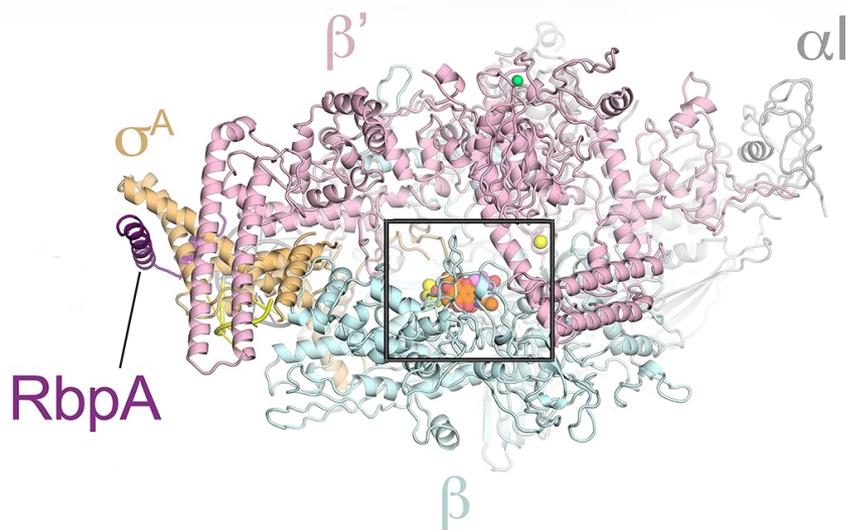


Searching for New Ways to Fight Bacterial Resistance



The image shows the overall structural basis for Kang A inhibition of rifamycin-resistant bacteria. The rifamycin scaffold is colored orange in the center of the RNA polymerase complex.

J. Peek, M. Lilic, D. Montiel, A. Milshteyn, I. Woodworth, J. Biggins, M. A. Ternei, P. Y. Calle, M. Danziger, T. Warrior, K. Saito, N. Braffman, A. Fay, M. S. Glickman, S. A. Darst, E. A. Campbell, S. F. Brady. *Nat. Commun.* **9**, 4147 (2018).

Work was performed in part at Brookhaven National Laboratory

Scientific Achievement

Scientists discovered that soil microbiomes could offer new analogues for clinically-used antibiotics.

Significance and Impact

The effectiveness of some antibiotics are threatened by the increasing resistance of bacteria; this study explores a new route to potential replacements for these antibiotics.

Research Details

- Surveying soil metagenomes helped to identify different rifamycin antibiotics, rifamycin congeners (kanglemycins, Kang), with potent activity against the most common resistance.
- Functional and mechanistic studies of the antibiotic effects on RNA polymerase, the target enzyme revealed a different inhibition mechanism.
- The x-ray crystallographic data were taken at NSLS-II AMX beamline and at APS beamline 24-ID-C.