  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(316)  
 <223> OTHER INFORMATION: putative epemerase/dehydratase

<400> SEQUENCE: 20

Met Gln Arg Asn Arg Lys Arg Ile Leu Val Thr Gly Gly Ala Gly Phe  
 1 5 10 15  
 Leu Gly Ser His Leu Cys Glu Arg Leu Val Glu Leu Gly His Asp Val  
 20 25 30  
 Leu Cys Val Asp Asn Tyr Phe Thr Gly Thr Lys Gln Asn Val Ala Thr  
 35 40 45  
 Leu Leu Gly Asn Pro Ser Phe Glu Ala Leu Arg His Asp Val Thr Phe  
 50 55 60  
 Pro Leu Tyr Val Glu Val Asp Glu Ile Tyr Asn Leu Ala Cys Pro Ala  
 65 70 75 80  
 Ser Pro Ile His Tyr Gln Phe Asp Pro Val Gln Thr Thr Lys Thr Ser  
 85 90 95  
 Val Met Gly Ala Ile Asn Met Leu Gly Leu Ala Lys Arg Thr His Ala  
 100 105 110  
 Arg Val Leu Gln Thr Ser Thr Ser Glu Val Tyr Gly Asp Pro Asp Val  
 115 120 125  
 His Pro Gln Pro Glu Ser Tyr Arg Gly Asn Val Asn Pro Leu Gly Pro  
 130 135 140  
 Arg Ala Cys Tyr Asp Glu Gly Lys Arg Cys Ala Glu Thr Leu Phe Phe  
 145 150 155 160  
 Asp Tyr His Arg Gln Gln Asn Val Arg Ile Lys Val Val Arg Ile Phe  
 165 170 175  
 Asn Thr Tyr Gly Pro Arg Met His Pro Asn Asp Gly Arg Val Val Ser  
 180 185 190  
 Asn Phe Ile Val Gln Ala Leu Arg Gly Glu Asp Ile Thr Leu Tyr Gly  
 195 200 205

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Asp Gly Ser Gln Thr Arg Ala Phe Cys Tyr Val Asp Asp Met Val Asp  
 210 215 220  
 Gly Leu Ile Arg Met Met Ala Thr Pro Ala Glu Leu Thr Gly Pro Ile  
 225 230 235 240  
 Asn Leu Gly Asn Pro His Glu Ile Ala Val Ser Glu Leu Ala Gln Ile  
 245 250 255  
 Ile Leu Arg Leu Thr Gly Ser Lys Ser Arg Leu Val Phe Arg Pro Leu  
 260 265 270  
 Pro Lys Asp Asp Pro Thr Gln Arg Cys Pro Asp Ile Ser Leu Ala Arg  
 275 280 285  
 Thr His Leu Asp Trp Glu Pro Thr Ile Gly Leu Glu Ala Gly Leu Gln  
 290 295 300  
 Arg Thr Ile Asp Tyr Phe Cys Ser Thr Leu Ala Ala  
 305 310 315

<210> SEQ ID NO 21  
 <211> LENGTH: 239  
 <212> TYPE: PRT  
 <213> ORGANISM: Burkholderia contaminans  
 <220> FEATURE:  
 <221> NAME/KEY: MISC\_FEATURE  
 <222> LOCATION: (1)..(239)  
 <223> OTHER INFORMATION: putative thioesterase

<400> SEQUENCE: 21

Met Arg Leu Ile Cys Phe Pro Tyr Ala Gly Gly Ser Ala Ala Val Tyr  
 1 5 10 15  
 Arg Thr Leu Gln Ala Ser Leu Pro Gly Ile Glu Val Cys Arg His Glu  
 20 25 30  
 Leu Ala Gly Arg Gly Ser Arg Leu Ser Glu Pro Ala Val Arg Asp Met  
 35 40 45  
 Ala Thr Leu Val Asp Thr Leu Leu Cys Asp Leu Asp Asp Cys Phe Asp  
 50 55 60  
 Arg Pro Phe Ala Leu Leu Gly His Ser Met Gly Ala Ala Ile Ala Ala  
 65 70 75 80  
 Glu Leu Ala Leu Arg Leu Pro Ala His Ala Arg Pro Asn Leu Arg His  
 85 90 95  
 Leu Phe Val Ser Ala Arg Ala Ala Pro Gly Lys Glu Arg His Asp Arg  
 100 105 110  
 Arg Met Gln Ala Leu Asp Asp Arg Ala Phe Ile Asp Ala Leu Arg Glu  
 115 120 125  
 Met Gly Gly Thr Pro Lys Ala Val Leu Asp Asn Ser Glu Leu Met Ala  
 130 135 140  
 Leu Leu Met Pro Ala Leu Arg Ala Asp Phe Thr Met Ile Glu Asn His  
 145 150 155 160  
 Arg Pro Val Pro Gly Pro Arg Leu Ala Val Asp Ile Thr Ala Phe Ala  
 165 170 175  
 Gly Arg Ala Asp Lys Glu Ile Pro Val Asp Ala Val Ala Gly Trp Gly  
 180 185 190  
 Ala Ala Thr Thr Gly Arg Phe Asp Phe His Val Ile Glu Gly Asp His  
 195 200 205  
 Phe Phe Leu Arg Asn Glu Met Arg Thr Met Ala Gly Ile Ile Ala Ala  
 210 215 220  
 Arg Met Arg Arg Pro Glu His Ala Ala Ser Ser Ala Leu Gln Ala  
 225 230 235



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<210> SEQ ID NO 22
<211> LENGTH: 95
<212> TYPE: PRT
<213> ORGANISM: Burkholderia contaminans
<220> FEATURE:
<221> NAME/KEY: MISC_FEATURE
<222> LOCATION: (1)..(95)
<223> OTHER INFORMATION: hypothetical protein

<400> SEQUENCE: 22

Met Gln His Arg Gln Lys Ala Val Pro Thr Gln Gln Val Ala Asn Glu
1          5          10          15
Arg Val Ile Val Thr Glu Trp Arg Phe Ala Pro Gly Ala Glu Thr Gly
          20          25          30
Trp His Val His Arg His Asp Tyr Val Val Val Pro Gln Thr Asp Gly
          35          40          45
Gln Leu Leu Leu Glu Thr Ala Gln Gly Asn Arg Glu Ser Gln Leu His
50          55          60
Ala Gly Arg Ser Tyr Ala Gly Leu Lys Gly Val Glu His Asn Val Val
65          70          75          80
Asn Ala Thr Asp His Glu Val Val Phe Val Glu Val Glu Ile Leu
          85          90          95

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<210> SEQ ID NO 23
<211> LENGTH: 58101
<212> TYPE: DNA
<213> ORGANISM: Burkholderia contaminans

<400> SEQUENCE: 23

aattcctgca gcacgggtgcg cgaccagccc cagatgtccc cgctgagcgt gagtgcgaga      60
ccggccgtcg tgatggccag ctgcgtctgg ccgaacacgc gcgtcaatgc gccttcgccg      120
ccgatcacga tccgcttgac gagatccgag atggactcgc agatcgaatc ggagaacgga      180
tagttgtaag gctgcgtgac ggcgcgcgac aggaacggct tgctgggctg cggcgtccag      240
accttgagcc acggtttggt cgtgaacggg aaccagatgg cttccaccgc gcccgagccg      300
tcgagaaaac atcgcatcgt gcggcccgtc gtgccggcgc cggcgaaacag ttcggaggcc      360
ggaatatcga cgtagctctg gcagcgtagc cgctggttcg gccctgccgt cagcgtgact      420
tcgacgacga gcgctcgcgc gatgtgcgcg aggaacgcgc cgatctcggg atcgctgcgc      480
tcgaaccggc gcagcacgta ttgctgcccg gccgtagcga acacgaccgc cgtgagcgcg      540
accacgagat tgctcagcga gccgtaggta tggcccgggt gcaaggtttc accggcccgcg      600
ggcacggcgg tgccgtgtgc atcgatcgcg agcgcgccgc cgagcgtgat gtcgcccggt      660
gccggcgcgg caatcacgcc gaggccaacc tgctcgagcg tcgagagcag cgactccagc      720
gagacgcccg tttggggcgt gacgcgcgcc ggacgcgccg acgtgtcgcg ggagacggcc      780
gtcagcgact tcgtcgtatc gacgacacg aggttcgcgg cgccggcgcg cgggtccagc      840
gtcagcggcg accagttgtg cgtgtagccg cgcgggcgta tccgatagcc gtttgcgcgc      900
gcccagttga cggttgcgac gacgtcgtcg gcggagcgcg gcgcggcggg ccatacgtcc      960
tgcaaggcga tctcgcgcgt ccagttcagg aacgcctgct tgtaaagctg gatgtcggcc      1020
gggaagccgg gcggtgtctc gccggccggt cgcgcgtgcg ccgcaacctg gtagagcggg      1080
gtccagcccg tgacgatgcc ggccgcccgc agcttcgcca tgtcggccag gaaggcgcga      1140
cgcggcgcag gttcgtctct gaagtcgtga ctcatggtgt gtcccaattt ttcggaattg      1200
ttttgcagat tggaaagacg acaaatgacg cgttgagact cgtgtggcaa ttcgagcagg      1260

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tgcgacgcgc	gggaagtgtt	gcgcgtgggt	gggccaggat	tgaaaaaaga	cggtgcgttc	1320
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gattgcttta	attggcgtc	ggccgttct	gtcggcaagc	agataggag	attcgacgga	1440
atcgcgcgcg	gcgaagcgt	agccgtggcg	atcgataaaa	gatgatttca	cgtgaatatt	1500
aatcttcatt	tttcgatttt	taaataaacc	cgccgcagc	tcaaggtga	ttgacgatgc	1560
gtcatgcatt	tcggtcgaaa	gcgtagcaat	ttatctatcg	ggtgacaagc	ggcggagttg	1620
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agctgaagcc	gggtcgatga	gcgggaggtg	gggtgaaatc	cgataattcc	tctctcgaat	1860
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ccggtatgact	gaatggacgc	tcgacgagcc	ggcggggcaa	gtcgtccgcg	tgcaatcgct	2640
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ggaaccggca	gcgatccgc	cgttgaaccg	gatcctggcg	gcctgcgacc	ggcagctcat	2760
tcacatcaat	ccgctgatgc	ggcgcggcaa	tggcggcgaa	gtcgcgcctg	cgcgcgggcc	2820
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gcgggaagg	gcgacggcac	ccgggtgcgac	ggcaggcatg	cccggcgtcg	aggcgtcgcg	3060
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gaaggagagc	acgatcgaga	cgtacatcaa	gcgagccgcg	gtgaagctcg	gcatcagcgg	3240
ccggcacggg	ctcacgaaat	ggatgatcga	cgattccgta	ccgtgcgcgt	cgccggcgtg	3300
acaccgtcac	gccatcacgc	cgccgacgcg	cgacgatcgc	cgccgggcat	gcgcgttcgg	3360
gccgcgggcc	ctcaggttcc	gagggcgggc	gacgcgtagt	cgccgcgcac	gcttctcatg	3420
tcgccccag	cagcttcctg	cgccggcccc	gccgtctcgt	cgagcatcca	gcgcgtcagg	3480
ccatgacgcc	cgctgaagcc	cagcttgacg	gccgcccgct	tcaggtaggt	ttcgacggtg	3540
ctttcgcgca	gcgcgaagcg	catggcgtatg	gcaggcaccg	tgtcaccggc	caggagtgcc	3600
gtgcatgcct	cgatctcgcg	cgctcagacg	ttgacgcccg	cttgcgtcag	gcgatcggcg	3660

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aatcgccgcg	ccacgcgctc	ccggcccgat	tgcgtcgccg	gcggcgcggc	ggtcgtcaca	3720
cgagcggcgg	gcggagccga	atcgagcgcc	gcgacatggc	tctcgacgat	cgaaaacagc	3780
acgtgcgaga	gttccttgag	gaaggtccgc	tctgcgccg	agaaatcgtc	gaacgtacag	3840
gtgcgataca	acgagatcac	gtaacagtgg	ccccgcttgc	gggtcacgag	gtggaattgc	3900
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atcggggcgt	ggccggcgag	cgtgtcgtcg	acgtgcaggg	ggctcgtgcc	cgggcgcggc	4020
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gcgcccaggc	tgcgcacctc	gggagggcgg	tccggcacgt	cgtcgatcgc	aagctccgaa	4140
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ccgtatcgga	gaagatcgcg	ccgctatcca	ggcgaatgac	ccgatccgcc	agcttgaagt	4560
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We claim:

1. A method of increasing production of occidiofungin diastereomers/conformers corresponding to the diastereomers/conformers having the total correlation spectroscopy (TOCSY) fingerprint set forth in FIG. 5C as the green NH correlations in a microorganism producing occidiofungin comprising:

decreasing the thioesterase activity of an occidiofungin gene D (ocfD) product (OcfD) in said microorganism, wherein said OcfD thioesterase activity is decreased by the introduction of a point mutation at the serine in position 2954 of the amino acid sequence of SEQ ID NO: 4, said point mutation replaces said serine with an alanine, glycine or proline, and culturing said microorganism under conditions to produce said occidiofungin diastereomers/conformers,

wherein said increase in production of occidiofungin diastereomers/conformers is as compared to a reference microorganism without said decrease in OcfD thioesterase activity cultured under the same conditions.

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