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Electrophile promiscuity of KdoS and AroS phosphoenolpyruvate (PEP) dependent aldolases in aldol addition reactions

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PURPOSE OF THE ABSTRACT

The formation of carbon–carbon bonds is strategic reaction in organic synthesis as it allows for the construction of the carbon scaffolds of organic molecules to generate more complex compounds from simpler ones¹. 2-keto-3-deoxyoctulosonic acid 8-phosphate synthase (KdoS, EC 2.5.1.55) and 3-deoxy-D-arabino-heptulosonic acid 7-phosphate synthase (AroS, EC 2.5.1.54) catalyze the irreversible addition of phosphoenolpyruvate (PEP) to D-arabinose 5-phosphate (Ara5P) and D-erythrose-4-phosphate (Ery4P) to yield 3-deoxy-D-manno-octulosonate-8-phosphate (KDO-8-P) and 3-deoxy-D-arabino-heptulosonic acid 7-phosphate (DAHP), respectively.

In this communication, metagenomic libraries of KdoS and AroS were screened to identify promising candidates capable of catalyzing the promiscuous aldol addition of phosphoenolpyruvate to aliphatic aldehydes (e.g. 2-(benzyloxy)acetaldehyde, 2-phenylacetaldehyde and 3-phenylpropanal)³ (Scheme 1).

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FIGURES

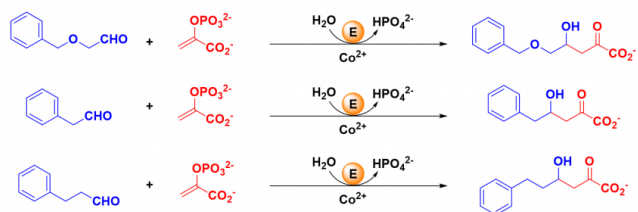


FIGURE 1

Scheme 1.

KdoS and AroS homologues from Prozomix panels catalyzed promiscuous aldol addition of phosphoenolpyruvate to aliphatic aldehydes.

FIGURE 2

KEYWORDS

PEP-Dependent Synthase | Enzyme Promiscuity | Aldol addition | Biocatalysis

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