

# The CopC Protein from *Pseudomonas syringae*: Intermolecular Transfer of Copper Occurs from Both the Copper(I) and Copper(II) Sites

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The CopC protein from *Pseudomonas syringae* pathovar *tomato* is expressed as one of four proteins encoded by the operon CopABCD that is responsible for copper resistance.<sup>1</sup> It is a small soluble molecule (10.5 kDa) with a  $\beta$ -barrel structure and features two distinct copper binding sites (see Figure<sup>2</sup>) which are highly specific for Cu<sup>I</sup> ( $K_D \sim 10^{-13}$ ) and Cu<sup>II</sup> ( $K_D < 10^{-15}$ ), respectively. These dissociation constants were estimated via ligand competition experiments monitored by electronic spectral and fluorescence probes.

The chemistries of the two copper sites are inter-dependent. When the Cu<sup>II</sup> site is empty, the Cu<sup>I</sup> ion is oxidized by air but when both sites are occupied, the molecule is stable in air. The availability of an unoccupied site of higher affinity induces intermolecular transfer of *either* Cu<sup>I</sup> or Cu<sup>II</sup> while maintaining free copper ion concentrations in solution at sub-*pico* molar levels.

This intriguing copper chemistry is consistent with the proposed role of CopC as a copper carrier in the oxidizing periplasmic space. These properties would allow it to exchange *either* Cu<sup>I</sup> or Cu<sup>II</sup> with its putative partners CopA, CopB and CopD, contrasting with the role of the Cu<sup>I</sup> (only) chaperones found in the reducing cytoplasm.

1. Cha, J. S.; Cooksey, D. A. *PNAS USA* **1991**, 88, 8915-9; Cooksey, D. A. *FEMS Microbiol. Rev* **1994**, 14, 381-6.

2. Arnesano F., Banci L., Bertini I. and Thompsett A. R., *Structure*, 2002, 10, 1337; Arnesano F., Banci L., Bertini I., Mangani S., and Thompsett A. R., *PNAS USA*, 2003, 100, 3814.

