

1 Nomenclature

EC number

1.14.19.5

Systematic name

acyl-CoA,hydrogen donor:oxygen Δ^{11} -oxidoreductase

Recommended name

 Δ^{11} -fatty-acid desaturase

Synonyms

APTQ desaturase <13> [15]

Cro-Z/E11 <6> [8]

 Δ^{11} desaturase <3,4,10> [6,12,13,16] Δ^{11} -(Z)-desaturase <1,2> [4] Δ^{11} -desaturase <5,6,8,9,14,15,16> (<5> several Δ^{11} -desaturase systems: one produces a large quantity of (Z)-11-hexadecenoic acid and another produces (E)1-tetradecenoic acid [7]) [2,7,8,9,17,18] Δ^{11} -fatty-acid desaturase <17,18> [19,20] Δ^{11} -myristoyl-CoA desaturase <4> [1] Δ^{11} -palmitoyl-CoA-desaturase <4> [3] Δ^{11} -palmitoyl-coenzyme A desaturase <6> [8]Dpu- Δ^{11} -APSQ <17> [19]Dpu- Δ^{11} -LPAE <17> [19]

LATPG1 <18> [20]

Lca-KPVQ <14> [17]

OfuZ/E11 protein <15> [18]

OscZ/E11 protein <16> [18]

PDesat-Tn Δ^{11} Z protein <7> [11]

SlS//E11 <10> [6]

TpDESN <9> [2]

Z/E11-desaturase <6> [8]

acyl-CoA <6> [8]

acyl-CoA Δ^{11} -desaturase <7> [11]

acyl-CoA desaturase <14> [17]

bifunctional Δ^{11} -desaturase <4> [12]bifunctional Z- Δ^{11} -desaturase <13> [15]fatty acid Δ^{11} -desaturase <6> [8]sphingolipid long chain base Δ^8 desaturase <8> [9]

CAS registry number

77000-04-5

2 Source Organism

- <1> *Bombyx mori* [4]
- <2> *Manduca sexta* [4]
- <3> *Trichoplusia ni* [13]
- <4> *Spodoptera littoralis* [1,3,10,12,13,16]
- <5> *Choristoneura fumiferana* [7]
- <6> *Choristoneura rosaceana* (UNIPROT accession number: Q8ISS3) [8]
- <7> *Trichoplusia ni* (UNIPROT accession number: O44390) [11]
- <8> *Thaumetopoea pityocampa* [9]
- <9> *Thalassiosira pseudonana* (UNIPROT accession number: Q6RT18) [2]
- <10> *Spodoptera littoralis* (UNIPROT accession number: Q6US81) [6]
- <11> *Argyrotaenia velutinana* [5]
- <12> *Ostrinia scapularis* [14]
- <13> *Manduca sexta* (UNIPROT accession number: Q4A181) [15]
- <14> *Lampronia capitella* (UNIPROT accession number: B6CBS5) [17]
- <15> *Ostrinia furnacalis* [18]
- <16> *Ostrinia scapularis* (UNIPROT accession number: Q2V0N6) [18]
- <17> *Dendrolimus punctatus* [19]
- <18> *Ostrinia latipennis* [20]

3 Reaction and Specificity**Catalyzed reaction**

acyl-CoA + reduced acceptor + O₂ = Δ^{11} -acyl-CoA + acceptor + 2 H₂O (<4> active site modeling [12])

Reaction type

oxidation
reduction

Natural substrates and products

- S** palmitic acid + reduced acceptor + O₂ <9> (<9> enzyme is not involved in production of polyunsaturated fatty acids [2]) (Reversibility: ?) [2]
- P** hexadec-11-enoic acid + acceptor + H₂O
- S** stearoyl-CoA + reduced acceptor + O₂ <7> (Reversibility: ?) [11]
- P** oleoyl-CoA + acceptor + H₂O
- S** Additional information <4,7,14,15,16,17> (<7> all unsaturated pheromone products are produced via a Δ^{11} Z-desaturation mechanism [11]; <4> the enzyme is involved in the biosynthesis of *Spodoptera littoralis* sex pheromone [1]; <14> catalyses key reactions leading to mono- and di-unsaturated fatty acyl-moities [17]; <15,16> Δ^{11} -desaturase is involved in the biosynthesis of Z/E11-14:OAc [18]; <17> heterologous expression

in yeast shows that Dpu- Δ^{11} -LPAE produces large amounts of Δ^{11} -monoenoic acids, in particular the Z11-16:Me and Z11-18:Me in a 1.5:1 ratio. When supplemented with the Z9-16:Me, yeast extracts of Dpu-D112-LPAE contained no 9,11-16:Me, but contain small amounts of E9,Z11- and E9,E11-16:Me when supplemented with E9-16:Me [19]; <17> heterologous expression in yeast shows that Dpu- Δ^{11} -APSQ produces a series of mono-unsaturated products. These monoenes are identified as Δ^8 -12:Me, Δ^8 -14:Me and Δ 8-16:Me. When supplemented with the Z9-16:acid, yeast cells transformed with Dpu- Δ^{11} -APSQ do not produce the Δ^8 -unsaturated fatty acid methyl esters, but produce significant amount of di-unsaturated 9,11- C_{16} methylesters [19]) (Reversibility: ?) [1,11,17,18,19]

P ?

Substrates and products

- S (Z)-11-tetradecenoyl-CoA + reduced acceptor + O₂ <4> (<4> (E,E)-10,12-tetradecadienoic acid is produced from (Z)-11-tetradecenoic acid by desaturation and concomitant migration of the precursor double bond [12]) (Reversibility: ?) [12]
- P (E,E)-10,12-tetradecadienoyl + acceptor + H₂O
- S methyl myristate + reduced electron acceptor + O₂ <6,11> (Reversibility: ?) [5,8]
- P methyl (11E)-tetradec-11-enoate + methyl (11Z)-tetradec-11-enoate + acceptor + H₂O (<6> in the ratio 7:1 [8])
- S myristic acid + NADH + ? <4> (<4> 50% of the activity with palmitic acid [3]) (Reversibility: ?) [3]
- P tetradec-11-enoic acid + NAD⁺ + ?
- S myristic acid + reduced acceptor + O₂ <10> (Reversibility: ?) [6]
- P (Z)-tetradec-11-enoic acid + (E)-tetradec-11-enoic acid + acceptor + H₂O
- S myristoyl-CoA + reduced acceptor + O₂ <4> (<4> reaction involves a first slow, isotope-sensitive C₁₁-H bond cleavage, with probable formation of an unstable intermediate, followed by a second fast C₁₂-H bond removal [1]) (Reversibility: ?) [1]
- P (Z)-tridec-11-enoyl-CoA + (E)-tridec-11-enoyl-CoA + acceptor + H₂O
- S palmitic acid + NADH + ? <4> (Reversibility: ?) [3]
- P (Z)-hexadec-11-enoic acid + NAD⁺ + ?
- S palmitic acid + reduced acceptor + O₂ <8> (Reversibility: ?) [9]
- P (Z)-11-hexadecenoic acid + acceptor + H₂O
- S palmitic acid + reduced acceptor + O₂ <1,2,10> (<1,2> pro-(R) C(11)-H and pro-(R) C(12)-H stereospecificity [4]) (Reversibility: ?) [4,6]
- P (Z)-hexadec-11-enoic acid + acceptor + H₂O
- S palmitic acid + reduced acceptor + O₂ <9> (<9> enzyme is not involved in production of polyunsaturated fatty acids [2]) (Reversibility: ?) [2]
- P hexadec-11-enoic acid + acceptor + H₂O
- S palmitoyl-CoA + NADH + ? <4> (Reversibility: ?) [3]
- P (Z)-hexadec-11-enoyl-CoA + NAD⁺ + ?

- S** stearic acid + NADH + ? <4> (<4> 18% of the activity with palmitic acid [3]) (Reversibility: ?) [3]
- P** oleic acid + NAD⁺ + ?
- S** stearyl-CoA + reduced acceptor + O₂ <7> (Reversibility: ?) [11]
- P** oleoyl-CoA + acceptor + H₂O
- S** Additional information <4,5,7,10,11,13,14,15,16,17,18> (<7> all unsaturated pheromone products are produced via a Δ^{11} Z-desaturation mechanism [11]; <4> the enzyme is involved in the biosynthesis of *Spodoptera littoralis* sex pheromone [1]; <5> several Δ^{11} -desaturase systems: one produces a large quantity of (Z)-11-hexadecenoic acid and another produces (E)-1-tetradecenoic acid [7]; <11> the enzyme produces a mixture of Z/E11-14:acids and exhibits no activity with C₁₆ and C₁₈ saturated fatty acid precursors [5]; <10> the recombinant enzyme expressed in yeast produces a mixture of E11-14:fatty acid, Z11-14:fatty acid, Z11-16:fatty acid and Z-11-18:fatty acid [6]; <13> MsexAPTQ desaturase catalyses the production of Z11-hexadecenoate and (Z10,E12)- and (E10,E12)-hexadecadienoates via 1,4-desaturation of the Z11-16 substrate, in a stereospecific manner, GC-MS analysis of conjugated dienes formed by APTQ desaturase in recombinant *Saccharomyces cerevisiae* cells, overview [15]; <4> substrate specificity, the enzyme catalyzes the formation of methylenecyclopropanes by enzymatic desaturation of 11-cyclopropylundecanoic acid and its disubstituted cis- and trans-derivatives 11-(cis-2-methylcyclopropyl)undecanoic acid, 11-(cis-2-ethylcyclopropyl)undecanoic acid, 11-(cis-2-propylcyclopropyl)undecanoic acid, 11-(trans-2-ethylcyclopropyl)undecanoic acid, and 11-(trans-2-propylcyclopropyl)undecanoic acid, detailed overview [16]; <14> catalyses key reactions leading to mono- and di-unsaturated fatty acyl-moieties [17]; <15,16> Δ^{11} -desaturase is involved in the biosynthesis of Z/E11-14:OAc [18]; <17> heterologous expression in yeast shows that Dpu-D11-LPAE produces large amounts of Δ^{11} -monoenoic acids, in particular the Z11-16:Me and Z11-18:Me in a 1.5:1 ratio. When supplemented with the Z9-16:Me, yeast extracts of Dpu-D112-LPAE contained no 9,11-16:Me, but contain small amounts of E9,Z11- and E9,E11-16:Me when supplemented with E9-16:Me [19]; <17> heterologous expression in yeast shows that Dpu- Δ^{11} -APSQ produces a series of mono-unsaturated products. These monoenes are identified as Δ^8 -12:Me, Δ 8-14:Me and Δ^8 -16:Me. When supplemented with the Z9-16:acid, yeast cells transformed with Dpu- Δ^{11} -APSQ do not produce the Δ^8 -unsaturated fatty acid methyl esters, but produce significant amount of di-unsaturated 9,11-C₁₆ methyl esters [19]; <18> in the presence of tetradecanoic acid, only (E)-11-tetradecenoic acid is produced in Sf9 cells infected with recombinant baculovirus expressing LATPG1. Sf9 cells infected with the control virus do not show Δ^{11} -desaturase activity [20]) (Reversibility: ?) [1,5,6,7,11,15,16,17,18,19,20]
- P** ?

Inhibitors

11-fluorotetradecanoic acid <4> (<4> 50% inhibition at 1:1 substrate/inhibitor ratio [10]) [10]

KCN <4> (<4> 83% inhibition by 1 mM, 95% inhibition by 5 mM [3]) [3]

NaN₃ <4> (<4> 92% inhibition b [3]) [3]

Cofactors/prosthetic groups

NADH <4> (<4> NADPH is a less effective electron donor. Highest activity with 1 mM of electron donors, reduced activity below [3]) [3]

pH-Optimum

6.8-7.2 <4> [3]

Temperature optimum (°C)

25 <4> (<4> assay at [16]) [3,16]

4 Enzyme Structure

Molecular weight

37800 <18> (<18> calculated from cDNA [20]) [20]

5 Isolation/Preparation/Mutation/Application

Source/tissue

abdomen <12> (<12> terminal abdominal segments with pheromone gland of female adults, RT-PCR [14]) [14]

fat body <17> [19]

pheromone gland <4,6,8,11,13,17> (<11> abdominal [5]) [3,5,8,9,10,15,19]

Additional information <12> (<12> in intersegmental membrane, 8th - 9th, in-situ hybridization [14]; <12> not in head, thorax, abdomen exclusive of the terminal abdominal segments [14]) [14]

Localization

microsome <4> [3]

Cloning

<3> (expressed in yeast) [13]

<4> (expressed as polyhistidine-tagged protein in elongase 1 and Δ^9 desaturase-deficient yeast cells) [12]

<4> (expressed in yeast) [13]

<4> (expression in *Saccharomyces cerevisiae* Δ elo1/ Δ ole mutant strain, which is both elongase 1 and Δ^9 desaturase-deficient, using a Cu²⁺ inducible expression vector, the recombinant expression leads to producing large quantities of C₁₁-monounsaturated fatty acids, mainly (Z)-11-hexadecenoic acid, (E,E)-10,12-tetradecadienoic acid, minor amounts of (E,Z)-10,12-hexadecadienoic acid, and very low amounts of (E,Z)-10,12-tetradecadienoic isomer) [12]

<6> [8]

<7> (genetic trans-formation of a desaturase-deficient strain of the yeast *Saccharomyces cerevisiae* with an expression plasmid encoding PDesat-Tn Δ^{11} Z results in complementation of the strain's fatty acid auxotrophy and the production of Δ^{11} Z-unsaturated fatty acids) [11]

<10> (expression in yeast) [6]

<11> [5]

<13> (gene d2, DNA and amino acid sequence determination and analysis, phylogenetic analysis, functional expression of MsexAPTQ desaturase in *Saccharomyces cerevisiae* strain W303) [15]

<14> (into a copper-inducible pYEX vector to assess its desaturase activity and then transformed into a desaturase- and elongase-deficient mutant *ole1 elo1* yeast strain) [17]

<16> (partial cDNAs are amplified and cloned into the vector pCold I DNA, recombinant proteins are produced by *Escherichia coli* BL21 cells and used as antigens to raise antibodies in mice) [18]

<17> (heterologously expressed in *Saccharomyces cerevisiae*) [19]

<18> (C-terminal His-tagged fusion protein expressed using recombinant baculoviruses) [20]

6 Stability

Storage stability

<4>, -80°C, stable for at least 1 h [3]

References

- [1] Pinilla, A.; Camps, F.; Fabrias, G.: Cryptoregiochemistry of the Δ^{11} -myristoyl-CoA desaturase involved in the biosynthesis of *Spodoptera littoralis* sex pheromone. *Biochemistry*, **38**, 15272-15277 (1999)
- [2] Tonon, T.; Harvey, D.; Qing, R.; Li, Y.; Larson, T.R.; Graham, I.A.: Identification of a fatty acid Δ^{11} -desaturase from the microalga *Thalassiosira pseudonana*. *FEBS Lett.*, **563**, 28-34 (2004)
- [3] Rodriguez, F.; Hallahan, D.L.; Pickett, J.A.; Camps, F.: Characterization of the Δ^{11} -palmitoyl-CoA-desaturase from *Spodoptera littoralis* (Lepidoptera: Noctuidae). *Insect Biochem. Mol. Biol.*, **22**, 143-148 (1992)
- [4] Svatos, A.; Kalinova, B.; Boland, W.: Stereochemistry of lepidopteran sex pheromone biosynthesis: a comparison of fatty acid-CoA Δ^{11} -(Z)-desaturases in *Bombyx mori* and *Manduca sexta* female moths. *Insect Biochem. Mol. Biol.*, **29**, 225-232 (1999)
- [5] Liu, W.; Jiao, H.; O'Connor, M.; Roelofs, W.L.: Moth desaturase characterized that produces both Z and E isomers of Δ^{11} -tetradecenoic acids. *Insect Biochem. Mol. Biol.*, **32**, 1489-1495 (2002)

- [6] Rodriguez, S.; Hao, G.; Liu, W.; Pina, B.; Rooney, A.P.; Camps, F.; Roelofs, W.L.; Fabrias, G.: Expression and evolution of Δ^9 and Δ^{11} desaturase genes in the moth *Spodoptera littoralis*. *Insect Biochem. Mol. Biol.*, **34**, 1315-1328 (2004)
- [7] Wolf, W.A.; Roelofs, W.L.: Reinvestigation confirms action of Δ^{11} -desaturases in spruce budworm moth sex pheromone biosynthesis. *J. Chem. Ecol.*, **13**, 1019-1027 (1987)
- [8] Hao, G.; O'Connor, M.; Liu, W.; Roelofs, W.L.: Characterization of Z/E11- and Z9-desaturases from the obliquebanded leafroller moth, *Choristoneura rosaceana*. *J. Insect Sci.*, **2**, 26 (2002)
- [9] Abad, J.L.; Villorbina, G.; Fabrias, G.; Camps, F.: Synthesis and use of stereospecifically deuterated analogues of palmitic acid to investigate the stereochemical course of the Δ^{11} desaturase of the processionary moth. *J. Org. Chem.*, **69**, 7108-7113 (2004)
- [10] Abad, J.L.; Villorbina, G.; Fabrias, G.; Camps, F.: Synthesis of fluorinated analogs of myristic acid as potential inhibitors of Egyptian armyworm (*Spodoptera littoralis*) DELA11 desaturase. *Lipids*, **38**, 865-871 (2003)
- [11] Knipple, D.C.; Rosenfield, C.L.; Miller, S.J.; Liu, W.; Tang, J.; Ma, P.W.; Roelofs, W.L.: Cloning and functional expression of a cDNA encoding a pheromone gland-specific acyl-CoA Δ^{11} -desaturase of the cabbage looper moth, *Trichoplusia ni*. *Proc. Natl. Acad. Sci. USA*, **95**, 15287-15292 (1998)
- [12] Serra, M.; Pina, B.; Bujons, J.; Camps, F.; Fabrias, G.: Biosynthesis of 10,12-dienoic fatty acids by a bifunctional Δ^{11} desaturase in *Spodoptera littoralis*. *Insect Biochem. Mol. Biol.*, **36**, 634-641 (2006)
- [13] Serra, M.; Gauthier, L.T.; Fabrias, G.; Buist, P.H.: Δ^{11} desaturases of *Trichoplusia ni* and *Spodoptera littoralis* exhibit dual catalytic behaviour. *Insect Biochem. Mol. Biol.*, **36**, 822-825 (2006)
- [14] Fukuzawa, M.; Fu, X.; Tatsuki, S.; Ishikawa, Y.: cDNA cloning and in situ hybridization of Δ^{11} -desaturase, a key enzyme of pheromone biosynthesis in *Ostrinia scapularis* (Lepidoptera: Crambidae). *J. Insect Physiol.*, **52**, 430-435 (2006)
- [15] Matouskova, P.; Pichova, I.; Svatos, A.: Functional characterization of a desaturase from the tobacco hornworm moth (*Manduca sexta*) with bifunctional Z11- and 10,12-desaturase activity. *Insect Biochem. Mol. Biol.*, **37**, 601-610 (2007)
- [16] Villorbina, G.; Roura, L.; Camps, F.; Joglar, J.; Fabrias, G.: Enzymatic desaturation of fatty acids: Δ^{11} desaturase activity on cyclopropane acid probes. *J. Org. Chem.*, **68**, 2820-2829 (2003)
- [17] Lienard, M.A.; Strandh, M.; Hedenstroem, E.; Johansson, T.; Loefstedt, C.: Key biosynthetic gene subfamily recruited for pheromone production prior to the extensive radiation of Lepidoptera. *BMC Evol. Biol.*, **8**, 270 (2008)
- [18] Sakai, R.; Fukuzawa, M.; Nakano, R.; Tatsuki, S.; Ishikawa, Y.: Alternative suppression of transcription from two desaturase genes is the key for species-specific sex pheromone biosynthesis in two *Ostrinia* moths. *Insect Biochem. Mol. Biol.*, **39**, 62-67 (2009)

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- [19] Lienard, M.A.; Lassance, J.M.; Wang, H.L.; Zhao, C.H.; Piskur, J.; Johansson, T.; Loefstedt, C.: Elucidation of the sex-pheromone biosynthesis producing 5,7-dodecadienes in *Dendrolimus punctatus* (Lepidoptera: Lasiocampidae) reveals Δ^{11} - and Δ^9 -desaturases with unusual catalytic properties. *Insect Biochem. Mol. Biol.*, **40**, 440-452 (2010)
- [20] Fujii, T.; Ito, K.; Tatematsu, M.; Shimada, T.; Katsuma, S.; Ishikawa, Y.: Sex pheromone desaturase functioning in a primitive *Ostrinia* moth is cryptically conserved in congeners genomes. *Proc. Natl. Acad. Sci. USA*, **108**, 7102-7106 (2011)