

## 1 Nomenclature

### EC number

2.5.1.80

### Systematic name

dimethylallyl-diphosphate:L-tryptophan 7-dimethylallyltransferase

### Recommended name

7-dimethylallyltryptophan synthase

### Synonyms

7-DMATS <1,2> [1,3]

dimethylallyltryptophan synthase <1> [3]

## 2 Source Organism

<1> *Aspergillus fumigatus* [3]

<2> *Aspergillus fumigatus* (UNIPROT accession number: A8JY03) [1,2]

## 3 Reaction and Specificity

### Catalyzed reaction

dimethylallyl diphosphate + L-tryptophan = diphosphate + 7-(3-methylbut-2-enyl)-L-tryptophan

### Reaction type

alkenyl group transfer

### Substrates and products

**S** dimethylallyl diphosphate + 1-methyl-L-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 1-methyl-7-(3-methylbut-2-enyl)-L-tryptophan (<2> 35.8% of the yield with L-tryptophan [1])

**S** dimethylallyl diphosphate + 11-methyl-DL-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 11-methyl-DL-7-( $\gamma,\gamma$ -dimethylallyl)tryptophan (<2> 19.1% of the yield with L-tryptophan [1])

**S** dimethylallyl diphosphate + 11-methyl-DL-tryptophan <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated in-

dole derivatives. Yield of diprenylated product: 33.5% (FgaPT2 followed by 7-DMATS), 4.8% (7-DMATS followed by FgaPT2) [3] (Reversibility: ?) [3]

**P** ?

**S** dimethylallyl diphosphate + 4-methyl-DL-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 4-methyl-7-(3-methylbut-2-enyl)-DL-tryptophan (<2> 89.4% of the yield with L-tryptophan [1])

**S** dimethylallyl diphosphate + 5-bromo-DL-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 5-bromo-7-(3-methylbut-2-enyl)-DL-tryptophan (<2> 7.2% of the yield with L-tryptophan [1])

**S** dimethylallyl diphosphate + 5-fluoro-L-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 5-fluoro-7-(3-methylbut-2-enyl)-L-tryptophan (<2> 10.7% of the yield with L-tryptophan [1])

**S** dimethylallyl diphosphate + 5-hydroxy-L-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 5-hydroxy-7-(3-methylbut-2-enyl)-L-tryptophan (<2> 26.7% of the yield with L-tryptophan [1])

**S** dimethylallyl diphosphate + 5-methoxy-DL-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 5-methoxy-7-(3-methylbut-2-enyl)-DL-tryptophan (<2> 13.3% of the yield with L-tryptophan [1])

**S** dimethylallyl diphosphate + 5-methoxy-DL-tryptophan <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 10.3% (FgaPT2 followed by 7-DMATS), 3.5% (7-DMATS followed by FgaPT2) [3] (Reversibility: ?) [3]

**P** ?

**S** dimethylallyl diphosphate + 5-methyl-DL-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 5-methyl-7-(3-methylbut-2-enyl)-DL-tryptophan (<2> 74.8% of the yield with L-tryptophan [1])

**S** dimethylallyl diphosphate + 5-methyl-DL-tryptophan <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 24.3% (FgaPT2 followed by 7-DMATS), 0.9% (7-DMATS followed by FgaPT2) [3] (Reversibility: ?) [3]

**P** ?

**S** dimethylallyl diphosphate + 6-fluoro-DL-tryptophan <2> (Reversibility: ?) [1]

**P** diphosphate + 6-fluoro-7-(3-methylbut-2-enyl)-DL-tryptophan (<2> 33.7% of the yield with L-tryptophan [1])

- S** dimethylallyl diphosphate + 6-methyl-DL-tryptophan <2> (Reversibility: ?) [1]
- P** diphosphate + 6-methyl-7-(3-methylbut-2-enyl)-DL-tryptophan (<2> 19.8% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + 6-methyl-DL-tryptophan <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 2.6% (FgaPT2 followed by 7-DMATS), 1.5% (7-DMATS followed by FgaPT2) [3]) (Reversibility: ?) [3]
- P** ?
- S** dimethylallyl diphosphate + 7-methyl-DL-tryptophan <2> (Reversibility: ?) [1]
- P** diphosphate + 7-(3-methylbut-2-enyl)-DL-tryptophan + ? (<2> 0.3% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + D-tryptophan <2> (Reversibility: ?) [1,2]
- P** diphosphate + D-7-(3-methylbut-2-enyl)-tryptophan (<2> 11.8% of the yield with L-tryptophan [1]; <2> 15.5% of the activity with L-tryptophan [2])
- S** dimethylallyl diphosphate + D-tryptophan <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 4.3% (FgaPT2 followed by 7-DMATS), 2.6% (7-DMATS followed by FgaPT2) [3]) (Reversibility: ?) [3]
- P** ?
- S** dimethylallyl diphosphate + DL-indole-3-lactic acid <2> (Reversibility: ?) [1]
- P** diphosphate + 7-(3-methylbut-2-enyl)-DL-indole-3-lactic acid (<2> 4.3% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + DL-indole-3-lactic acid <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 35.7% (FgaPT2 followed by 7-DMATS), 16.7% (7-DMATS followed by FgaPT2) [3]) (Reversibility: ?) [3]
- P** ?
- S** dimethylallyl diphosphate + L-Trp-Gly <2> (Reversibility: ?) [2]
- P** diphosphate + L-7-(3-methylbut-2-enyl)-Trp-Gly (<2> 10.9% of the activity with L-tryptophan [2])
- S** dimethylallyl diphosphate + L-abrine <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 92.7% (FgaPT2 followed by 7-DMATS), 11.3% (7-DMATS followed by FgaPT2) [3]) (Reversibility: ?) [3]
- P** ?
- S** dimethylallyl diphosphate + L- $\beta$ -homotryptophan <2> (Reversibility: ?) [1]
- P** diphosphate + 7-(3-methylbut-2-enyl)-L- $\beta$ -homotryptophan (<2> 28.2% of the yield with L-tryptophan [1])

- S** dimethylallyl diphosphate + L- $\beta$ -homotryptophan <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 31.5% (FgaPT2 followed by 7-DMATS), 6.6% (7-DMATS followed by FgaPT2) [3]) (Reversibility: ?) [3]
- P** ?
- S** dimethylallyl diphosphate + L-tryptophan <2> (Reversibility: ?) [1,2]
- P** diphosphate + L-7-(3-methylbut-2-enyl)-tryptophan
- S** dimethylallyl diphosphate + L-tryptophan <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 89.7% (FgaPT2 followed by 7-DMATS), 20% (7-DMATS followed by FgaPT2) [3]) (Reversibility: ?) [3]
- P** ?
- S** dimethylallyl diphosphate + L-tryptophan hydroxamate <2> (Reversibility: ?) [1]
- P** diphosphate + L-7-( $\gamma,\gamma$ -dimethylallyl)tryptophan hydroxamate (<2> 4.6% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + L-tryptophan methyl ester <2> (Reversibility: ?) [1]
- P** diphosphate + 7-(3-methylbut-2-enyl)-L-tryptophan methyl ester (<2> 7.2% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + N-acetyl-DL-tryptophan <2> (Reversibility: ?) [1]
- P** diphosphate + N-acetyl-DL-7-( $\gamma,\gamma$ -dimethylallyl)tryptophan (<2> 7.7% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + N $^{\alpha}$ -methyl-L-tryptophan <2> (<2> i.e. L-abrine [1]) (Reversibility: ?) [1]
- P** diphosphate + N $^{\alpha}$ -methyl-L-7-( $\gamma,\gamma$ -dimethylallyl)tryptophan (<2> 82.2% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + cyclo-L-Trp-Gly <2> (Reversibility: ?) [2]
- P** diphosphate + cyclo-L-7-(3-methylbut-2-enyl)-Trp-Gly (<2> 1.8% of the activity with L-tryptophan [2])
- S** dimethylallyl diphosphate + indole-3-acetic acid <2> (Reversibility: ?) [1]
- P** diphosphate + 7-(3-methylbut-2-enyl)-indole-3-acetic acid (<2> 10.8% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + indole-3-butyric acid <2> (Reversibility: ?) [1]
- P** diphosphate + 7-(3-methylbut-2-enyl)-indole-3-butyric acid (<2> 9.1% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + indole-3-propionic acid <2> (Reversibility: ?) [1]
- P** diphosphate + 7-(3-methylbut-2-enyl)-indole-3-propionic acid (<2> 8.5% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + indole-3-pyruvic acid <2> (Reversibility: ?) [1]

- P** diphosphate + 7-(3-methylbut-2-enyl)-indole-3-pyruvic acid (<2> 4.8% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + tryptamine <2> (Reversibility: ?) [1]
- P** diphosphate + 7-(3-methylbut-2-enyl)-tryptamine (<2> 7.5% of the yield with L-tryptophan [1])
- S** dimethylallyl diphosphate + tryptamine <1> (<1> tandem incubation of dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) leads to the production of diprenylated indole derivatives. Yield of diprenylated product: 11% (FgaPT2 followed by 7-DMATS), 6.3% (7-DMATS followed by FgaPT2) [3]) (Reversibility: ?) [3]
- P** ?
- S** Additional information <2> (<2> no substrate: geranyl diphosphate, L-phenylalanine, L-tyrosine [2]) (Reversibility: ?) [2]
- P** ?

**Metals, ions**

Ca<sup>2+</sup> <2> (<2> slight stimulation [2]) [2]

Additional information <2> (<2> addition of divalent cations is not required [2]) [2]

**K<sub>m</sub>-Value (mM)**

0.067 <2> (dimethylallyl diphosphate, <2> pH 7.5, 37°C [2]) [2]

0.137 <2> (L-tryptophan, <2> pH 7.5, 37°C [2]) [2]

**4 Enzyme Structure****Molecular weight**

65000 <2> (<2> gel filtration [1]) [1]

**Subunits**

? <2> (<2> x \* 53000, calculated [2]) [2]

monomer <2> (<2> 1 \* 50000, SDS-PAGE, 1 \* 54000, calculated, His-tagged recombinant protein [1]) [1]

**5 Isolation/Preparation/Mutation/Application****Purification**

<2> (recombinant protein) [2]

**Cloning**

<1> (expressed as a recombinant protein) [3]

<2> (expression in Escherichia coli, His-tagged protein) [1,2]

**Application**

biotechnology <1> (<1> the combination of the two dimethylallyltryptophan synthases FgaPT2 and 7-DMATS (EC 2.5.1.34 and EC 2.5.1.80) can be successfully used for chemoenzymatic synthesis of the diprenylated derivatives. The

potential of recombinant enzymes from secondary metabolite biosynthesis as promising tools for the production of designed compounds is demonstrated [3]) [3]

## 6 Stability

### General stability information

<2>, 37°C, absence of substrate, 16 h, 80% residual activity [1]

## References

- [1] Kremer, A.; Li, S.M.: Potential of a 7-dimethylallyltryptophan synthase as a tool for production of prenylated indole derivatives. *Appl. Microbiol. Biotechnol.*, **79**, 951-961 (2008)
- [2] Kremer, A.; Westrich, L.; Li, S.M.: A 7-dimethylallyltryptophan synthase from *Aspergillus fumigatus*: overproduction, purification and biochemical characterization. *Microbiology*, **153**, 3409-3416 (2007)
- [3] Ruan, H.L.; Stec, E.; Li, S.M.: Production of diprenylated indole derivatives by tandem incubation of two recombinant dimethylallyltryptophan synthases. *Arch. Microbiol.*, **191**, 791-795 (2009)