Supporting Information

Process Development and Scale-up Total Synthesis of Largazole, a Potent Class I Histone Deacetylase Inhibitor

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Figure S1. Analytical HPLC profile of compound 2



Figure S2. Analytical HPLC profile of compound 8



Figure S3. Analytical HPLC profile of compound 5



Figure S4. Analytical HPLC profile of compound 3



Figure S5. Analytical HPLC profile of compound 1 (fraction 1)



Figure S6. Analytical HPLC profile of compound 1 (fraction 2)

Figure S7. Picture of product largazole



Figure S8. ¹H NMR Spectrum of 2 in CDCl₃ (400 MHz) at 25 °C



S9

Figure S9. ¹³C NMR Spectrum of 2 in CDCl₃ (100 MHz) at 25 °C





S10



Figure S10. ¹H NMR Spectrum of 18 in CDCl₃ (400 MHz) at 25 °C







Figure S12. ¹H NMR Spectrum of 20 in CDCl₃ (400 MHz) at 25 °C



Figure S13. ¹³C NMR Spectrum of 20 in CDCl₃ (100 MHz) at 25 °C

Figure S14. ¹H NMR Spectrum of 7 in D2O (400 MHz) at 25 °C









Figure S16. ¹H NMR Spectrum of 23 in CDCl₃ (400 MHz) at 25 °C



Figure S17. ¹³C NMR Spectrum of 23 in CDCl₃ (100 MHz) at 25 $^{\circ}$ C



Figure S18. ¹H NMR Spectrum of 24 in CDCl₃ (500 MHz) at 25 °C



Figure S19. ¹³C NMR Spectrum of 24 in CDCl₃ (125 MHz) at 25 °C



Figure S20. ¹H NMR Spectrum of 8 in CDCl₃ (400 MHz) at 25 °C



Figure S21. ¹³C NMR Spectrum of 8 in CDCl₃ (100 MHz) at 25 °C



Figure S22. ¹H NMR Spectrum of **5** in CDCl₃ (400 MHz) at 25 °C



Figure S23. ¹³C NMR Spectrum of **5** in CDCl₃ (100 MHz) at 25 °C



Figure S24. ¹H NMR Spectrum of (*RS*)-9 in CDCl₃ (400 MHz) at 25 °C



Figure S25. ¹³C NMR Spectrum of (*RS*)-9 in CDCl₃ (100 MHz) at 25 $^{\circ}$ C



Figure S26. ¹H NMR Spectrum of (*R*)-9 in CDCl₃ (400 MHz) at 25 °C



Figure S27. ¹³C NMR Spectrum of (R)-9 in CDCl₃ (100 MHz) at 25 °C



Figure S28. ¹H NMR Spectrum of (S)-31 in CDCl₃ (400 MHz) at 25 °C



Figure S29. ¹³C NMR Spectrum of (S)-31 in CDCl₃ (100 MHz) at 25 $^{\circ}$ C



Figure S30. ¹H NMR Spectrum of (*S*)-9 in CDCl₃ (400 MHz) at 25 °C



Figure S31. ¹³C NMR Spectrum of (S)-9 in CDCl₃ (100 MHz) at 25 $^{\circ}$ C



Figure S32. ¹H NMR Spectrum of 33 in CDCl₃ (400 MHz) at 25°C



Figure S33. ¹³C NMR Spectrum of 33 in CDCl₃ (100 MHz) at 25 °C



Figure S34. ¹H NMR Spectrum of 6 in CDCl₃ (400 MHz) at 25 °C



Figure S35. 13 C NMR Spectrum of 6 in CDCl₃ (100 MHz) at 25 °C



Figure S36. ¹H NMR Spectrum of **4** in CDCl₃ (400 MHz) at 25 °C



Figure S37. ¹³C NMR Spectrum of 4 in CDCl₃ (100 MHz) at 25 °C



Figure S38. ¹H NMR Spectrum of 3 in CDCl₃ (400 MHz) at 25 °C



Figure S39. ¹³C NMR Spectrum of 3 in CDCl₃ (100 MHz) at 25 °C







