

## Supplementary Information

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***Streptomyces natalensis* programmed cell death and morphological differentiation are dependent on oxidative stress**

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**Supplementary Table S1** – *S. natalensis* homologues.

**Supplementary Table S2** – Primers.

**Supplementary Figure S1** - Construction of *S. natalensis*  $\Delta catR$  strain by gene replacement.

**Supplementary Figure S2** – 2D-PAGE gels.

**Supplementary Figure S3** – SDS-PAGE.

22 **Table S1.** *S. natalensis* homologues of known *S. coelicolor* development related proteins.

Protein name	SNA_code	Identity	Coverage	e-value	<i>S. coelicolor</i>
					RefSeq
<b>BldA</b>	SNA_04225	93%	100%	4.00E-28	Y00209.1
<b>BldB</b>	SNA_08815	68%	93%	1.00E-23	NP_629849.1
<b>BldC</b>	SNA_03650	100%	100%	6.00E-44	NP_628272.1
<b>BldD</b>	SNA_13365	98%	100%	4.00E-113	NP_625769.1
<b>BldG</b>	SNA_00365	96%	100%	1.00E-73	NP_627747.1
<b>BldH (AdpA)</b>	SNA_06025	92%	79%	0	NP_627022.1
<b>BldKA</b>	SNA_34770	60%	99%	8.00E-146	NP_629262.1
<b>BldKB</b>	SNA_34775	49%	97%	0	NP_629263.1
<b>BldKC</b>	SNA_34780	70%	99%	5.00E-166	NP_629264.1
<b>BldKD</b>	SNA_34785	78%	99%	0	NP_629265.1
<b>BldKE</b>	SNA_34790	80%	93%	0	NP_733655.1
<b>BldM</b>	SNA_39180	100%	100%	7.00E-142	NP_628926.1
<b>BldN</b>	SNA_38050	96%	67%	7.00E-123	NP_627533.1
<b>WhiA</b>	SNA_10605	92%	99%	0	NP_626214.1
<b>WhiB</b>	SNA_03895	95%	98%	7.00E-56	NP_627256.1
<b>WhiD</b>	SNA_39175	97%	84%	4.00E-64	NP_628925.1
<b>WhiEI</b>	SNA_26210	57%	96%	2.00E-138	NP_629462.1
<b>WhiEII</b>	SNA_26205	72%	91%	9.00E-65	NP_629461.1
<b>WhiEIII</b>	SNA_26200	76%	99%	0	NP_629460.1
<b>WhiEIV</b>	SNA_26195	74%	96%	0	NP_629459.1
<b>WhiEV</b>	SNA_26190	52%	98%	9.00E-23	NP_629458.1
<b>WhiEVI</b>	SNA_26185	68%	91%	7.00E-69	NP_629457.1
<b>WhiEVII</b>	SNA_26180	72%	95%	2.00E-52	NP_629456.1
<b>WhiEVIII</b>	SNA_26175	77%	96%	0	NP_629463.1
<b>WhiG</b>	SNA_32585	90%	99%	0	NP_629755.1
<b>WhiH</b>	SNA_31720	76%	99%	7.00E-162	NP_629942.1
<b>WhiI</b>	SNA_30685	95%	99%	9.00E-140	NP_630140.1
<b>ChpA</b>	SNA_06310	50%	56%	3.00E-37	NP_733581.1
<b>ChpC</b>	SNA_01235	46%	99%	5.00E-38	NP_625949.1
<b>ChpD</b>	SNA_00490	67%	97%	2.00E-23	NP_626950.1
<b>ChpD2</b>	SNA_12255	77%	60%	1.00E-20	NP_626950.1

Protein name	SNA_code	Identity	Coverage	e-value	<i>S. coelicolor</i>
					RefSeq
<b>ChpE</b>	SNA_11245	64%	97%	6.00E-20	NP_626070.1
<b>ChpE2</b>	SNA_11250	67%	64%	1.00E-17	NP_626070.1
<b>ChpG</b>	SNA_00485	67%	49%	2.00E-16	NP_626939.1
<b>ChpG2</b>	SNA_05975	73%	48%	3.00E-15	NP_626939.1
<b>ChpH</b>	SNA_12040	86%	98%	8.00E-24	NP_625950.1
<b>RdIA</b>	SNA_27905	53%	71%	2.00E-25	NP_626951.1
<b>RamR</b>	SNA_20600	50%	98%	1.00E-50	NP_733711.1
<b>RamB</b>	SNA_20605	56%	96%	1.00E-174	NP_630759.1
<b>RamA</b>	SNA_20610	56%	92%	8.00E-147	NP_630758.1
<b>RamS</b>	SNA_20615	76%	97%	3.00E-17	NP_630757.1
<b>RamC</b>	SNA_20620	66%	99%	0	NP_630756.1
<b>DivIVA</b>	SNA_09545	84 %	54 %	4E-99	NP_626336
<b>FilP</b>	SNA_33665	92%	100 %	0	NP_629535
<b>FtsK</b>	SNA_11150	61 %	88 %	1E-152	NP_733622
<b>FtsW</b>	SNA_09505	79 %	94%	0	NP_626344
<b>FtsZ</b>	SNA_09520	86 %	100 %	0	NP_626341
<b>Mbl</b>	SNA_06725	97 %	100 %	0	NP_626848
<b>MreB</b>	SNA_07640	71 %	98 %	4E-170	NP_626694
<b>ParA</b>	SNA_02365	89 %	100 %	4E-173	NP_628072
<b>ParB</b>	SNA_02360	74 %	99 %	1E-149	NP_628073
<b>Scy</b>	SNA_33660	60 %	100 %	0	NP_629536
<b>SigF</b>	SNA_01565	92 %	94 %	8E-171	NP_628217
<b>Smc</b>	SNA_32725	82 %	86 %	0	NP_629712
<b>SsgA</b>	SNA_26640	49 %	97 %	4E-40	NP_630795
<b>SsgB</b>	SNA_00325	88 %	92 %	2E-159	NP_627756
<b>SspA</b>	SNA_27035	44 %	61 %	3E-3	NP_631483

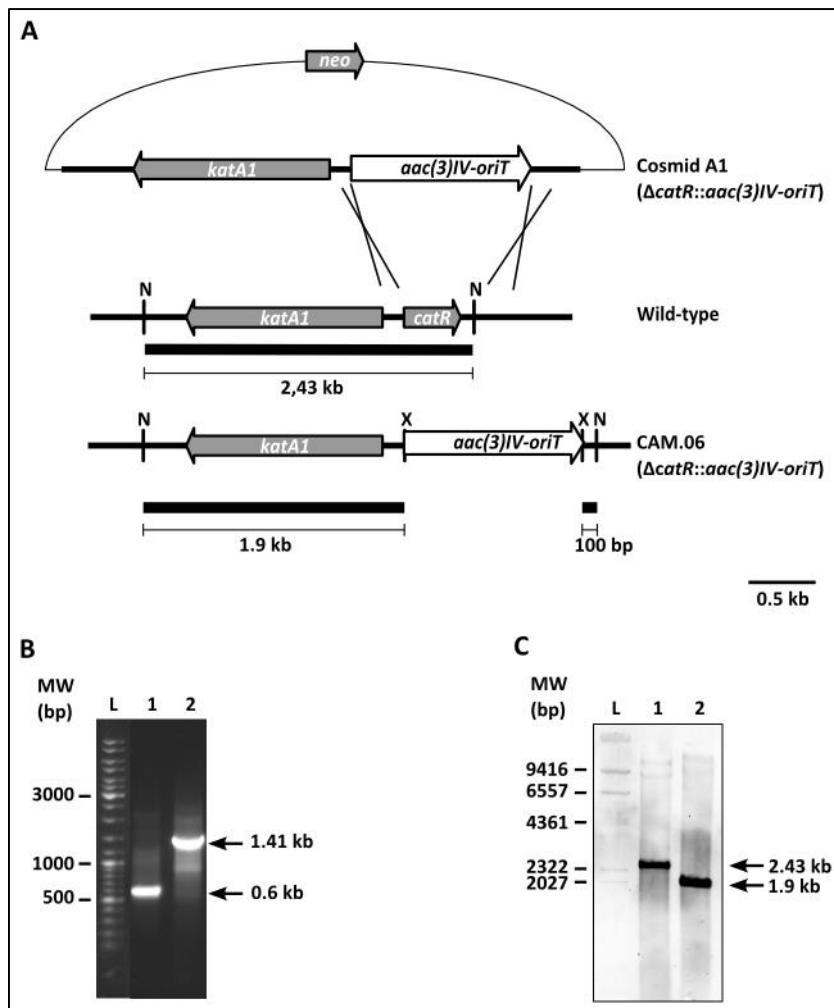
24 **Table S2.** Primers used in PCR and RT-qPCR.

Primer	Sequence	T.A. (°C)	Amplicon (bp)
D_catR_F	ATTCCGGGGATCCGTCGACCGTACCCTGGGCTGTATCTGACTGATAGGTGATTGGCATG	60	1462
D_catR_R	TGTAGGCTGGAGCTGCTCCCTGGGGGCGGGCCGGACGCGGGACGCCGACCGCGGTCA		
C_catR_F	GTGCGCCTCCTGAAGTGATC	60	584
C_catR_R	TCAGGCGTGGTGGCCGCT		
adpA_S	CCCGCGACCACCCATTGGA	65	105
adpA_AS	GGTGAGCACATCGCCGTCGTC		
bldD_S	CTGGAAGGCCGTGGTGGTGG	65	105
bldD_AS	AGCTCCTGGACCGGCACCC		
bldN_S	CACGGCCGAGGACCTCACCA	65	102
bldN_AS	ATGGTCACCAGCCAGGCGC		
chpC_S	CACCAGGGCGGGTCGGA	60	109
chpC_AS	TGGCCGGTACCGCGTAGC		
ramC_S	ATCCATATCTCGGCCTGCCTGG	60	102
ramC_AS	CCGGCTCGGCATGCATTG		
ramS_S	TCGACCTGCAGACGATGGAAC	60	99
ramS_AS	TCAGACTGCTGTCGCCGCA		
rdIA_S	GAGTAACCGCCACGCCAGCA	60	103
rdIA_AS	CCCTGCGGGCTCATGTCCG		
bfrA_S	CGGCTGGCCGAAGCTTGC	60	103
bfrA_AS	CGGCAGCCCTTCGAGGAAGA		
catR_S	CAACGCCACCCACGACCA	65	75
catR_AS	TCGCCCTGGAGATGGACG		
cpx_S	GAGCGGCTCCAGCAGGACT	60	106

<b>Primer</b>	<b>Sequence</b>	<b>T.A. (°C)</b>	<b>Amplicon (bp)</b>
<b>cpx_AS</b>	TCGCGGCTTGTTGACG		
<b>dpsA_S</b>	CACTGGAACGTCGTCGGCC	60	150
<b>dps_AS</b>	CTTGCGACCGTCGAGGC		
<b>katA1_S</b>	CCTGGTCGGTCTTGAAGTGGTA	60	80
<b>katA1_AS</b>	CTACGGCTCGCACACCTTC		
<b>katA3_S</b>	CACACCAAGGACGACGACTT	60	81
<b>katA3_AS</b>	CTCCACCAGACGCTTCTTCTC		
<b>16S_S</b>	CAGGCTAGAGTTCGGTAGG	65	117
<b>16S_AS</b>	CTCCTCAGCGTCAGTATCG		

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27 **Fig. S1 - Construction of *S. natalensis*  $\Delta catR$  strain by gene replacement.**

28 (A) Predicted restriction enzyme polymorphism caused by gene replacement. The *NcoI*-

29 *XbaI* restriction pattern before and after replacement is shown. The probe used for

30 southern hybridization is indicated by thick lines. N, *NcoI*; X, *XbaI*. (B) Confirmation of gene

31 disruption by PCR. A pair of primers, C\_catR\_F and C\_catR\_R, covering the deleted region

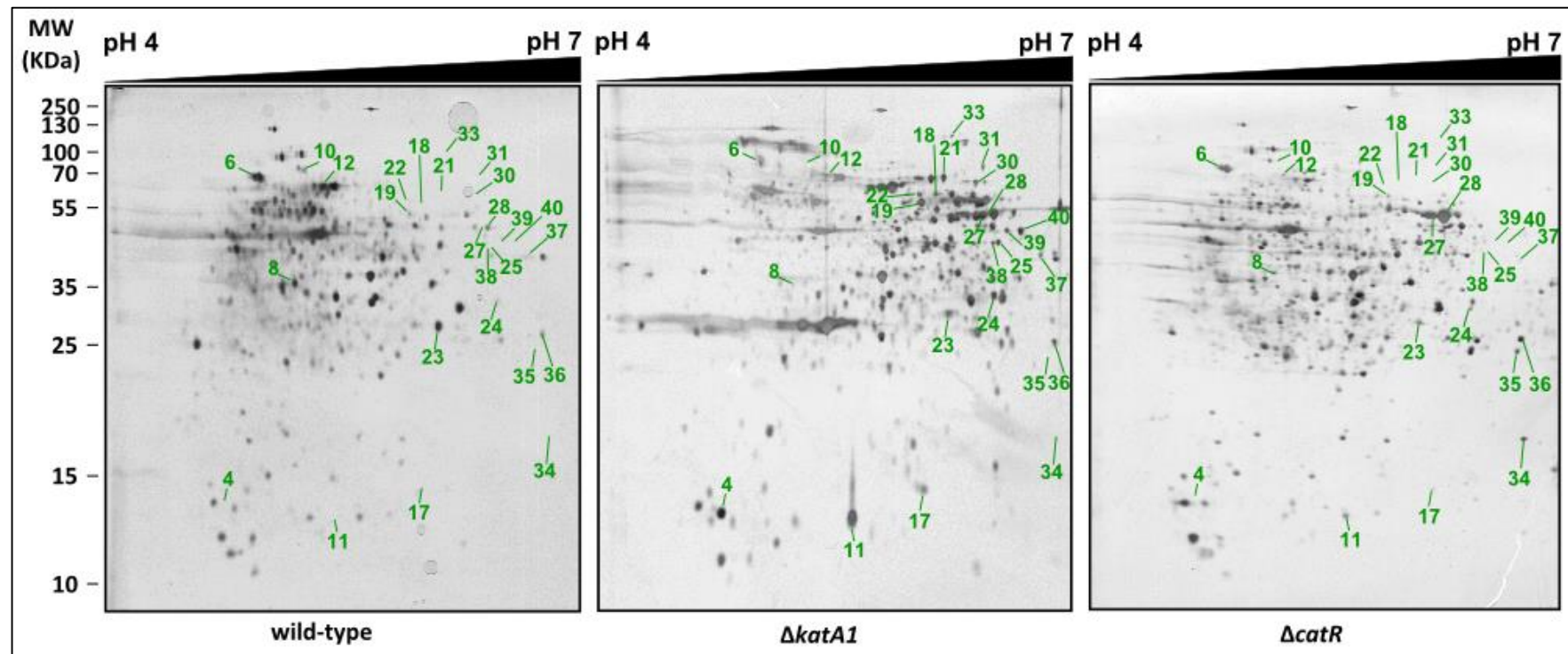
32 in the chromosome were to identify double crossover mutants. Lane L, GeneRuler™

33 (Thermo Scientific). (C) Confirmation of gene disruption by Southern blot of the *NcoI*-*XbaI*

34 digested chromosomal DNA of the wild type (lane 1), and  $\Delta catR$  (lane 2) strains. Lane L,

35 DIG-labelled DNA Molecular Weight Marker II (Roche).

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**Fig. S2 - Proteome analysis by 2D-PAGE.**

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The presented 2D gels are representative of biological duplicates (100  $\mu$ g of crude protein extracts per gel). The spots indicated in the

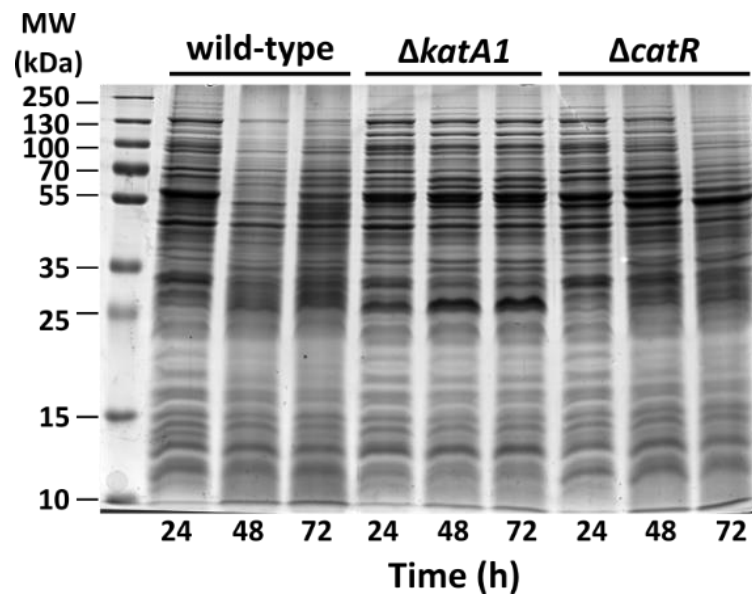
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figure correspond to the proteins identified by mass spectrometry (for SNA code correspondence to spot ID number see Table 1).

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**Fig. S3** - Coomassie coloration of SDS-PAGE with protein crude extracts (20  $\mu$ g of

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protein per lane) used as loading control for the nuclease activity assay (Fig. 4B).

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