Supplementary File 2

A. MultiGeneBlast search with the *Leptosphaeria maculans* sirodesmin biosynthesis gene cluster as query. The results match those of Fox & Howlett (2008), who found similarity between this sirodesmin biosynthesis gene cluster and the gene cluster for the biosynthesis of gliotoxin from *Aspergillus fumigatus* (shown in the results as hit nr 5). Two gene clusters with still different architectures from *Aspergillus flavus* and *Penicillium marneffei* show up as hits nrs 3 and 4, and may encode different and yet unknown molecules from the same structural family.



B. MultiGeneBlast search with the *Aspergillus nidulans* aflatoxin/sterigmatocystin biosynthesis gene cluster as query. The results match those of Slot & Rokas (2011), who showed that *Podospora anserina* possesses a gene cluster highly similar to those in *Aspergillus nidulans* and *Aspergillus flavus*. The *P. anserina* gene cluster is hit nr 3. Additionally, a number of other homologous gene clusters from *Aspergillus* and one from *Emericella* are identified by MultiGeneBlast.

multigene Search Results BLAST Results pages: 1, 2, 3, 4, 5
MultiGeneBlast hits
Select gene cluster alignment
Query: Aspergillus nidulans FGSC A4 chromosome IV ANcontig1.132, whole
1. ; AACD01000132 Aspergillus nidulans FG5C A4 Total score; 31.5 Cumulative Blast bit score; 276040
2. ; AY092402 Aspergillus ochraceoroseus strain SRRC 1492 allatoxin/sterigmatocystin gene cluster 🛛 Total score; 29.0 — Cumulative Blast bit score; 196760
3. ; CU640366 Podospora anserina 5 mat+ genomic DNA chromosome 2, supercontig 2. Total score; 29.0 Cumulative Blast bit score; 160040
4. ; EQ963478 Aspergillus flavus NRRL3357 scf_1106286417182 genomic scaffold 🛛 Total score ; Z2.5 🦳 Cumulative Blast bit score ; 144450
5. ; U34740 Emericella nidulans sterigmatocystin biosynthetic gene cluster: (stcA), (stcB), (stcC), Total score: 22.0 Cumulative Blast bit score: 221930
6. : AKHY01000106 Aspergillus oryzae 3.042 Total score: ZZ.0 Cumulative Blast bit score: 141170
7. ; AY510451 Aspergillus flavus isolate AF13 aflatoxin biosynthesis gene cluster Total score; 21.5 Cumulative Blast bit score; 143430
8. ; AY510453 Aspergillus flavus isolate AF70 aflatoxin biosynthesis gene cluster Total score; 21.5 Cumulative Blast bit score; 142320
9. ; AB196490 Aspergillus oryzae DNA, aflatoxin biosynthesis gene cluster Total score; 21.5 Cumulative Blast bit score; 140870
10. : A Y510454 Aspergillus nomius isolate AN13137 affatoxin biosynthesis gene cluster Total score: 21.0 Cumulative Blast bit score: 146400
11. : AY510452 Aspergillus flavus isolate BN008 aflatoxin biosynthesis gene cluster Total score: 21.0 Cumulative Blast bit score: 145520
12. : A Y 3/ 149V Aspergillus parasilicus altatoxin paltway gene cluster Total score: 21.0 Cumulative Blast bit score: 145460

C. MultiGeneBlast search with the clorobiocin biosynthesis gene cluster as query, matching the results of Pojer (2011), who compared the gene clusters encoding the biosynthesis of clorobiocin, coumermycin (hit nr 2) and novobiocin (hit nr 3), which have different combinations of subclusters encoding different combinations of chemical moieties in the final compound. Besides these gene clusters, evolutionary similarities with the gene clusters encoding phenalinolactone (hit nr 4) and simocyclinone (hits nr 8-10) are also detected, matching subclusters encoding the shared pyrrole and coumarin chemical moieties, respectively.

multigene Search Results BLAST Results pages: 1, 2, 3, 4, 5	
MultiGeneBlast hits	?
Select gene cluster alignment Query: Streptomyces roseochromogenes subsp. oscitans clorobiocin	Display all
1, : AF329398 Streptomyces roseochromogenes subsp. oscitans clorobiocin biosynthetic gene cluster Total score: 41.5 Cumulative Blast bit	score: 190660
2. : AF235050 Streptomyces rishiniensis strain D5M 40489 hypothetical protein gene Total score: 32.0 Cumulative Blast bit score: 138770	
3. : AF170880 Streptomyces caeruleus strain NCIB 11891 novobiocin biosynthetic gene cluster Total score: 25.0 Cumulative Blast bit score:	109370
4. : DQ230532 Streptomyces sp. Tu6071 phenalinolactone biosynthetic gene cluster Total score: 9.0 Cumulative Blast bit score: 30790	
	·
6. ; CP002040 Nocardiopsis dassonvillei subsp. dassonvillei D5M 43111 chromosome 1 Total score; 7.5 Cumulative Blast bit score; 17920	
	·
8. : AF324838 Streptomyces antibioticus simocyclinone biosynthetic gene cluster Total score: 6.0 Cumulative Blast bit score: 18640	
9. : AF322256 Streptomyces antibioticus simocyclinone biosynthetic gene cluster Total score: 6.0 Cumulative Blast bit score: 18580	
10. : AF321122 Streptomyces antibioticus simocyclinone biosynthetic gene cluster Total score: 6.0 Cumulative Blast bit score: 18580	
11. : CP002047 Streptomyces bingchenggensis BCW-1 Total score: 5.5 Cumulative Blast bit score: 10410 DDDDDXDDXDDX CV CV CV 12. : AM889123 Streptomyces olivaceus NDP-L-rhamnose biosynthesis gene cluster, galE3 and proB genes, Total score: 5.0 Cumulative B]

D. MultiGeneBlast search with the cladoniamide biosynthesis gene cluster as query, matching the results of Ryan (2011), who compared the gene clusters encoding the biosynthesis of cladoniamide, BE-54017, rebeccamycin, AT2433, K252a and staurosporine.

multigene BLAST	Search Results Results pages: 1, 2, 3, 4, 5
MultiGeneBlast hits	•
Select gene cluster alignment Query: Streptomyces uncialis putative	Display all Display all
1. : IN165773 Streatomyres uorialis au	tative RibD domain-containing organic, gutative transcriptional Total score: 19.0 Cumulative Rist bit score: 128599
2. : JF439215 Uncultured bacterium AB	1650 antitumor substance BE-54017 gene cluster Total score: 18.0 Cumulative Blast bit score: 74930
3. : AJ414559 Saccharothrix aerocoloni	igenes gene cluster for rebeccamycin biosynthesis, Total scorer 7.5 Cumulative Blast bit scorer 30370
4. : AF534707 Lechevalleria aerocolonig	genes putative regulatory protein, putative antibiotic antiport Total score: 7.5 Cumulative Blast bit score: 30370
5. : AB090952 Lechevalleria aerocoloni <u>v</u>	genes rebeccamycin biosynthetic gene cluster Total score: 7.5 Cumulative Blast bit score: 30370
6. : AB071405 Lechevalleria aerocolonig	genes rebeccamycin biosynthetic gene cluster (orf6, rebG, rebG,, Total score: 7.5 Cumulative Blast bit score: 30370
7. : DQ297453 Actinomadura melliaura .	strain SCC 1655 NADH oxidase and hypothetical proteins genes Total score: 6.5 Cumulative Blast bit score: 29440
8. : F3031030 Nonomuraea longicalena	strain NRRL 15532 putative transcriptional regulator gene Total score: 5.5 Cumulative Blast bit score: 23370
9. : CP000850 Salinispora arenicola CN	5-205 Total score: 5.0 Cumulative Blast bit score: 29670
10. : AB088119 Streptomyces sp. TP-A	0274 staurosponine biosynthetic gene cluster (staR, staB, staA, st Total score: 5.0 Cumulative Blast bit score: 29350
11, : CM000914 Streptomyces clavulige	rus ATCC 27064 plasmid p5CL4 Total score: 5.0 Cumulative Blast bit score: 26920
12. : FR845719 Streptomyces venezuel.	ae ATCC 10712 complete genome, Total score: 5.0 Cumulative Blast bit score: 13630

E. MultiGeneBlast search with the *Xanthomonas gardneri* lipopolysaccharide biosynthesis gene cluster as query. The results match those of Potnis et al. (2011), who compared the gene clusters of *Xanthomonas gardneri*, *Xanthomonas vesicatoria*, and *Xanthomonas campestris pv. vesicatoria*. Additionally, a number of additional gene cluster variants from other species can be observed.

multigene Search Results BLAST Results pages: 1, 2, 3, 4, 5	
MultiGeneBlast hits	?
Select gene cluster alignment Image: Control of the second se	Display all
1. ; AEQX01000135 Xanthomonas gardneri ATCC 19865 Total score; 22.0 Cumulative Blast bit score; 122270	
2. : AEQV01000058 Xanthomonas vesicatoria ATCC 35937 Total score: 22.0 Cumulative Blast bit score: 107480	
3. : AF204145 Xanthomonas campestris pv. campestris LPS biosynthetic cluster Total score: 22.0 Cumulative Blast bit score: 99210	
4. : AM920689 Xanthomonas campestris pv. campestris complete genome, strain B100. Total score: 22.0 Cumulative Blast bit score: 99100	
5. : CP002789 Xanthomonas campestris pv. raphani 756C Total score: Z2.0 Cumulative Blast bit score: 97590	
6. : CP000050 Xanthomonas campestris pv. campestris str. 8004 Total score: 20.0 Cumulative Blast bit score: 93680	
8. : AM039952 Xanthomonas campestris pv. vesicatoria complete genome. Total score: 15.0 Cumulative Blast bit score: 69030	
9. : A Y442352 Aneurinibacillus thermoaerophilus strain L420-91T 5-layer glycan biosynthesis gene cluster 🛛 Total score: 8.0 Cumulative Blast b	it score: 15340
10, : ACPX01000172 Xanthomonas fuscans subsp. aurantifolii str. ICPB 11122 — Total score: 7.5 — Cumulative Blast bit score: 32040	
11. : ACPY01000143 Xanthomonas fuscans subsp. aurantifolii str. ICPB 10535 Total score: 7.5 Cumulative Blast bit score: 32030	
12. ; FP565176 Xanthomonas albilineans GPE PC73 complete genome. Total score: 7.5 Cumulative Blast bit score: 20520	

References

- 1. Fox E.M., Howlett B.J. (2008) Biosynthetic gene clusters for epipolythiodioxopiperazines in filamentous fungi. *Mycological Research* 112: 162-169.
- 2. Slot J.C., Rokas A. (2011) Horizontal transfer of large and highly toxic secondary metabolic gene cluster between fungi. *Current Biology* 21: 134-139.
- 3. F. Pojer, S.M. Li, L. Heide (2002) Molecular cloning and sequence analysis of the clorobiocin biosynthetic gene cluster: new insights into the biosynthesis of aminocoumarin antibiotics. *Microbiology* 148: 3901-3911.
- 4. Ryan K.S. (2011) Biosynthetic gene cluster for the cladoniamides, bis-indoles with a rearranged scaffold. *PLoS ONE* 6(8): e23694.
- Potnis N., Krasileva K., Chow V., Almeida N.F., Patil P.B., Ryan R.P., Sharlach M., Behlau F., Dow J.M., Momol M.T., White F.F., Preston J.F., Vinatzer B.A., Koebnik R., Setubal J.C., Norman D.J., Staskawicz B.J., Jones J.B. (2011) Comparative genomics reveals diversity among xanthomonads infecting tomato and pepper. *BMC Genomics* 12: 146.