

Figure S1 Morphological features of 58 endophytic strains (Red box: SH-1.2-R-15).

## Table S1 16S rRNA analysis of 58 endophytic strains.

NO.	Strains	Host plant	Parts	Closest sequences by BlastN	Percent	Genbank accession
					Identity	
1	SH-1.1-R-1	Dendrobium	Root	Bacillus sp. (in: Bacteria) strain 12D6	97.14%	MN784185
2	SH-1.1-R-2	officinale -	Root	Stenotrophomonas rhizophila strain PgBE58	99.72%	MN784186
3	SH-1.1-R-3	one year	Root	Bacillus sp. BS98	99.90%	MN784187
4	SH-1.1-R-4	old-1	Root	Streptomyces sp. strain SHP 1-2	99.80%	MN784188
5	SH-1.1-R-5		Root	Stenotrophomonas sp. strain H9	100.00%	MN784189
6	SH-1.1-R-6		Root	Streptomyces sp.SR3-82 gene	99.85%	MN784190
7	SH-1.1-R-7		Root	Bacillus thuringiensis strain ZLynn1000-39	97.59%	MN784191
8	SH-1.1-R-8		Root	Same as 5	-	-
9	SH-1.1-R-9		Root	Aerococcus viridans strain NBRC 12219	99.78%	MN784192
10	SH-1.1-L-1		Leaf	Bacillus cereus strain BC2	99.90%	MN784193
11	SH-1.1-L-2		Leaf	Bacillus thuringiensis strain GA-A07	99.90%	MN784194
12	SH-1.1-L-3		Leaf	Same as 58	-	-
13	SH-1.1-S-1		Stem	Stenotrophomonas sp. Strain FB14	99.93%	MN784195
14	SH-1.2-R-1	Dendrobium	Root	Streptomyces sp. NK04104	96.58%	MN784196
15	SH-1.2-R-2	officinale -	Root	Streptomyces sp. CdTB01	97.13%	MN784197
16	SH-1.2-R-3	one year	Root	Pseudomonas sp. BMS12	99.33%	MN784198
17	SH-1.2-R-4	old-2	Root	Streptomyces sp. g2b	99.79%	MN784199
18	SH-1.2-R-5		Root	Streptomyces sp. HF-2	96.82%	MN784200
19	SH-1.2-R-6		Root	Pseudomonas sp. Strain IAE244	99.92%	MN784201
20	SH-1.2-R-7		Root	Streptomyces sp. LCB 0297	97.94%	MN784202
21	SH-1.2-R-8		Root	Streptomyces sp. strain QJSt8	97.47%	MN784203
22	SH-1.2-R-9		Root	Same as 7	-	-
23	SH-1.2-R-10		Root	Same as 8	-	-
24	SH-1.2-R-11		Root	Same as 21	-	-
25	SH-1.2-R-12		Root	Bacillus cereus strain ULT15	99.60%	MN784204
26	SH-1.2-R-13		Root	Streptomyces sp. TM-74	97.51%	MN784205
27	SH-1.2-R-14		Root	Streptomyces fulvissimus strain DSM 40593	97.64%	MN784206
28	SH-1.2-R-15		Root	Streptomyces chartreusis strain ISP 5085	99.21%	MN784207
29	SH-1.2-R-16		Root	Streptomyces variabilis strain NRRL B-3984	97.52%	MN784208
30	SH-1.2-R-17		Root	Streptomyces sp. strain TM-A158	99.86%	MN784209
31	SH-1.2-R-18	]	Root	Bacillus thuringiensis strain GA-A07 chromosome	99.93%	MN784210
32	SH-1.2-R-19	1	Root	Same as 58	-	-
33	SH-1.2-L-1		Leaf	Micromonospora aurantiaca strain IMB16-201	99.92%	MN784211
34	SH-3.1-R-1	Dendrobium	Root	Bacillus sp. (in: Bacteria) strain 12D6	98.67%	MN784212
35	SH-3.1-R-2	officinale -	Root	Same as 36	-	-
36	SH-3.1-R-3	three years	Root	Streptomyces sp. GSENDO-0578	96.80%	MN784213
37	SH-3.1-R-4	old-1	Root	Same as 50	-	-
38	SH-3.1-R-5		Root	Streptomyces sp. 13-22	96.92%	MN784214
39	SH-3.1-R-6		Root	Streptomyces sp. MI02-7b	99.58%	MN784215
40	SH-3.1-R-7	1	Root	Streptomyces anulatus strain 174456	97.06%	MN784216
41	SH-3.1-R-8	1	Root	Same as 38	-	-
42	SH-3.1-R-9		Root	Streptomyces sp. DSM 40835 clone K12	99.70%	MN784217
43	SH-3.1-R-10		Root	Bacillus thuringiensis strain JW-1	99.81%	MN784218
44	SH-3.1-R-11	1	Root	Bacillus thuringiensis strain GA-A07	100.00%	MN784219

45	SH-3.1-R-12		Root	Same as 66	-	-
46	SH-3.1-R-13		Root	Bacillus sp. JAS24-2 chromosome	100.00%	MN784220
47	SH-3.1-R-14		Root	Streptomyces sp. strain MUSC11	98.46%	MN784221
48	SH-3.1-R-15		Root	Same as 66	-	-
49	SH-3.1-R-16		Root	Streptomyces sp. strain BOR09	93.69%	MN784222
50	SH-3.1-R-17		Root	Viridibacillus sp. strain TM-B117	99.69%	MN784223
51	SH-3.1-R-19		Root	Streptomyces sp. SR3-82	96.17%	MN784224
52	SH-3.1-R-20		Root	Bacillus thuringiensis strain GCU1	99.79%	MN784225
53	SH-3.1-R-21		Root	Streptomyces prunicolor NBRC 13075	99.80%	MN784226
54	SH-3.1-R-22		Root	Streptomyces sp. strain GDMCC 60254	96.51%	MN784227
55	SH-3.1-L-1		Leaf	Same as 59	-	-
56	SH-3.1-L-2		Leaf	Same as 65	-	-
57	SH-3.1-L-3		Leaf	Same as 60	-	-
58	SH-3.1-S-1		Stem	Streptomyces sp. strain GDMCC 60254	96.92%	MN784228
59	SH-3.1-S-2		Stem	Bacillus cereus strain NRRL B-23957	98.73%	MN784229
60	SH-3.1-S-3		Stem	Streptomyces sp. SM17	99.31%	MN784230
61	SH-3.1-S-4		Stem	Same as 65	-	-
62	SH-3.2-R-1	Dendrobium	Root	Bacillus sp. HT-Z74-B2	98.55%	MN784231
63	SH-3.2-R-2	officinale -	Root	Same as 50	-	-
64	SH-3.2-R-3	three years	Root	Same as 60	-	-
65	SH-3.2-R-4	old-2	Root	Bacillus cereus strain GE16	98.78%	MN784232
66	SH-3.2-R-5		Root	Streptomyces sp. GKU 867	100.00%	MN784233
67	SH-3.2-R-6		Root	Bacillus thuringiensis strain QZL38 chromosome	100.00%	MN784234
68	SH-3.2-R-7		Root	Streptomyces rochei 7434AN4 DNA	99.79%	MN784235
69	SH-3.2-R-8		Root	Streptomyces sp. Endophyte N2 chromosome	100.00%	MN784236
70	SH-3.2-R-9		Root	Bacillus cereus strain Sneb2000	100.00%	MN784237
71	SH-3.2-R-10		Root	Brevundimonas sp. strain 7002-176	100.00%	MN784238
72	SH-3.2-R-11		Root	Corynebacterium sp. strain ABYHD3-2	99.85%	MN784239
73	SH-3.2-R-12		Root	Streptomyces sp. TJ-27	97.98%	MN784240
74	SH-3.2-R-13		Root	Bacillus sp. AR4-2 chromosome	100.00%	MN784241
75	SH-3.2-L-1		Leaf	Bacillus thuringiensis strain GA-A07	100.00%	MN784242



Figure S2 Scatterplot of 75 cultured endophytic strains screened in triplicate at 25 or 50  $\mu$ g/mL in Hep3B2.1-7 cell viability assay. Data are shown normalized to percent max response of the high control (Medium only group). Each data point represents the mean and standard deviation of three replicates in 96 wells. The hit cutoff calculated as the average percent inhibition plus three times the standard deviation of the low control wells, is shown as the black line at 11.7%.

Table S2. Strains showed inhibition a	against Hep3B2.1-7	' cell more than the hi	t cutoff (>11.7%).
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No.	Endophytic strains	%Inhibition	No.	Endophytic strains	%Inhibition
2	SH-1.1-R-2	22.14±4.46	54	SH-3.1-R-22	35.74±2.31
4	SH-1.1-R-4	44.24±1.39	56	SH-3.1-L-2	18.64±1.16
16	SH-1.2-R-3	14.67±1.99	58	SH-3.1-S-1	14.87±3.17
18	SH-1.2-R-5	97.24±0.38	59	SH-3.1-S-2	47.80±0.99
19	SH-1.2-R-6	89.27±9.58	61	SH-3.1-S-4	12.00±1.38
24	SH-1.2-R-11	23.97±4.63	62	SH-3.2-R-1	33.77±2.22
27	SH-1.2-R-14	98.44±0.36	63	SH-3.2-R-2	41.40±1.71
28	SH-1.2-R-15	20.30±3.76	64	SH-3.2-R-3	31.27±4.01
31	SH-1.2-R-18	34.47±2.74	65	SH-3.2-R-4	12.30±3.31
33	SH-1.2-L-1	14.47±1.28	66	SH-3.2-R-5	47.07±1.86
42	SH-3.1-R-9	23.14±1.77	67	SH-3.2-R-6	62.20±6.76
43	SH-3.1-R-10	26.37±2.21	68	SH-3.2-R-7	98.67±0.62
44	SH-3.1-R-11	12.17±4.47	69	SH-3.2-R-8	89.50±2.46
47	SH-3.1-R-14	45.34±1.71	73	SH-3.2-R-12	49.97±1.41



Figure S3 Scatterplot of 75 cultured endophytic strains screened in triplicate at 25 or 50  $\mu$ g/mL in *S. aureus* inhibitory assay. Data are shown normalized to percent max response of the high control (Bacterial treated with 100  $\mu$ g/mL Ampicillin). Each data point represents the mean and standard deviation of three replicates in 96 wells. The hit cutoff calculated as the average percent inhibition plus three times the standard deviation of the low control wells, is shown as the black line at 36.2%.

Table S3 Strains showed inhibition against S	S. aureus more than the hit cutoff	(>36.2%).
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No.	Endophytic strains	%Inhibition	No.	Endophytic strains	%Inhibition
12	SH-1.1-L-3	41.86±3.50	37	SH-3.1-R-4	51.50±1.96
13	SH-1.1-S-1	50.13±5.59	44	SH-3.1-R-11	41.81±8.64
14	SH-1.2-R-1	36.70±6.74	48	SH-3.1-R-15	40.45±21.15
15	SH-1.2-R-2	40.17±5.94	50	SH-3.1-R-17	48.24±15.86
18	SH-1.2-R-5	95.12±3.34	52	SH-3.1-R-20	47.21±7.18
23	SH-1.2-R-10	54.04±7.21	56	SH-3.1-L-2	46.37±3.41
24	SH-1.2-R-11	72.75±6.14	57	SH-3.1-L-3	38.59±3.27
26	SH-1.2-R-13	51.38±3.27	68	SH-3.2-R-7	100.00±0.24
27	SH-1.2-R-14	99.66±0.58	69	SH-3.2-R-8	99.32±0.53
28	SH-1.2-R-15	100.12±0.09	70	SH-3.2-R-9	51.01±6.67
29	SH-1.2-R-16	40.21±5.97			



Figure S4 Scatterplot of 75 samples screened in triplicate at 25 or 50  $\mu$ g/mL in *E. coli* inhibitory assay. Data are shown normalized to percent max response of the high control (Bacterial treated with 100  $\mu$ g/mL Ampicillin). Each data point represents the mean and standard deviation of three replicates in 96 wells. The hit cutoff calculated as the average percent inhibition plus three times the standard deviation of the low control wells, is shown as the black line at 26.4%.

Table S4 Strains showed inhibition against E. coli more than the hit cutoff (>26.4%).

No.	Endophytic strains	%Inhibition
59	SH-3.1-S-2	31.49±6.86



Scheme S1 Work-up scheme for the Streptomyces sp. SH-1.2-R-15.



Figure S5. (+) and (–)-ESI-MS of compound 1.



Figure S6. HRESI-MS spectrum of compound 1.



Figure S7. <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound 1.



Figure S8. <sup>1</sup>H NMR (500 MHz, CDCl<sub>3</sub>) spectrum of compound **1**.



Figure S9. <sup>13</sup>C NMR (125 MHz, CDCl<sub>3</sub>) spectrum of compound **1**.



Figure S10. <sup>1</sup>H-<sup>1</sup>H COSY (500 MHz, CDCl<sub>3</sub>) spectrum of compound **1**.



Figure S11. HSQC (500 MHz, CDCl<sub>3</sub>) spectrum of compound 1.



Figure S12. HMBC (500 MHz,  $CDCI_3$ ) spectrum of compound 1.



Figure S13. ROESY (500 MHz, CDCl<sub>3</sub>) spectrum of compound 1.



Figure S14. (+) and (–)-ESI-MS of compound 2.



Figure S15. HRESI-MS spectrum of compound 2.



**Figure S16.** <sup>1</sup>H NMR (500 MHz, DMSO- $d_6$ ) spectrum of compound **2**.



Figure S17. <sup>1</sup>H NMR (500 MHz, CDCl<sub>3</sub>) spectrum of compound 2.



Figure S18. <sup>13</sup>C NMR (125 MHz, CDCl<sub>3</sub>) spectrum of compound 2.



**Figure S19.** <sup>1</sup>H-<sup>1</sup>H COSY (500 MHz, CDCl<sub>3</sub>) spectrum of compound **2**.



Figure S20. HSQC (500 MHz,  $CDCI_3$ ) spectrum of compound 2.



Figure S21. HMBC (500 MHz, CDCl<sub>3</sub>) spectrum of compound 2.



Figure S22. ROESY (500 MHz,  $CDCI_3$ ) spectrum of compound 2.



Figure S23. (+) and (–)-ESI-MS of compound 3.



Figure S24. HRESI-MS spectrum of compound 3.



Figure S25. <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound **3**.



Figure S26. <sup>13</sup>C NMR (125 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound **3**.



**Figure S27.** <sup>1</sup>H-<sup>1</sup>H COSY (500 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound **3**.



Figure S28. HSQC (500 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound 3.



Figure S29. HMBC (500 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound 3.



Figure S30. (+) and (–)-ESI-MS of compound 4.



Figure S31. HRESI-MS spectrum of compound 4.



Figure S32. <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound **4**.



Figure S33. <sup>13</sup>C NMR (125 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound **4**.

Print of window 80: Current Mass Spectra Data File : D:\DATA\Wang Group\Zhaohuimin\ZHAOHUIMIN 2019-06-25 15-06-12\dj60-3lh12P2.D Sample Name : dj60-31h12P2

\_\_\_\_\_ Acq. Operator : SYSTEM Seq. Line : 2 Acq. Instrument : LCMS-Q Location : P1-F2 Injection Date : 6/25/2019 3:19:08 PM Inj: 1 Inj Volume : 2.000 µl Different Inj Volume from Sample Entry! Actual Inj Volume : 1.000 µl : D:\user1\data\ZHAOHUIMIN\ZHAOHUIMIN 2019-06-25 15-06-12\LC-MS-general.M Acq. Method Last changed : 5/27/2019 9:17:15 AM by SYSTEM Analysis Method : D:\DATA\Wang Group\Zhaohuimin\ZHAOHUIMIN 2019-06-25 15-06-12\LC-MS-general.M ( Sequence Method)

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Figure S34. (+) and (–)-ESI-MS of compound 5.



Figure S35. HRESI-MS spectrum of compound 5.



Figure S36. <sup>1</sup>H NMR (500 MHz, CDOD<sub>3</sub>) spectrum of compound 5.



Figure S37. <sup>13</sup>C NMR (125 MHz, CDOD<sub>3</sub>) spectrum of compound **5**.

Print of window 80: Current Mass Spectra Data File : D:\DATA\Wang Group\Zhaohuimin\ZHAOHUIMIN 2019-06-25 15-06-12\dj60-31h12P1.D Sample Name : dj60-31h12P1

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Different Inj Volume from Sample Entry! Actual Inj Volume : 1.000 µl
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Last changed
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Figure S38. (+) and (–)-ESI-MS of compound 6.



Figure S39. HRESI-MS spectrum of compound 6.



Figure S40. <sup>1</sup>H NMR (500 MHz, CDOD<sub>3</sub>) spectrum of compound **6**.



Figure S41. <sup>13</sup>C NMR (125 MHz, CDOD<sub>3</sub>) spectrum of compound 6.



Figure S42. (+) and (–)-ESI-MS of compound 7.



Figure S43. HRESI-MS spectrum of compound 7.



**Figure S44.** <sup>1</sup>H NMR (500 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound **7**.



Figure S45. <sup>13</sup>C NMR (125 MHz, DMSO-*d*<sub>6</sub>) spectrum of compound **7**.