## D-Sedoheptulose Efficient Production of Rare Sugar -

NAGASE & CO. targeted the D-sedoheptulose produced by *Streptomyces* and developed technology to mass produce it using N-STePP<sup>™</sup>

## [What is D-Sedoheptulose?]

D-Sedoheptulose (sedoheptulose) is a rare naturally-occurring sugar. While glucose is a six-carbon monosaccharide and xylose is a five-carbon monosaccharide, sedoheptulose is unique in being a seven-carbon monosaccharide, which is very rare in nature (Figure 1). Sedoheptulose was discovered in *Sedum spectabile* in 1917<sup>1</sup>, yet due to its rarity almost no research was carried out on its bioactive properties. It is an interesting question whether or not there is a causal relationship between *Sedum spectabile's* desiccation tolerance and the ample amount of sedoheptulose it contains.

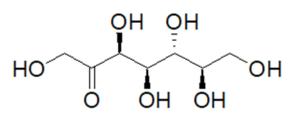
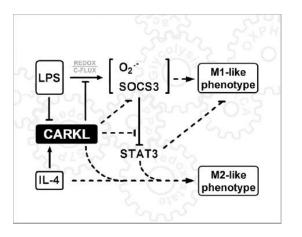


Figure 1 D-Sedoheptulose



Sedoheptulose 7-phosphate, an organophosphate of sedoheptulose, is an intermediate in the pentose phosphate pathway, part of the central metabolism of organisms, and is also an intermediate in the Calvin-Benson pathway, a carbon fixation pathway in photosynthesis, so it fulfills an important role in the metabolism of organisms.

Recently, it was found that a sedoheptulose kinase (CARKL) that catalyzes the phosphorylation of sedoheptulose can control the inflammatory immune response through metabolic control (Figure 2)<sup>2</sup>. These results corroborate the existence of a mutually dependent relationship between metabolism and immunity, and further research may show what role sedoheptulose plays in this relationship.

Figure 2 CARKL Regulation Cell Metabolism (2012) 15:813-826 Figure 7.

## [Production of Sedoheptulose by Streptomyces]

It has long been known that *Streptomyces* produces sedoheptulose, but the amount was minimal<sup>3</sup>. And while sedoheptulose is anticipated to have uses in treating and preventing inflammation<sup>4</sup>, or in supplements and health foods<sup>5</sup>, it has not yet been put to practical use. The Nagase BIO-INNOVATION CENTER, while developing technology to produce materials using *Streptomyces* as a host, discovered that sedoheptulose production could be dramatically increased through metabolic engineering<sup>6</sup>. By developing this technology, the Nagase BIO-INNOVATION CENTER was able to accelerate research into the functions of sedoheptulose and contribute to studies for its practical applications.

## [References]

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- 2. Haschemi A, et al. The Sedoheptulose Knase CARKL Directs Macrophage Polarization through Control of Glucose Metabolism. Cell Metabolism (2012) 15:813-826
- 3. Okuda T, et al. Accumulation of sedoheptulose by Streptomyces. J. Biochem. (1963) 54:107-108
- 4. WO2014147214
- 5. WO2014147213
- 6. WO2019208747

The NAGASE BIO-INNOVATION CENTER is committed to developing processes for efficiently producing a wide range of compounds with proprietary fermentation technologies.

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