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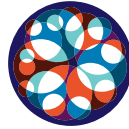
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The cover shows a metaphorical representation of the anti-CRISPR AcrIIA6, represented as handcuffs, sequestering two *Streptococcus thermophilus* CRISPR1-Cas9 (StlCas9) molecules at a time and preventing conformational changes associated with DNA recognition and binding. In the absence of AcrIIA6, StlCas9 tightly binds to its target DNA, and can proceed to target cleavage. This cover artwork has been made by Beata Edyta Mierzwa ([www.BeataScienceArt.com](http://www.BeataScienceArt.com)).

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