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(71) Applicant: BASF AGRO B. V. [NL/NL]; Groningsingel 1, NL-6835 EA Arnhem (NL).

(71) Applicant (for MN only): BASF (CHINA) COMPANY LIMITED [CN/CN]; 300 Jiangxinsha Road, Shanghai 200137 (CN).

(72) Inventors: APONTE, Raphael; G7 18, 68159 Mannheim (DE). TRESCH, Stefan; Rieslingweg 18, 67281 Kirchheim (DE). WITSCH, Matthias; Höhenweg 12b, 67098 Bad Dürkheim (DE). LERCHL, Jens; Golmer Fichten 5, 14476 Golm (DE). MASSA, Dario; N3, 3, 68161 Mannheim (DE). SEISER, Tobias; Waldparkstr. 34, 68163 Mannheim (DE). MIETZNER, Thomas; Rehbergstr. 62, 76855 Annweiler (DE). PAULIK, Jill Marie; 215 King George Loop, Cary, NC 27511 (US). BROMMER, Chad; 1903 Ridley St., Raleigh, NC 27608 (US).

(74) Agent: PROBST, Joseph-Christopher; BASF Construction Chemicals GmbH, Dr.-Albert-Frank-Str. 32, 83308 Trostberg (DE).

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(54) Title: PLANTS HAVING INCREASED TOLERANCE TO HERBICIDES

(57) Abstract: Provided a method for controlling undesired vegetation at a plant cultivation site, the method comprising the steps of providing, at said site, a plant that comprises at least one nucleic acid comprising a nucleotide sequence encoding a wild-type or a mutated protoporphyrinogen oxidase (PPO) which is resistant or tolerant to a PPO-inhibiting herbicide by applying to said site an effective amount of said herbicide. Further provided plants comprising wild-type or mutated PPO enzymes, and methods of obtaining such plants.



PLANTS HAVING INCREASED TOLERANCE TO HERBICIDES

This application claims priority to US provisional applications number US 61/864671 and US 61/864672 the contents of which are incorporated herein by reference in its entirety.

FIELD OF THE INVENTION

The present invention relates in general to methods for conferring on plants agricultural level tolerance to a herbicide. Particularly, the invention refers to plants having an increased tolerance to PPO-inhibiting herbicides. More specifically, the present invention relates to methods and plants obtained by mutagenesis and cross-breeding and transformation that have an increased tolerance to PPO-inhibiting herbicides.

BACKGROUND OF THE INVENTION

Herbicides that inhibit protoporphyrinogen oxidase (hereinafter referred to as Protoporphyrinogen oxidase or PPO; EC:1.3.3.4), a key enzyme in the biosynthesis of protoporphyrin IX, have been used for selective weed control since the 1960s. PPO catalyzes the last common step in chlorophyll and heme biosynthesis which is the oxidation of protoporphyrinogen IX to protoporphyrin IX. (Matringe et al. 1989. *Biochem. J.* 260: 231). PPO-inhibiting herbicides include many different structural classes of molecules (Duke et al. 1991. *Weed Sci.* 39: 465; Nandihalli et al. 1992. *Pesticide Biochem. Physiol.* 43: 193; Matringe et al. 1989. *FEBS Lett.* 245: 35; Yanase and Andoh. 1989. *Pesticide Biochem. Physiol.* 35: 70). These herbicidal compounds include the diphenylethers {e.g. lactofen, (+-)-2-ethoxy-1-methyl-2-oxoethyl 5-{2-chloro-4-(trifluoromethyl)phenoxy}-2-nitrobenzoate; acifluorfen, 5-{2-chloro-4-(trifluoromethyl)phenoxy}-2-nitrobenzoic acid; its methyl ester; or oxyfluorfen, 2-chloro-1-(3-ethoxy-4-nitrophenoxy)-4-(trifluorobenzene)}, oxidiazoles, (e.g. oxidiazon, 3-{2,4-dichloro-5-(1-methylethoxy)phenyl}-5-(1,1-dimethylethyl)-1,3,4-oxadiazol-2-(3H)-one), cyclic imides (e.g. S-23142, N-(4-chloro-2-fluoro-5-propargyloxyphenyl)-3,4,5,6-tetrahydrophthalimide; chlorophthalim, N-(4-chlorophenyl)-3,4,5,6-tetrahydrophthalimide), phenyl pyrazoles (e.g. TNPP-ethyl, ethyl 2-{1-(2,3,4-trichlorophenyl)-4-nitropyrzoly-5-oxo}propionate; M&B 39279), pyridine derivatives (e.g. LS 82-556), and phenopylate and its O-phenylpyrrolidino- and piperidinocarbamate analogs. Many of these compounds competitively inhibit the normal reaction catalyzed by the enzyme, apparently acting as substrate analogs.

Application of PPO-inhibiting herbicides results in the accumulation of protoporphyrinogen IX in the chloroplast and mitochondria, which is believed to leak into the cytosol where it is oxidized by a peroxidase. When exposed to light, protoporphyrin IX causes formation of singlet oxygen in the cytosol and the formation of other reactive oxygen species, which can cause lipid peroxidation and membrane disruption leading to rapid cell death (Lee et al. 1993. *Plant Physiol.* 102: 881).

Not all PPO enzymes are sensitive to herbicides which inhibit plant PPO enzymes. Both the *Escherichia coli* and *Bacillus subtilis* PPO enzymes (Sasarmen et al. 1993. *Can. J. Microbiol.*

39: 1155; Dailey et al. 1994. J. Biol. Chem. 269: 813) are resistant to these herbicidal inhibitors. Mutants of the unicellular alga *Chlamydomonas reinhardtii* resistant to the phenylimide herbicide S-23142 have been reported (Kataoka et al. 1990. J. Pesticide Sci. 15: 449; Shibata et al. 1992. In Research in Photosynthesis, Vol. III, N. Murata, ed. Kluwer: Netherlands. pp. 567-70). At least one of these mutants appears to have an altered PPO activity that is resistant not only to the herbicidal inhibitor on which the mutant was selected, but also to other classes of protox inhibitors (Oshio et al. 1993. Z. Naturforsch. 48c: 339; Sato et al. 1994. In ACS Symposium on Porphyrin Pesticides, S. Duke, ed. ACS Press: Washington, D.C.). A mutant tobacco cell line has also been reported that is resistant to the inhibitor S-21432 (Che et al. 1993. Z. Naturforsch. 48c: 350). Auxotrophic *E. coli* mutants have been used to confirm the herbicide resistance of cloned plant PPO-inhibiting herbicides.

Three main strategies are available for making plants tolerant to herbicides, i.e. (1) detoxifying the herbicide with an enzyme which transforms the herbicide, or its active metabolite, into non-toxic products, such as, for example, the enzymes for tolerance to bromoxynil or to basta (EP242236, EP337899); (2) mutating the target enzyme into a functional enzyme which is less sensitive to the herbicide, or to its active metabolite, such as, for example, the enzymes for tolerance to glyphosate (EP293356, Padgett S. R. et al., J. Biol. Chem., 266, 33, 1991); or (3) overexpressing the sensitive enzyme so as to produce quantities of the target enzyme in the plant which are sufficient in relation to the herbicide, in view of the kinetic constants of this enzyme, so as to have enough of the functional enzyme available despite the presence of its inhibitor. The third strategy was described for successfully obtaining plants which were tolerant to PPO inhibitors (see e.g. US5,767,373 or US5,939,602, and patent family members thereof.). In addition, US 2010/0100988 and WO 2007/024739 discloses nucleotide sequences encoding amino acid sequences having enzymatic activity such that the amino acid sequences are resistant to PPO inhibitor herbicidal chemicals, in particular 3-phenyluracil inhibitor specific PPO mutants.

WO 2012/080975 discloses plants the tolerance of which to a PPO-inhibiting herbicide named "benzoxazinone-derivative" herbicide (1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione) had been increased by transforming said plants with nucleic acids encoding mutated PPO enzymes. In particular, WO 2012/080975 discloses that the introduction of nucleic acids which code for a mutated PPO of an *Amaranthus* type II PPO in which the Arginine at position 128 had been replaced by a leucine, alanine, or valine, and the phenylalanine at position 420 had been replaced by a methionine, cysteine, isoleucine, leucine, or threonine, confers increased tolerance/resistance to a benzoxazinone-derivative herbicide.

The inventors of the present invention have now surprisingly found that those types of double-mutants and, furthermore, novel substitutions for R128 and F420 which are not disclosed in WO 2012/080975 confer increased tolerance/resistance to a wide variety of PPO inhibitors including, but not limited to a "benzoxazinone-derivative" (1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione) herbicide described in WO 2012/080975. Thus, to date, the prior art has not described

PPO-inhibiting herbicide tolerant plants containing a mutated PPO nucleic acid according to the present invention, which are tolerant/resistant to a broad spectrum of PPO inhibitors. Therefore, what is needed in the art are crop plants and crop plants having increased tolerance to herbicides such as PPO-inhibiting herbicide and containing at least one wildtype and/or mutated PPO nucleic acid according to the present invention. Also needed are methods for controlling weed growth in the vicinity of such crop plants or crop plants. These compositions and methods would allow for the use of spray over techniques when applying herbicides to areas containing crop plants or crop plants.

10 SUMMARY OF THE INVENTION

The problem is solved by the present invention which refers to a method for controlling undesired vegetation at a plant cultivation site, the method comprising the steps of:

- 15 a) providing, at said site, a plant that comprises at least one nucleic acid comprising a nucleotide sequence encoding a wild type protoporphyrinogen oxidase (PPO) or a mutated protoporphyrinogen oxidase (PPO) which is resistant or tolerant to a PPO-inhibiting herbicide,
- b) applying to said site an effective amount of said herbicide.

- 20 In addition, the present invention refers to a method for identifying a PPO-inhibiting herbicide by using a wild-type or mutated PPO of the present invention encoded by a nucleic acid which comprises the nucleotide sequence of SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47, or a variant thereof.

- 25 Said method comprises the steps of:

- a) generating a transgenic cell or plant comprising a nucleic acid encoding a mutated PPO of the present invention, wherein the mutated PPO of the present invention is expressed;
- b) applying a PPO-inhibiting herbicide to the transgenic cell or plant of a) and to a control cell or plant of the same variety;
- 30 c) determining the growth or the viability of the transgenic cell or plant and the control cell or plant after application of said test compound, and
- d) selecting test compounds which confer reduced growth to the control cell or plant as compared to the growth of the transgenic cell or plant.

- 35 Another object refers to a method of identifying a nucleotide sequence encoding a mutated PPO which is resistant or tolerant to a PPO-inhibiting herbicide, the method comprising:

- a) generating a library of mutated PPO-encoding nucleic acids,
- b) screening a population of the resulting mutated PPO-encoding nucleic acids by expressing each of said nucleic acids in a cell or plant and treating said cell or plant with a PPO-inhibiting herbicide,
- 40 c) comparing the PPO-inhibiting herbicide-tolerance levels provided by said population of mutated PPO encoding nucleic acids with the PPO-inhibiting herbicide-tolerance level provided by a control PPO-encoding nucleic acid,
- d) selecting at least one mutated PPO-encoding nucleic acid that provides a significantly

increased level of tolerance to a PPO-inhibiting herbicide as compared to that provided by the control PPO-encoding nucleic acid.

In a preferred embodiment, the mutated PPO-encoding nucleic acid selected in step d) provides at least 2-fold as much tolerance to a PPO-inhibiting herbicide as compared to that provided by the control PPO-encoding nucleic acid.

The resistance or tolerance can be determined by generating a transgenic plant comprising a nucleic acid sequence of the library of step a) and comparing said transgenic plant with a control plant.

Another object refers to a method of identifying a plant or algae containing a nucleic acid encoding a mutated PPO which is resistant or tolerant to a PPO-inhibiting herbicide, the method comprising:

- a) identifying an effective amount of a PPO-inhibiting herbicide in a culture of plant cells or green algae.
- b) treating said plant cells or green algae with a mutagenizing agent,
- c) contacting said mutagenized cells population with an effective amount of PPO-inhibiting herbicide, identified in a),
- d) selecting at least one cell surviving these test conditions,
- e) PCR-amplification and sequencing of PPO genes from cells selected in d) and comparing such sequences to wild-type PPO gene sequences, respectively.

In a preferred embodiment, the mutagenizing agent is ethylmethanesulfonate.

Another object refers to an isolated and/or recombinantly produced and/or chemically synthesized (synthetic) nucleic acid encoding a mutated PPO, the nucleic acid comprising the sequence of SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47, or a variant thereof, as defined hereinafter.

Another object refers to an isolated mutated PPO polypeptide, the polypeptide comprising the sequence set forth in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, a variant, derivative, orthologue, paralogue or homologue thereof, as defined hereinafter.

In a preferred embodiment, the nucleic acid being identifiable by a method as defined above.

In another embodiment, the invention refers to a plant cell transformed by and expressing a wild-type or a mutated PPO nucleic acid according to the present invention or a plant which has been mutated to obtain a plant expressing, preferably over-expressing a wild-type or a mutated PPO nucleic acid according to the present invention, wherein expression of said nucleic acid in the plant cell results in increased resistance or tolerance to a PPO-inhibiting herbicide as compared to a wild type variety of the plant cell.

In another embodiment, the invention refers to a plant comprising a plant cell according to the

present invention, wherein expression of the nucleic acid in the plant results in the plant's increased resistance to PPO-inhibiting herbicide as compared to a wild type variety of the plant.

In another embodiment, the invention refers to a plant that expresses a mutagenized or recombinant mutated PPO polypeptide, and wherein said mutated PPO confers upon the plant increased herbicide tolerance as compared to the corresponding wild-type variety of the plant when expressed therein

The plants of the present invention can be transgenic or non-transgenic.

Preferably, the expression of the nucleic acid of the invention in the plant results in the plant's increased resistance to PPO-inhibiting herbicides as compared to a wild type variety of the plant.

In another embodiment, the invention refers to a method for growing the plant according to the present invention while controlling weeds in the vicinity of said plant, said method comprising the steps of:

- a) growing said plant ; and
- b) applying a herbicide composition comprising a PPO-inhibiting herbicide to the plant and weeds, wherein the herbicide normally inhibits protoporphyrinogen oxidase, at a level of the herbicide that would inhibit the growth of a corresponding wild-type plant.

In another embodiment, the invention refers to a seed produced by a transgenic plant comprising a plant cell of the present invention, or to a seed produced by the non-transgenic plant that expresses a mutagenized PPO polypeptide, wherein the seed is true breeding for an increased resistance to a PPO-inhibiting herbicide as compared to a wild type variety of the seed.

In another embodiment, the invention refers to a method of producing a transgenic plant cell with an increased resistance to a PPO-inhibiting herbicide as compared to a wild type variety of the plant cell comprising, transforming the plant cell with an expression cassette comprising a wild-type or a mutated PPO nucleic acid.

In another embodiment, the invention refers to a method of producing a transgenic plant comprising, (a) transforming a plant cell with an expression cassette comprising a wild-type or a mutated PPO nucleic acid, and (b) generating a plant with an increased resistance to PPO-inhibiting herbicide from the plant cell.

Preferably, the expression cassette further comprises a transcription initiation regulatory region and a translation initiation regulatory region that are functional in the plant.

In another embodiment, the invention relates to using the mutated PPO of the invention as selectable marker. The invention provides a method of identifying or selecting a transformed plant cell, plant tissue, plant or part thereof comprising a) providing a transformed plant cell,

plant tissue, plant or part thereof, wherein said transformed plant cell, plant tissue, plant or part thereof comprises an isolated nucleic acid encoding a mutated PPO polypeptide of the invention as described hereinafter, wherein the polypeptide is used as a selection marker, and wherein said transformed plant cell, plant tissue, plant or part thereof may optionally comprise a further isolated nucleic acid of interest; b) contacting the transformed plant cell, plant tissue, plant or part thereof with at least one PPO-inhibiting inhibiting compound; c) determining whether the plant cell, plant tissue, plant or part thereof is affected by the inhibitor or inhibiting compound; and d) identifying or selecting the transformed plant cell, plant tissue, plant or part thereof.

10 The invention is also embodied in purified mutated PPO proteins that contain the mutations described herein, which are useful in molecular modeling studies to design further improvements to herbicide tolerance. Methods of protein purification are well known, and can be readily accomplished using commercially available products or specially designed methods, as set forth for example, in Protein Biotechnology, Walsh and Headon (Wiley, 1994).

15 In another embodiment, the invention relates to a combination useful for weed control, comprising (a) a polynucleotide encoding a mutated PPO polypeptide according to the present invention, which polynucleotide is capable of being expressed in a plant to thereby provide to that plant tolerance to a PPO inhibiting herbicide; and (b) a PPO inhibiting herbicide.

20 In another embodiment, the invention relates to a process for preparing a combination useful for weed control comprising (a) providing a polynucleotide encoding a mutated PPO polypeptide according to the present invention, which polynucleotide is capable of being expressed in a plant to thereby provide to that plant tolerance to a PPO inhibiting herbicide; and (b) providing a PPO inhibiting herbicide.

In a preferred embodiment, said step of providing a polynucleotide comprises providing a plant containing the polynucleotide.

30 In another preferred embodiment, said step of providing a polynucleotide comprises providing a seed containing the polynucleotide.

In another preferred embodiment, said process further comprises a step of applying the PPO inhibiting herbicide to the seed.

35 In another embodiment, the invention relates to the use of a combination useful for weed control, comprising (a) a polynucleotide encoding a mutated PPO polypeptide according to the present invention, which polynucleotide is capable of being expressed in a plant to thereby provide to that plant tolerance to a PPO inhibiting herbicide; and (b) a PPO inhibiting herbicide, to control weeds at a plant cultivation site.

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BRIEF DESCRIPTION OF THE DRAWINGS

Figure 1 shows an amino acid sequence alignment of *Amaranthus tuberculatus* (A.tuberculatus), *Amaranthus tuberculatus* resistant (A.tuberculatus_R), *Arabidopsis thaliana* long (A.thaliana_2), *Spinacia oleracea* short (S.oleracea_2), *Nicotiana tabacum* short (N.tabacum_2), *Glycine max* (Glycine_max), *Arabidopsis thaliana* short (A.thaliana_1), *Nicotiana tabacum* long (N.tabacum_1), *Chlamydomonas reinhardtii* long (C.reinhardtii_1), *Zea mays* (Z.mays), *Oryza sativa* (O.sativa_1), *Solanum tuberosum* (S.tuberosum), *Cucumis sativus* (C.sativus), *Cichorium intybus* (C.intybus_1), *Spinacia oleracea* long (S.oleracea_1), *Polytomella* sp. Pringsheim 198.80 (*Polytomella*) PPO sequences. Conserved regions are indicated in light grey, grey and black.

Figure 2 shows wildtype and transgenic *Arabidopsis* plants comprising a nucleic acid encoding a mutated PPO polypeptide (based on SEQ ID NO:2; AMATU_PPO2_R128A_420V); 1 = Kixor [saflufenacil]; 2 = BAS 850H [1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione]; 3 = Spotlight [fluroxypyr]; 4 = Kixor + Spotlight; A = non-transgenic (for any PPOi treatment); B = AMATU_PPO2_R128A_420V transgenic plants

Figure 3 shows T1 Transformed corn 7 days after treatment with 100 g saflufenacil + 50 g ai/ha BAS 850H + 1% (v/v) MSO. Plants were sprayed at the V2-V3 stage. 1 = untransformed control; 2 = Tp-Fdx_AmtuPPX2L_R128A_F420V (Transit peptide of *Silene pratensis* Ferredoxin fused to mutated PPO); 3 = AmtuPPX2L_R128A_F420L

Figure 4 shows T0 Transformed corn 3 days after treatment. Plants were sprayed with 0 or 50 g ai/ha BAS 850H + 1% MSO at the V2-V3 stage. 1 = wildtype, 2 = AmtuPPX2L_R128L_F420M; 3 = AmtuPPX2L_R128A_F420I; 4 = AmtuPPX2L_R128A_F420V; 5 = AmtuPPX2L_R128A_F420L; 6 = AmtuPPX2L_R128M_F420I; 7 = AmtuPPX2L_R128M_F420L; 8 = AmtuPPX2L_R128M_F420V

Figure 5 shows T1 transformed soybean 7 days after treatment with the indicated herbicide + 1% (v/v) MSO. Plants were sprayed at the V2-V3 stage; A = unsprayed; B = saflufenacil 150 g ai/ha; C = BAS 850H 100 g ai/ha; 1 = wildtype control plant; 2 = AmtuPPX2L_R128A_F420M; 3 = AmtuPPX2L_R128A_F420I; 4 = AmtuPPX2L_R128A_F420V;

Figure 6 shows T0 Transformed soybean clones 7 days after indicated treatment. Plants were sprayed at the V2-V3 stage; 1 = wildtype control; 2 = AmtuPPX2L_R128L_F420V; A = saflufenacil g ai/ha + 1% MSO; B = BAS 850H g ai/ha + 1% MSO

Figure 7 shows T2 Transformed soybean 4 days after the indicated treatment. Plants were sprayed at the V2-V3 stage. Treatments contained 1% (v/v) MSO (methylated soy oil – based spray adjuvant; also known as Destiny HC); 1 = wildtype; 2 = AmtuPPX2L_R128A_F420V; 3 = AmtuPPX2L_R128A_F420L; 4 = AmtuPPX2L_R128A_F420M; 5 = AmtuPPX2L_R128A_F420I; A = unsprayed; B = 100 g ai/ha saflufenacil+50 g ai/ha BAS 850H; C = 200 g ai/ha saflufenacil+100 g ai/ha BAS 850H; D = 100 g ai/ha saflufenacil+140 g ai/ha flumioxazin; E = 100 g ai/ha saflufenacil+560 g ai/ha sulfentrazone;

KEY TO SEQUENCE LISTING

Table 1

SEQ. ID NO:	Description	Organism	Gene	Accession No:
1	PPO nucleic acid	Amaranthus tuberculatus	PPX2L_WC	DQ386114
2	PPO amino acid	Amaranthus tuberculatus	ABD52326	
3	PPO nucleic acid	Amaranthus tuberculatus	PPX2L_AC	DQ386117
4	PPO amino acid	Amaranthus tuberculatus	ABD52329	
5	PPO nucleic acid	Amaranthus tuberculatus	PPX2L_CC_R	DQ386118
6	PPO amino acid	Amaranthus tuberculatus	ABD52330	
7	PPO nucleic acid	Amaranthus tuberculatus	PPX2L_AC_R	DQ386116
8	PPO amino acid	Amaranthus tuberculatus	ABD52328	
9	PPO nucleic acid	Arabidopsis thaliana	PPX	AB007650
10	PPO amino acid	Arabidopsis thaliana	BAB08301	
11	PPO nucleic acid	Nicotiana tabacum	ppxl	AF044128
12	PPO amino acid	Nicotiana tabacum	AAD02290	
13	PPO nucleic acid	Cichorium intybus	PPX1	AF160961
14	PPO amino acid	Cichorium intybus	AF160961_1	
15	PPO nucleic acid	Spinacia oleracea	SO-POX1	AB029492
16	PPO amino acid	Spinacia oleracea	BAA96808	
17	PPO nucleic acid	Spinacia oleracea	SO-POX2	AB046993
18	PPO amino acid	Spinacia oleracea	BAB60710	
19	PPO nucleic acid	Solanum tuberosum	PPOX	AJ225107

20	PPO amino acid	Solanum tuberosum	CAA12400	
21	PPO nucleic acid	Zea mays	ZM_BFc0091B03	BT063659
22	PPO amino acid	Zea mays	ACN28356	
23	PPO nucleic acid	Zea mays	prpo2	NM_001111534
24	PPO amino acid	Zea mays	NP_001105004	
25	PPO nucleic acid	Chlamydomonas	Ppx1	AF068635
26	PPO amino acid	Chlamydomonas	AAC79685	
27	PPO nucleic acid	Polytomella	PPO	AF332964
28	PPO amino acid	Polytomella	AF332964_1	
29	PPO nucleic acid	Sorghum bicolor	Hyp. Protein	XM_002446665
30	PPO amino acid	Sorghum bicolor	XP_002446710	
31	PPO nucleic acid	Oryza sativa	PPOX1	AB057771
32	PPO amino acid	Oryza sativa	BAB39760	
33	PPO nucleic acid	Amaranthus tuberculatus	PPX2	DQ386113
34	PPO amino acid	Amaranthus tuberculatus	ABD52325	
35	PPO nucleic acid	Arabidopsis thaliana	PPOX	NM_178952
36	PPO amino acid	Arabidopsis thaliana	NP_849283	
37	PPO nucleic acid	Nicotiana tabacum	ppxII	AF044129
38	PPO amino acid	Nicotiana tabacum	AAD02291	
39	PPO nucleic acid	Glycine max	hemG	AB025102
40	PPO amino acid	Glycine max	BAA76348	

41	PPO nucleic acid	Cucumis sativus	CsPPO	AB512426
42	PPO amino acid	Cucumis sativus	BAH84864.1	
43	PPO nucleic acid	Oryza sativa	Hyp. Protein	AL606613
44	PPO amino acid	Oryza sativa	CAE01661	
45	PPO nucleic acid	Oryza sativa	amine oxidase	
46	PPO amino acid	Oryza sativa	Os04g41260.1	
47	PPO nucleic acid	Amaranthus tuberculatus	PPX1	
48	PPO amino acid	Amaranthus tuberculatus	PPO1	

DETAILED DESCRIPTION

The articles "a" and "an" are used herein to refer to one or more than one (i.e., to at least one) of the grammatical object of the article. By way of example, "an element" means one or more elements.

As used herein, the word "comprising," or variations such as "comprises" or "comprising," will be understood to imply the inclusion of a stated element, integer or step, or group of elements, integers or steps, but not the exclusion of any other element, integer or step, or group of elements, integers or steps.

The inventors of the present invention have found, that the tolerance or resistance of a plant to a PPO-inhibiting herbicide could be remarkably increased by overexpressing a nucleic acid encoding a mutated PPO polypeptide comprising the sequence set forth in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, a variant, derivative, orthologue, paralogue or homologue thereof.

The present invention refers to a method for controlling undesired vegetation at a plant cultivation site, the method comprising the steps of:

- a) providing, at said site, a plant that comprises at least one nucleic acid comprising a nucleotide sequence encoding a wild-type protoporphyrinogen oxidase or a mutated protoporphyrinogen oxidase (mutated PPO) which is resistant or tolerant to a PPO-inhibiting herbicide,
- b) applying to said site an effective amount of said herbicide.

The term "control of undesired vegetation" is to be understood as meaning the killing of weeds and/or otherwise retarding or inhibiting the normal growth of the weeds. Weeds, in the broadest

sense, are understood as meaning all those plants which grow in locations where they are undesired, e.g. (crop) plant cultivation sites. The weeds of the present invention include, for example, dicotyledonous and monocotyledonous weeds. Dicotyledonous weeds include, but are not limited to, weeds of the genera: *Sinapis*, *Lepidium*, *Galium*, *Stellaria*, *Matricaria*, *Anthemis*,
 5 *Galinsoga*, *Chenopodium*, *Urtica*, *Senecio*, *Amaranthus*, *Portulaca*, *Xanthium*, *Convolvulus*,
Ipomoea, *Polygonum*, *Sesbania*, *Ambrosia*, *Cirsium*, *Carduus*, *Sonchus*, *Solanum*, *Rorippa*,
Rotala, *Lindernia*, *Lamium*, *Veronica*, *Abutilon*, *Emex*, *Datura*, *Viola*, *Galeopsis*, *Papaver*,
Centaurea, *Trifolium*, *Ranunculus*, and *Taraxacum*. Monocotyledonous weeds include, but are not limited to, weeds of the genera: *Echinochloa*, *Setaria*, *Panicum*, *Digitaria*, *Phleum*, *Poa*,
 10 *Festuca*, *Eleusine*, *Brachiaria*, *Lolium*, *Bromus*, *Avena*, *Cyperus*, *Sorghum*, *Agropyron*,
Cynodon, *Monochoria*, *Fimbristylis*, *Sagittaria*, *Eleocharis*, *Scirpus*, *Paspalum*, *Ischaemum*,
Sphenoclea, *Dactyloctenium*, *Agrostis*, *Alopecurus*, and *Apera*. In addition, the weeds of the present invention can include, for example, crop plants that are growing in an undesired location. For example, a volunteer maize plant that is in a field that predominantly comprises
 15 soybean plants can be considered a weed, if the maize plant is undesired in the field of soybean plants.

The term “plant” is used in its broadest sense as it pertains to organic material and is intended to encompass eukaryotic organisms that are members of the Kingdom Plantae, examples of
 20 which include but are not limited to vascular plants, vegetables, grains, flowers, trees, herbs, bushes, grasses, vines, ferns, mosses, fungi and algae, etc, as well as clones, offsets, and parts of plants used for asexual propagation (e.g. cuttings, pipings, shoots, rhizomes, underground stems, clumps, crowns, bulbs, corms, tubers, rhizomes, plants/tissues produced in tissue culture, etc.). The term “plant” further encompasses whole plants, ancestors and progeny
 25 of the plants and plant parts, including seeds, shoots, stems, leaves, roots (including tubers), flowers, florets, fruits, pedicles, peduncles, stamen, anther, stigma, style, ovary, petal, sepal, carpel, root tip, root cap, root hair, leaf hair, seed hair, pollen grain, microspore, cotyledon, hypocotyl, epicotyl, xylem, phloem, parenchyma, endosperm, a companion cell, a guard cell, and any other known organs, tissues, and cells of a plant, and tissues and organs, wherein
 30 each of the aforementioned comprise the gene/nucleic acid of interest. The term “plant” also encompasses plant cells, suspension cultures, callus tissue, embryos, meristematic regions, gametophytes, sporophytes, pollen and microspores, again wherein each of the aforementioned comprises the gene/nucleic acid of interest.

35 Plants that are particularly useful in the methods of the invention include all plants which belong to the superfamily Viridiplantae, in particular monocotyledonous and dicotyledonous plants including fodder or forage legumes, ornamental plants, food crops, trees or shrubs selected from the list comprising *Acer* spp., *Actinidia* spp., *Abelmoschus* spp., *Agave sisalana*,
Agropyron spp., *Agrostis stolonifera*, *Allium* spp., *Amaranthus* spp., *Ammophila arenaria*,
 40 *Ananas comosus*, *Annona* spp., *Apium graveolens*, *Arachis* spp., *Artocarpus* spp., *Asparagus officinalis*, *Avena* spp. (e.g. *Avena sativa*, *Avena fatua*, *Avena byzantina*, *Avena fatua* var. *sativa*, *Avena hybrida*), *Averrhoa carambola*, *Bambusa* sp., *Benincasa hispida*, *Bertholletia excelsa*, *Beta vulgaris*, *Brassica* spp. (e.g. *Brassica napus*, *Brassica rapa* ssp. [canola, oilseed rape, turnip rape]), *Cadaba farinosa*, *Camellia sinensis*, *Canna indica*, *Cannabis sativa*,

Capsicum spp., Carex elata, Carica papaya, Carissa macrocarpa, Carya spp., Carthamus
 tinctorius, Castanea spp., Ceiba pentandra, Cichorium endivia, Cinnamomum spp., Citrullus
 lanatus, Citrus spp., Cocos spp., Coffea spp., Colocasia esculenta, Cola spp., Corchorus sp.,
 Coriandrum sativum, Corylus spp., Crataegus spp., Crocus sativus, Cucurbita spp., Cucumis
 5 spp., Cynara spp., Daucus carota, Desmodium spp., Dimocarpus longan, Dioscorea spp.,
 Diospyros spp., Echinochloa spp., Elaeis (e.g. Elaeis guineensis, Elaeis oleifera), Eleusine
 coracana, Eragrostis tef, Erianthus sp., Eriobotrya japonica, Eucalyptus sp., Eugenia uniflora,
 Fagopyrum spp., Fagus spp., Festuca arundinacea, Ficus carica, Fortunella spp., Fragaria spp.,
 Ginkgo biloba, Glycine spp. (e.g. Glycine max, Soja hispida or Soja max), Gossypium hirsutum,
 10 Helianthus spp. (e.g. Helianthus annuus), Hemerocallis fulva, Hibiscus spp., Hordeum spp. (e.g.
 Hordeum vulgare), Ipomoea batatas, Juglans spp., Lactuca sativa, Lathyrus spp., Lens
 culinaris, Linum usitatissimum, Litchi chinensis, Lotus spp., Luffa acutangula, Lupinus spp.,
 Luzula sylvatica, Lycopersicon spp. (e.g. Lycopersicon esculentum, Lycopersicon lycopersicum,
 Lycopersicon pyriforme), Macrotyloma spp., Malus spp., Malpighia emarginata, Mammea
 15 americana, Mangifera indica, Manihot spp., Manilkara zapota, Medicago sativa, Melilotus spp.,
 Mentha spp., Miscanthus sinensis, Momordica spp., Morus nigra, Musa spp., Nicotiana spp.,
 Olea spp., Opuntia spp., Ornithopus spp., Oryza spp. (e.g. Oryza sativa, Oryza latifolia),
 Panicum miliaceum, Panicum virgatum, Passiflora edulis, Pastinaca sativa, Pennisetum sp.,
 Persea spp., Petroselinum crispum, Phalaris arundinacea, Phaseolus spp., Phleum pratense,
 20 Phoenix spp., Phragmites australis, Physalis spp., Pinus spp., Pistacia vera, Pisum spp., Poa
 spp., Populus spp., Prosopis spp., Prunus spp., Psidium spp., Punica granatum, Pyrus
 communis, Quercus spp., Raphanus sativus, Rheum rhabarbarum, Ribes spp., Ricinus
 communis, Rubus spp., Saccharum spp., Salix sp., Sambucus spp., Secale cereale, Sesamum
 spp., Sinapis sp., Solanum spp. (e.g. Solanum tuberosum, Solanum integrifolium or Solanum
 25 lycopersicum), Sorghum bicolor, Spinacia spp., Syzygium spp., Tagetes spp., Tamarindus
 indica, Theobroma cacao, Trifolium spp., Tripsacum dactyloides, Triticosecale rimpai, Triticum
 spp. (e.g. Triticum aestivum, Triticum durum, Triticum turgidum, Triticum hybernum, Triticum
 macha, Triticum sativum, Triticum monococcum or Triticum vulgare), Tropaeolum minus,
 Tropaeolum majus, Vaccinium spp., Vicia spp., Vigna spp., Viola odorata, Vitis spp., Zea mays,
 30 Zizania palustris, Ziziphus spp., amaranth, artichoke, asparagus, broccoli, Brussels sprouts,
 cabbage, canola, carrot, cauliflower, celery, collard greens, flax, kale, lentil, oilseed rape, okra,
 onion, potato, rice, soybean, strawberry, sugar beet, sugar cane, sunflower, tomato, squash, tea
 and algae, amongst others. According to a preferred embodiment of the present invention, the
 plant is a crop plant. Examples of crop plants include inter alia soybean, sunflower, canola,
 35 alfalfa, rapeseed, cotton, tomato, potato or tobacco. Further preferably, the plant is a
 monocotyledonous plant, such as sugarcane. Further preferably, the plant is a cereal, such as
 rice, maize, wheat, barley, millet, rye, sorghum or oats.

In a preferred embodiment, the plant has been previously produced by a process comprising
 40 recombinantly preparing a plant by introducing and over-expressing a wild-type or mutated PPO
 transgene according to the present invention, as described in greater detail hereinafter.

In another preferred embodiment, the plant has been previously produced by a process
 comprising in situ mutagenizing plant cells, to obtain plant cells which express a mutated PPO.

As disclosed herein, the nucleic acids of the invention find use in enhancing the herbicide tolerance of plants that comprise in their genomes a gene encoding a herbicide-tolerant wild-type or mutated PPO protein. Such a gene may be an endogenous gene or a transgene, as described hereinafter.

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Therefore, in another embodiment the present invention refers to a method of increasing or enhancing the PPO-inhibitor herbicide tolerance or resistance of a plant, the method comprising overexpressing a nucleic acid encoding a mutated PPO polypeptide comprising the sequence set forth in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, a variant, derivative, orthologue, paralogue or homologue thereof.

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Additionally, in certain embodiments, the nucleic acids of the present invention can be stacked with any combination of polynucleotide sequences of interest in order to create plants with a desired phenotype. For example, the nucleic acids of the present invention may be stacked with any other polynucleotides encoding polypeptides having pesticidal and/or insecticidal activity, such as, for example, the *Bacillus thuringiensis* toxin proteins (described in U.S. Patent Nos. 5,366,892; 5,747,450; 5,737,514; 5,723,756; 5,593,881; and Geiser et al (1986) Gene 48: 109).

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By way of example, polynucleotides that may be stacked with the nucleic acids of the present invention include nucleic acids encoding polypeptides conferring resistance to pests/pathogens such as viruses, nematodes, insects or fungi, and the like. Exemplary polynucleotides that may be stacked with nucleic acids of the invention include polynucleotides encoding: polypeptides having pesticidal and/or insecticidal activity, such as other *Bacillus thuringiensis* toxic proteins (described in U.S. Pat. Nos. 5,366,892; 5,747,450; 5,737,514; 5,723,756; 5,593,881; and Geiser et al., (1986) Gene 48:109), lectins (Van Damme et al. (1994) Plant Mol. Biol. 24:825, pentin (described in U.S. Pat. No. 5,981,722), and the like; traits desirable for disease or herbicide resistance (e.g., fumonisin detoxification genes (U.S. Pat. No. 5,792,931); avirulence and disease resistance genes (Jones et al. (1994) Science 266:789; Martin et al., (1993) Science 262:1432; Mindrinos et al. (1994) Cell 78:1089); acetolactate synthase (ALS) mutants that lead to herbicide resistance such as the S4 and/or Hra mutations; glyphosate resistance (e.g., 5-enol-pyrovyl-shikimate-3-phosphate-synthase (EPSPS) gene, described in U.S. Pat. Nos. 4,940,935 and 5,188,642; or the glyphosate N-acetyltransferase (GAT) gene, described in Castle et al. (2004) Science, 304:1151-1154; and in U.S. Patent App. Pub. Nos. 20070004912, 20050246798, and 20050060767)); glufosinate resistance (e.g, phosphinothricin acetyl transferase genes PAT and BAR, described in U.S. Pat. Nos. 5,561,236 and 5,276,268); resistance to herbicides including sulfonyl urea, DHT (2,4D), and PPO herbicides (e.g., glyphosate acetyl transferase, aryloxy alkanoate dioxygenase, acetolactate synthase, and protoporphyrinogen oxidase); a cytochrome P450 or variant thereof that confers herbicide resistance or tolerance to, inter alia, HPPD herbicides (U.S. patent application Ser. No. 12/156,247; U.S. Pat. Nos. 6,380,465; 6,121,512; 5,349,127; 6,649,814; and 6,300,544; and PCT Patent App. Pub. No. WO2007000077); and traits desirable for processing or process products such as high oil (e.g., U.S. Pat. No. 6,232,529); modified oils (e.g., fatty acid desaturase genes (U.S. Pat. No. 5,952,544; WO 94/11516)); modified starches (e.g., ADPG pyrophosphorylases (AGPase), starch synthases (SS), starch branching enzymes (SBE), and

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starch debranching enzymes (SDBE)); and polymers or bioplastics (e.g., U.S. Pat. No. 5,602,321; beta-ketothiolase, polyhydroxybutyrate synthase, and acetoacetyl-CoA reductase (Schubert et al. (1988) J. Bacteriol. 170:5837-5847) facilitate expression of polyhydroxyalkanoates (PHAs)); the disclosures of which are herein incorporated by reference.

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In a particularly preferred embodiment, the plant comprises at least one additional heterologous nucleic acid comprising a nucleotide sequence encoding a herbicide tolerance enzyme selected, for example, from the group consisting of 5-enolpyruvylshikimate-3-phosphate synthase (EPSPS), Glyphosate acetyl transferase (GAT), Cytochrome P450, phosphinothricin acetyltransferase (PAT), Acetohydroxyacid synthase (AHAS; EC 4.1.3.18, also known as acetolactate synthase or ALS), Protoporphyrinogen oxidase (PPGO), Phytoene desaturase (PD) and dicamba degrading enzymes as disclosed in WO 02/068607. The combinations generated can also include multiple copies of any one of the polynucleotides of interest.

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Generally, the term "herbicide" is used herein to mean an active ingredient that kills, controls or otherwise adversely modifies the growth of plants. The preferred amount or concentration of the herbicide is an "effective amount" or "effective concentration." By "effective amount" and "effective concentration" is intended an amount and concentration, respectively, that is sufficient to kill or inhibit the growth of a similar, wild-type, plant, plant tissue, plant cell, or host cell, but that said amount does not kill or inhibit as severely the growth of the herbicide-resistant plants, plant tissues, plant cells, and host cells of the present invention. Typically, the effective amount of a herbicide is an amount that is routinely used in agricultural production systems to kill weeds of interest. Such an amount is known to those of ordinary skill in the art. Herbicidal activity is exhibited by herbicides useful for the the present invention when they are applied directly to the plant or to the locus of the plant at any stage of growth or before planting or emergence. The effect observed depends upon the plant species to be controlled, the stage of growth of the plant, the application parameters of dilution and spray drop size, the particle size of solid components, the environmental conditions at the time of use, the specific compound employed, the specific adjuvants and carriers employed, the soil type, and the like, as well as the amount of chemical applied. These and other factors can be adjusted as is known in the art to promote non-selective or selective herbicidal action. Generally, it is preferred to apply the herbicide postemergence to relatively immature undesirable vegetation to achieve the maximum control of weeds.

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By a "herbicide-tolerant" or "herbicide-resistant" plant, it is intended that a plant that is tolerant or resistant to at least one herbicide at a level that would normally kill, or inhibit the growth of, a normal or wild-type plant. By "herbicide-tolerant wildtype or mutated PPO protein" or "herbicide-resistant wildtype or mutated PPO protein", it is intended that such a PPO protein displays higher PPO activity, relative to the PPO activity of a wild-type PPO protein, when in the presence of at least one herbicide that is known to interfere with PPO activity and at a concentration or level of the herbicide that is known to inhibit the PPO activity of the wild-type mutated PPO protein. Furthermore, the PPO activity of such a herbicide-tolerant or herbicide-resistant mutated PPO protein may be referred to herein as "herbicide-tolerant" or "herbicide-resistant" PPO activity.

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Generally, if the PPO-inhibiting herbicides (also referred to as compounds A) and/or the herbicidal compounds B as described herein, which can be employed in the context of the present invention, are capable of forming geometrical isomers, for example E/Z isomers, it is possible to use both, the pure isomers and mixtures thereof, in the compositions useful for the present invention. If the PPO-inhibiting herbicides A and/or the herbicidal compounds B as described herein have one or more centers of chirality and, as a consequence, are present as enantiomers or diastereomers, it is possible to use both, the pure enantiomers and diastereomers and their mixtures, in the compositions according to the invention. If the PPO-inhibiting herbicides A and/or the herbicidal compounds B as described herein have ionizable functional groups, they can also be employed in the form of their agriculturally acceptable salts. Suitable are, in general, the salts of those cations and the acid addition salts of those acids whose cations and anions, respectively, have no adverse effect on the activity of the active compounds. Preferred cations are the ions of the alkali metals, preferably of lithium, sodium and potassium, of the alkaline earth metals, preferably of calcium and magnesium, and of the transition metals, preferably of manganese, copper, zinc and iron, further ammonium and substituted ammonium in which one to four hydrogen atoms are replaced by C₁-C₄-alkyl, hydroxy-C₁-C₄-alkyl, C₁-C₄-alkoxy-C₁-C₄-alkyl, hydroxy-C₁-C₄-alkoxy-C₁-C₄-alkyl, phenyl or benzyl, preferably ammonium, methylammonium, isopropylammonium, dimethylammonium, diisopropylammonium, trimethylammonium, heptylammonium, dodecylammonium, tetradecylammonium, tetramethylammonium, tetraethylammonium, tetrabutylammonium, 2-hydroxyethylammonium (olamine salt), 2-(2-hydroxyethyl-1-oxy)ethyl-1-ylammonium (diglycolamine salt), di(2-hydroxyethyl-1-yl)ammonium (diolamine salt), tris(2-hydroxyethyl)ammonium (trolamine salt), tris(2-hydroxypropyl)ammonium, benzyltrimethylammonium, benzyltriethylammonium, N,N,N-trimethylethanolammonium (choline salt), furthermore phosphonium ions, sulfonium ions, preferably tri(C₁-C₄-alkyl)sulfonium, such as trimethylsulfonium, and sulfoxonium ions, preferably tri(C₁-C₄-alkyl)sulfoxonium, and finally the salts of polybasic amines such as N,N-bis-(3-aminopropyl)methylamine and diethylenetriamine. Anions of useful acid addition salts are primarily chloride, bromide, fluoride, iodide, hydrogensulfate, methylsulfate, sulfate, dihydrogenphosphate, hydrogenphosphate, nitrate, bicarbonate, carbonate, hexafluorosilicate, hexafluorophosphate, benzoate and also the anions of C₁-C₄-alkanoic acids, preferably formate, acetate, propionate and butyrate.

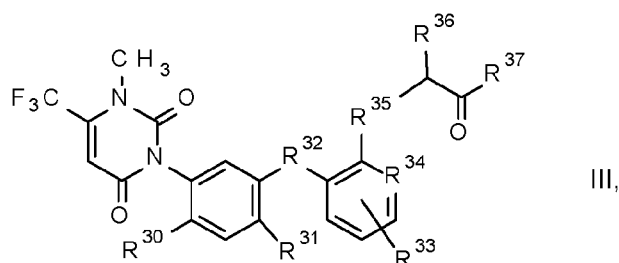
The PPO-inhibiting herbicides A and/or the herbicidal compounds B as described herein having a carboxyl group can be employed in the form of the acid, in the form of an agriculturally suitable salt as mentioned above or else in the form of an agriculturally acceptable derivative, for example as amides, such as mono- and di-C₁-C₆-alkylamides or arylamides, as esters, for example as allyl esters, propargyl esters, C₁-C₁₀-alkyl esters, alkoxyalkyl esters, tetrahydrofuran-2-ylmethyl esters and also as thioesters, for example as C₁-C₁₀-alkylthio esters. Preferred mono- and di-C₁-C₆-alkylamides are the methyl and the dimethylamides. Preferred arylamides are, for example, the anilides and the 2-chloroanilides. Preferred alkyl esters are, for example, the methyl, ethyl, propyl, isopropyl, butyl, isobutyl, pentyl, hexyl (1-methylhexyl), heptyl (1-methylheptyl), heptyl, octyl or isooctyl (2-ethylhexyl) esters. Preferred C₁-C₄-alkoxy-C₁-C₄-alkyl esters are the straight-chain or branched C₁-C₄-alkoxy ethyl esters, for example the 2-methoxyethyl, 2-ethoxyethyl, 2-butoxyethyl (butotyl), 2-butoxypropyl or 3-

butoxypropyl ester. An example of a straight-chain or branched C₁-C₁₀-alkylthio ester is the ethylthio ester.

Examples of PPO inhibiting herbicides which can be used according to the present invention

- 5 are acifluorfen, acifluorfen-sodium, aclonifen, azafenidin, bencarbazon, benzfendizon, bifenox, butafenacil, carfentrazone, carfentrazone-ethyl, chlomethoxyfen, cinidon-ethyl, fluazolate, flufenpyr, flufenpyr-ethyl, flumiclorac, flumiclorac-pentyl, flumioxazin, fluoroglycofen, fluoroglycofen-ethyl, fluthiacet, fluthiacet-methyl, fomesafen, halosafen, lactofen, oxadiargyl, oxadiazon, oxyfluorfen, pentoxazone, profluazol, pyraclonil, pyraflufen, pyraflufen-ethyl, saflufenacil, sulfentrazone, thidiazimin, tiafenacil, chlornitrofen, flumipropyn, fluoronitrofen, flupropacil, furyloxyfen, nitrofluorfen, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), N-ethyl-3-2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452098-92-9), N-tetrahydrofurfuryl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 915396-43-9), N-ethyl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452099-05-7), N-tetrahydrofurfuryl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452100-03-7), 3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2*H*-benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-[1,3,5]triazinan-2,4-dione (CAS 451484-50-7), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2*H*-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), 2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2*H*-benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isoindole-1,3-dione (CAS 1300118-96-0), 1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2*H*-benzo[1,4]oxazin-6-yl)-1*H*-pyrimidine-2,4-dione, methyl (*E*)-4-[2-chloro-5-[4-chloro-5-(difluoromethoxy)-1*H*-methyl-pyrazol-3-yl]-4-fluorophenoxy]-3-methoxy-but-2-enoate [CAS 948893-00-3], 3-[7-Chloro-5-fluoro-2-(trifluoromethyl)-1*H*-benzimidazol-4-yl]-1-methyl-6-(trifluoromethyl)-1*H*-pyrimidine-2,4-dione (CAS 212754-02-4), and

uracils of formula III



wherein

R³⁰ and R³¹ independently of one another are F, Cl or CN;

R³² is O or S;

R³³ is H, F, Cl, CH₃ or OCH₃;

R³⁴ is CH or N;

R³⁵ is O or S;

R³⁶ is H, CN, CH₃, CF₃, OCH₃, OC₂H₅, SCH₃, SC₂H₅, (CO)OC₂H₅ or CH₂R³⁸,

wherein R³⁸ is F, Cl, OCH₃, SCH₃, SC₂H₅, CH₂F, CH₂Br or CH₂OH;

and

R³⁷ is (C₁-C₆-alkyl)amino, (C₁-C₆-dialkyl)amino, (NH)OR³⁹, OH, OR⁴⁰ or SR⁴⁰

wherein R³⁹ is CH₃, C₂H₅ or phenyl; and

R⁴⁰ is independently of one another C₁-C₆-alkyl, C₂-C₆-alkenyl, C₃-C₆-alkynyl, C₁-C₆-haloalkyl, C₁-C₆-alkoxy-C₁-C₆-alkyl, C₁-C₆-alkoxy-C₁-C₆-alkoxy-C₁-C₆-alkyl, C₂-C₆-cyanoalkyl, C₁-C₄-alkoxy-carbonyl-C₁-C₄-alkyl, C₁-C₄-alkyl-carbonyl-amino, C₁-C₆-alkylsulfinyl-C₁-C₆-alkyl, C₁-C₆-alkyl-sulfonyl-C₁-C₆-alkyl, C₁-C₆-dialkoxo-C₁-C₆-alkyl, C₁-C₆-alkyl-carbonyloxy-C₁-C₆-alkyl, phenyl-carbonyl-C₁-C₆-alkyl, tri(C₁-C₃-alkyl)-silyl-C₁-C₆-alkyl, tri(C₁-C₃-alkyl)-silyl-C₁-C₆-alkenyl, tri(C₁-C₃-alkyl)-silyl-C₁-C₆-alkynyl, tri(C₁-C₃-alkyl)-silyl-C₁-C₆-alkoxy-C₁-C₆-alkyl, dimethylamino, tetrahydropyranyl, tetrahydrofuran-yl-C₁-C₃-alkyl, phenyl-C₁-C₆-alkoxy-C₁-C₆-alkyl, phenyl-C₁-C₃-alkyl, pyridyl-C₁-C₃-alkyl, pyridyl, phenyl,

which pyridyls and phenyls independently of one another are substituted by one to five substituents selected from the group consisting of halogen, C₁-C₃-alkyl or C₁-C₂-haloalkyl;

C₃-C₆-cycloalkyl or C₃-C₆-cycloalkyl-C₁-C₄-alkyl,

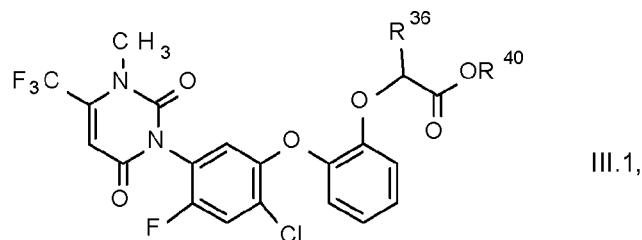
which cycloalkyls independently of one another are unsubstituted or substituted by one to five substituents selected from the group consisting of halogen, C₁-C₃-alkyl and C₁-C₂-haloalkyl;

including their agriculturally acceptable alkali metal salts or ammonium salts.

Preferred PPO-inhibiting herbicides that can be used according to the present invention are: Acifluorfen, acifluorfen-sodium, azafenidin, bencarbazone, benzfendizone, butafenacil, carfentrazone-ethyl, cinidon-ethyl, flufenpyr-ethyl, flumiclorac-pentyl, flumioxazin, fluoroglycofen-ethyl, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxadiazon, oxyfluorfen, pentoxazone, pyraflufen-ethyl, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]-acetate (CAS 353292-31-6; S-3100), N-ethyl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452098-92-9), N-tetrahydrofurfuryl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 915396-43-9), N-ethyl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452099-05-7), N-tetrahydrofurfuryl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452100-03-7), 3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2*H*-benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-[1,3,5]triazinan-2,4-dione (CAS 451484-50-7), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2*H*-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinan-2,4-dione (CAS 1258836-72-4), 2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2*H*-benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isoindole-1,3-dione (CAS 1300118-96-0); 1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2*H*-benzo[1,4]oxazin-6-yl)-1*H*-pyrimidine-2,4-dione (CAS 1304113-05-0), 3-[7-Chloro-5-fluoro-2-(trifluoromethyl)-1*H*-benzimidazol-4-yl]-1-methyl-6-(trifluoromethyl)-1*H*-pyrimidine-2,4-dione

(CAS 212754-02-4)

uracils of formula III.1 (corresponding to uracils of formula III, wherein R^{30} is F, R^{31} is Cl, R^{32} is O; R^{33} is H; R^{34} is CH; R^{35} is O and R^{37} is OR^{40})



wherein

R^{36} is OCH_3 , OC_2H_5 , SCH_3 or SC_2H_5 ;

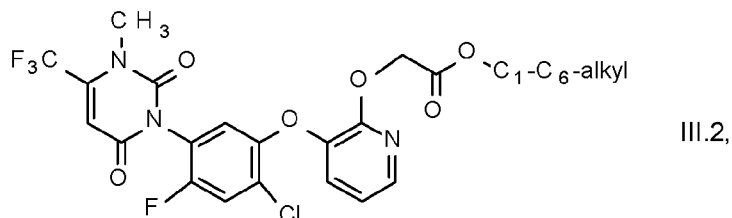
and

R^{40} is C_1 - C_6 -alkyl, C_2 - C_6 -alkenyl, C_3 - C_6 -alkynyl, C_1 - C_6 -haloalkyl, C_1 - C_6 -alkoxy- C_1 - C_6 -alkyl, C_1 - C_6 -alkoxy- C_1 - C_6 -alkoxy- C_1 - C_6 -alkyl, C_1 - C_3 -cyanoalkyl, phenyl- C_1 - C_3 -alkyl, pyridyl- C_1 - C_3 -alkyl, C_3 - C_6 -cycloalkyl or C_3 - C_6 -cycloalkyl- C_1 - C_4 -alkyl,

which cycloalkyls are unsubstituted or substituted by one to five substituents selected from the group consisting of halogen, C_1 - C_3 -alkyl and C_1 - C_2 -haloalkyl;

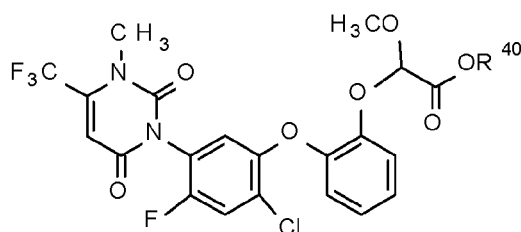
and

uracils of formula III.2 (corresponding to uracils of formula III, wherein R^{30} is F; R^{31} is Cl; R^{32} is O; R^{33} is H; R^{34} is N; R^{35} is O and R^{37} is OR^{40} with R^{40} is C_1 - C_6 -alkyl)



Particularly preferred PPO-inhibiting herbicides that can be used according to the present invention are:

acifluorfen, acifluorfen-sodium, butafenacil, carfentrazone-ethyl, cinidon-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)-phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-[1,3,5]triazinan-2,4-dione (CAS 451484-50-7), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), and 2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isoindole-1,3-dione (CAS 1300118-96-0), 1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-1H-pyrimidine-2,4-dione (CAS 1304113-05-0), uracils of formula III.1.1 (corresponding to uracils of formula III, wherein R^{30} is F, R^{31} is Cl, R^{32} is O; R^{33} is H; R^{34} is CH; R^{35} is O, R^{36} is OCH_3 and R^{37} is OR^{40})

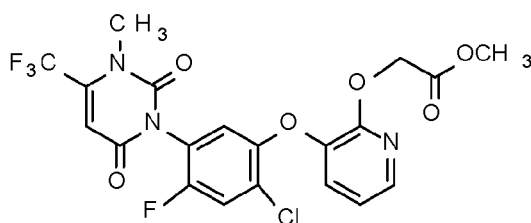


III.1.1,

wherein

R^{40} is C_1 - C_6 -alkyl, C_2 - C_6 -alkenyl, C_3 - C_6 -alkynyl, C_1 - C_6 -haloalkyl, C_1 - C_6 -alkoxy- C_1 - C_6 -alkyl, C_1 - C_6 -alkoxy- C_1 - C_6 -alkoxy- C_1 - C_6 -alkyl, C_1 - C_3 -cyanoalkyl, phenyl- C_1 - C_3 -alkyl, pyridyl- C_1 - C_3 -alkyl, C_3 - C_6 -cycloalkyl or C_3 - C_6 -cycloalkyl- C_1 - C_4 -alkyl, which cycloalkyls are unsubstituted or substituted by one to five substituents selected from the group consisting of halogen, C_1 - C_3 -alkyl and C_1 - C_2 -haloalkyl; is preferably CH_3 , $CH_2CH_2OC_2H_5$, CH_2CHF_2 , cyclohexyl, (1-methylcyclopropyl)methyl or CH_2 (pyridine-4-yl);

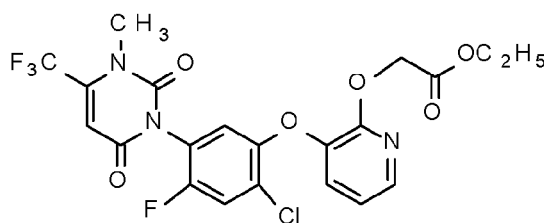
uracils of formula III.2.1 (corresponding to uracils of formula III, wherein R^{30} is F; R^{31} is Cl; R^{32} is O; R^{33} is H; R^{34} is N; R^{35} is O and R^{37} is OR^{40} with R^{40} is CH_3)



III.2.1,

and

uracils of formula III.2.2 (corresponding to uracils of formula III, wherein R^{30} is F; R^{31} is Cl; R^{32} is O; R^{33} is H; R^{34} is N; R^{35} is O and R^{37} is OR^{40} with R^{40} is C_2H_5)



III.2.2,

Especially preferred PPO-inhibiting herbicides are the PPO-inhibiting herbicides.1 to A.14 listed below in table A:

Table A

A.1	acifluorfen
A.2	butafenacil
A.3	carfentrazone-ethyl
A.4	cinidon-ethyl
A.5	flumioxazin
A.6	fluthiacet-methyl

A.7	fomesafen
A.8	lactofen
A.9	oxadiargyl
A.10	oxyfluorfen
A.11	saflufenacil
A.12	sulfentrazone
A.13	ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6)
A.14	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4)

The PPO-inhibiting herbicides described above that are useful to carry out the present invention are often best applied in conjunction with one or more other herbicides to obtain control of a wider variety of undesirable vegetation. For example, PPO-inhibiting herbicides may further be used in conjunction with additional herbicides to which the crop plant is naturally tolerant, or to which it is resistant via expression of one or more additional transgenes as mentioned supra, or to which it is resistant via mutagenesis and breeding methods as described hereinafter. When used in conjunction with other targeting herbicides, the PPO-inhibiting herbicides, to which the plant of the present invention had been made resistant or tolerant, can be formulated with the other herbicide or herbicides, tank mixed with the other herbicide or herbicides, or applied sequentially with the other herbicide or herbicides.

Suitable components for mixtures are, for example, selected from the herbicides of class b1) to b15)

B) herbicides of class b1) to b15):

- b1) lipid biosynthesis inhibitors;
- b2) acetolactate synthase inhibitors (ALS inhibitors);
- b3) photosynthesis inhibitors;
- b4) protoporphyrinogen-IX oxidase inhibitors,
- b5) bleacher herbicides;
- b6) enolpyruvyl shikimate 3-phosphate synthase inhibitors (EPSP inhibitors);
- b7) glutamine synthetase inhibitors;
- b8) 7,8-dihydropteroate synthase inhibitors (DHP inhibitors);
- b9) mitosis inhibitors;
- b10) inhibitors of the synthesis of very long chain fatty acids (VLCFA inhibitors);
- b11) cellulose biosynthesis inhibitors;
- b12) decoupler herbicides;
- b13) auxinic herbicides;
- b14) auxin transport inhibitors; and
- b15) other herbicides selected from the group consisting of bromobutide, chlorflurenol, chlorflurenol-methyl, cinmethylin, cumyluron, dalapon, dazomet, difenzoquat, difenzoquat-metilsulfate, dimethipin, DSMA, dymron, endothal and its salts, etobenzanid, flumprop, flumprop-isopropyl, flumprop-methyl, flumprop-M-

isopropyl, flamprop-M-methyl, flurenol, flurenol-butyl, flurprimidol, fosamine, fosamine-ammonium, indanofan, indaziflam, maleic hydrazide, mefluidide, metam, methiozolin (CAS 403640-27-7), methyl azide, methyl bromide, methyl-dymron, methyl iodide, MSMA, oleic acid, oxaziclomefone, pelargonic acid, pyributicarb, quinoclamine, triaziflam, tridiphane and 6-chloro-3-(2-cyclopropyl-6-methylphenoxy)-4-pyridazinol (CAS 499223-49-3) and its salts and esters;

including their agriculturally acceptable salts or derivatives.

- 10 Examples of herbicides B which can be used in combination with the PPO-inhibiting herbicides according to the present invention are:

b1) from the group of the lipid biosynthesis inhibitors:

- ACC-herbicides such as alloxydim, alloxydim-sodium, butroxydim, clethodim, clodinafop, clodinafop-propargyl, cycloxydim, cyhalofop, cyhalofop-butyl, diclofop, diclofop-methyl, fenoxaprop, fenoxaprop-ethyl, fenoxaprop-P, fenoxaprop-P-ethyl, fluazifop, fluazifop-butyl, fluazifop-P, fluazifop-P-butyl, haloxyfop, haloxyfop-methyl, haloxyfop-P, haloxyfop-P-methyl, metamifop, pinoxaden, profoxydim, propaquizafop, quizalofop, quizalofop-ethyl, quizalofop-tefuryl, quizalofop-P, quizalofop-P-ethyl, quizalofop-P-tefuryl, sethoxydim, tepraloxym, tralkoxydim,
- 4-(4'-Chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1312337-72-6); 4-(2',4'-Dichloro-4-cyclopropyl[1,1'-biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1312337-45-3); 4-(4'-Chloro-4-ethyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1033757-93-5); 4-(2',4'-Dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-2,2,6,6-tetramethyl-2H-pyran-3,5(4H,6H)-dione (CAS 1312340-84-3); 5-(Acetyloxy)-4-(4'-chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1312337-48-6); 5-(Acetyloxy)-4-(2',4'-dichloro-4-cyclopropyl- [1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one; 5-(Acetyloxy)-4-(4'-chloro-4-ethyl-2'-fluoro[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1312340-82-1); 5-(Acetyloxy)-4-(2',4'-dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1033760-55-2); 4-(4'-Chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1312337-51-1); 4-(2',4'-Dichloro -4-cyclopropyl- [1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester; 4-(4'-Chloro-4-ethyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1312340-83-2); 4-(2',4'-Dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1033760-58-5); and non ACC herbicides such as benfuresate, butylate, cycloate, dalapon, dimepiperate, EPTC, esprocarb, ethofumesate, flupropanate, molinate, orbencarb, pebulate, prosulfocarb, TCA, thiobencarb, tiocarbazil, triallate and vernolate;

b2) from the group of the ALS inhibitors:

sulfonylureas such as amidosulfuron, azimsulfuron, bensulfuron, bensulfuron-methyl, chlorimuron, chlorimuron-ethyl, chlorsulfuron, cinosulfuron, cyclosulfamuron, ethametsulfuron,

- ethametsulfuron-methyl, ethoxysulfuron, flazasulfuron, flucetosulfuron, flupyrsulfuron, flupyrsulfuron-methyl-sodium, foramsulfuron, halosulfuron, halosulfuron-methyl, imazosulfuron, iodosulfuron, iodosulfuron-methyl-sodium, iofensulfuron, iofensulfuron-sodium, mesosulfuron, metazosulfuron, metsulfuron, metsulfuron-methyl, nicosulfuron, orthosulfamuron, oxasulfuron,
- 5 primisulfuron, primisulfuron-methyl, propyrisulfuron, prosulfuron, pyrazosulfuron, pyrazosulfuron-ethyl, rimsulfuron, sulfometuron, sulfometuron-methyl, sulfosulfuron, thifensulfuron, thifensulfuron-methyl, triasulfuron, tribenuron, tribenuron-methyl, trifloxysulfuron, triflusulfuron, triflusulfuron-methyl and tritosulfuron,
- imidazolinones such as imazamethabenz, imazamethabenz-methyl, imazamox, imazapic,
- 10 imazapyr, imazaquin and imazethapyr, triazolopyrimidine herbicides and sulfonanilides such as cloransulam, cloransulam-methyl, diclosulam, flumetsulam, florasulam, metosulam, penoxsulam, pyrimisulfan and pyroxsulam,
- pyrimidinylbenzoates such as bispyribac, bispyribac-sodium, pyribenzoxim, pyriftalid, pyriminobac, pyriminobac-methyl, pyrithiobac, pyrithiobac-sodium, 4-[[[2-[(4,6-dimethoxy-2-
- 15 pyrimidinyl)oxy]phenyl]methyl]amino]-benzoic acid-1-methylethyl ester (CAS 420138-41-6), 4-[[[2-[(4,6-dimethoxy-2-pyrimidinyl)oxy]phenyl]methyl]amino]-benzoic acid propyl ester (CAS 420138-40-5), N-(4-bromophenyl)-2-[(4,6-dimethoxy-2-pyrimidinyl)oxy]benzenemethanamine (CAS 420138-01-8),
- sulfonylaminocarbonyl-triazolinone herbicides such as flucarbazone, flucarbazone-sodium,
- 20 propoxycarbazone, propoxycarbazone-sodium, thienencarbazone and thienencarbazone-methyl; and triafamone;
- among these, a preferred embodiment of the invention relates to those compositions comprising at least one imidazolinone herbicide;
- 25 b3) from the group of the photosynthesis inhibitors:
- amicarbazone, inhibitors of the photosystem II, e.g. triazine herbicides, including of chlorotriazine, triazinones, triazindiones, methylthiotriazines and pyridazinones such as ametryn, atrazine, chloridazone, cyanazine, desmetryn, dimethametryn, hexazinone, metribuzin, prometon, prometryn, propazine, simazine, simetryn, terbumeton, terbuthylazin, terbutryn and
- 30 trietazin, aryl urea such as chlorobromuron, chlorotoluron, chloroxuron, dimefuron, diuron, fluometuron, isoproturon, isouron, linuron, metamitron, methabenzthiazuron, metobenzuron, metoxuron, monolinuron, neburon, siduron, tebuthiuron and thiadiazuron, phenyl carbamates such as desmedipham, karbutilat, phenmedipham, phenmedipham-ethyl, nitrile herbicides such as bromofenoxim, bromoxynil and its salts and esters, ioxynil and its salts and esters, uraciles
- 35 such as bromacil, lenacil and terbacil, and bentazon and bentazon-sodium, pyridate, pyridafol, pentanochlor and propanil and inhibitors of the photosystem I such as diquat, diquat-dibromide, paraquat, paraquat-dichloride and paraquat-dimetilsulfate. Among these, a preferred embodiment of the invention relates to those compositions comprising at least one aryl urea herbicide. Among these, likewise a preferred embodiment of the invention relates to those
- 40 compositions comprising at least one triazine herbicide. Among these, likewise a preferred embodiment of the invention relates to those compositions comprising at least one nitrile herbicide;

b4) from the group of the protoporphyrinogen-IX oxidase inhibitors:

acifluorfen, acifluorfen-sodium, azafenidin, bencarbazone, benzfendizone, bifenox, butafenacil, carfentrazone, carfentrazone-ethyl, chlomethoxyfen, cinidon-ethyl, fluazolate, flufenpyr, flufenpyr-ethyl, flumiclorac, flumiclorac-pentyl, flumioxazin, fluoroglycofen, fluoroglycofen-ethyl, fluthiacet, fluthiacet-methyl, fomesafen, halosafen, lactofen, oxadiargyl, oxadiazon, oxyfluorfen, pentoxazone, proflumazone, pyraclonil, pyraflufen, pyraflufen-ethyl, saflufenacil, sulfentrazone, thidiazimin, tienfenacil, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100, N-ethyl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 452098-92-9), N-tetrahydrofurfuryl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 915396-43-9), N-ethyl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 452099-05-7), N-tetrahydrofurfuryl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 452100-03-7), 3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-[1,3,5]triazinan-2,4-dione, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), 2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isindole-1,3-dione, 1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-1H-pyrimidine-2,4-dione (CAS 1304113-05-0), methyl (E)-4-[2-chloro-5-[4-chloro-5-(difluoromethoxy)-1H-methyl-pyrazol-3-yl]-4-fluoro-phenoxy]-3-methoxy-but-2-enoate [CAS 948893-00-3], and 3-[7-Chloro-5-fluoro-2-(trifluoromethyl)-1H-benzimidazol-4-yl]-1-methyl-6-(trifluoromethyl)-1H-pyrimidine-2,4-dione (CAS 212754-02-4);

b5) from the group of the bleacher herbicides:

PDS inhibitors: beflubutamid, diflufenican, fluridone, flurochloridone, flurtamone, norflurazon, picolinafen, and 4-(3-trifluoromethylphenoxy)-2-(4-trifluoromethylphenyl)pyrimidine (CAS 180608-33-7), HPPD inhibitors: benzobicyclon, benzofenap, clomazone, isoxaflutole, mesotrione, pyrasulfotole, pyrazolynate, pyrazoxyfen, sulcotrione, tefuryltrione, tembotrione, topramezone and bicyclopyrone, bleacher, unknown target: aclonifen, amitrole and flumeturon;

b6) from the group of the EPSP synthase inhibitors:

glyphosate, glyphosate-isopropylammonium, glyposate-potassium and glyphosate-trimesium (sulfosate);

b7) from the group of the glutamine synthase inhibitors:

bilanaphos (bialaphos), bilanaphos-sodium, glufosinate, glufosinate-P and glufosinate-ammonium;

b8) from the group of the DHP synthase inhibitors:

asulam;

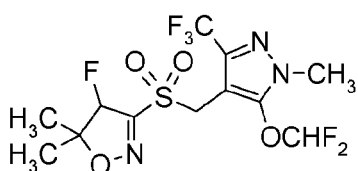
b9) from the group of the mitosis inhibitors:

compounds of group K1: dinitroanilines such as benfluralin, butralin, dinitramine, ethalfluralin, fluchloralin, oryzalin, pendimethalin, prodiamine and trifluralin, phosphoramidates such as amiprofos, amiprofos-methyl, and butamipfos, benzoic acid herbicides such as chlorthal,

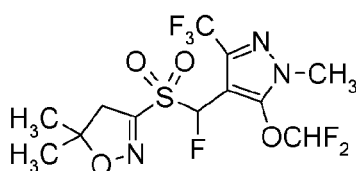
chlorthal-dimethyl, pyridines such as dithiopyr and thiazopyr, benzamides such as propyzamide and tebutam; compounds of group K2: chlorpropham, propham and carbetamide, among these, compounds of group K1, in particular dinitroanilines are preferred;

5 b10) from the group of the VLCFA inhibitors:

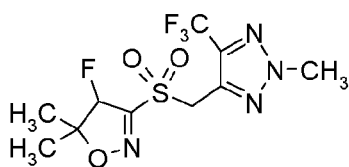
chloroacetamides such as acetochlor, alachlor, butachlor, dimethachlor, dimethenamid, dimethenamid-P, metazachlor, metolachlor, metolachlor-S, pethoxamid, pretilachlor, propachlor, propisochlor and thenylchlor, oxyacetanilides such as flufenacet and mefenacet, acetanilides
 10 such as diphenamid, naproanilide and napropamide, tetrazolinones such fentrazamide, and other herbicides such as anilofos, cafenstrole, fenoxasulfone, ipfencarbazone, piperophos, pyroxasulfone and isoxazoline compounds of the formulae II.1, II.2, II.3, II.4, II.5, II.6, II.7, II.8 and II.9



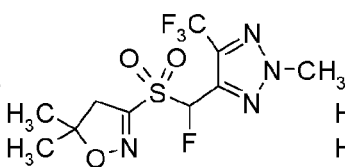
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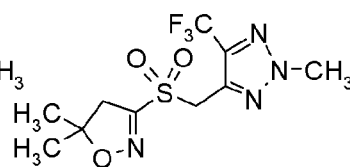
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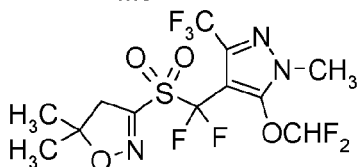
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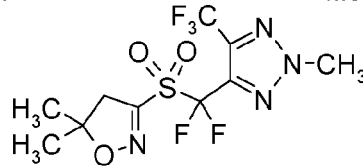
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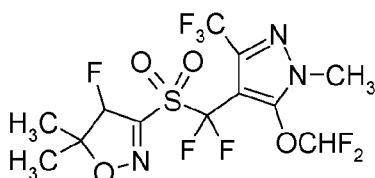
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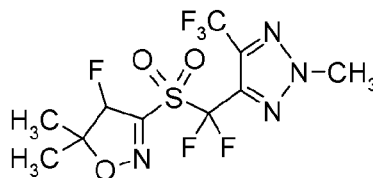
II.6



II.7



II.8



II.9

the isoxazoline compounds of the formula (I) are known in the art, e.g. from WO 2006/024820, WO 2006/037945, WO 2007/071900 and WO 2007/096576;

among the VLCFA inhibitors, preference is given to chloroacetamides and oxyacetamides;

b11) from the group of the cellulose biosynthesis inhibitors:

chlorthiamid, dichlobenil, flupoxam, indaziflam, triaziflam, isoxaben and 1-Cyclohexyl-5-pentafluorophenoxy-1⁴-[1,2,4,6]thiatriazin-3-ylamine;

b12) from the group of the decoupler herbicides:

5 dinoseb, dinoterb and DNOC and its salts;

b13) from the group of the auxinic herbicides:

2,4-D and its salts and esters such as clacyfos, 2,4-DB and its salts and esters,

10 aminocyclopyrachlor and its salts and esters, aminopyralid and its salts such as aminopyralid-tris(2-hydroxypropyl)ammonium and its esters, benazolin, benazolin-ethyl, chloramben and its salts and esters, clomeprop, clopyralid and its salts and esters, dicamba and its salts and

esters, dichlorprop and its salts and esters, dichlorprop-P and its salts and esters, fluroxypyr, fluroxypyr-butometyl, fluroxypyr-meptyl, halauxifen and its salts and esters (CAS 943832-60-8); MCPA and its salts and esters, MCPA-thioethyl, MCPB and its salts and esters, mecoprop and
15 its salts and esters, mecoprop-P and its salts and esters, picloram and its salts and esters, quinclorac, quinmerac, TBA (2,3,6) and its salts and esters and triclopyr and its salts and esters;

b14) from the group of the auxin transport inhibitors: diflufenzopyr, diflufenzopyr-sodium, naptalam and naptalam-sodium;

20

b15) from the group of the other herbicides: bromobutide, chlorflurenol, chlorflurenol-methyl, cinmethylin, cumyluron, cyclopyrimorate (CAS 499223-49-3) and its salts and esters, dalapon, dazomet, difenzoquat, difenzoquat-metilsulfate, dimethipin, DSMA, dymron, endothal and its salts, etobenzanid, flamprop, flamprop-isopropyl, flamprop-methyl, flamprop-M-isopropyl,
25 flamprop-M-methyl, flurenol, flurenol-butyl, flurprimidol, fosamine, fosamine-ammonium, indanofan, indaziflam, maleic hydrazide, mefluidide, metam, methiozolin (CAS 403640-27-7), methyl azide, methyl bromide, methyl-dymron, methyl iodide, MSMA, oleic acid, oxaziclomefone, pelargonic acid, pyributicarb, quinoclamine, triaziflam and tridiphane..

30 Preferred herbicides B that can be used in combination with the PPO-inhibiting herbicides according to the present invention are:

b1) from the group of the lipid biosynthesis inhibitors:

35 clethodim, clodinafop-propargyl, cycloxydim, cyhalofop-butyl, diclofop-methyl, fenoxaprop-P-ethyl, fluazifop-P-butyl, haloxyfop-P-methyl, metamifop, pinoxaden, profoxydim, propaquizafop, quizalofop-P-ethyl, quizalofop-P-tefuryl, sethoxydim, tepraloxym, tralkoxydim, 4-(4'-Chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1312337-72-6); 4-(2',4'-Dichloro-4-cyclopropyl[1,1'-biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1312337-45-3); 4-(4'-Chloro-4-ethyl-2'-fluoro[1,1'-
40 biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1033757-93-5); 4-(2',4'-Dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-2,2,6,6-tetramethyl-2H-pyran-3,5(4H,6H)-dione (CAS 1312340-84-3); 5-(Acetyloxy)-4-(4'-chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1312337-48-6); 5-(Acetyloxy)-4-(2',4'-dichloro-4-cyclopropyl-[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one; 5-

- (Acetyloxy)-4-(4'-chloro-4-ethyl-2'-fluoro[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1312340-82-1); 5-(Acetyloxy)-4-(2',4'-dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1033760-55-2); 4-(4'-Chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1312337-51-1); 4-(2',4'-Dichloro-4-cyclopropyl-[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester; 4-(4'-Chloro-4-ethyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1312340-83-2); 4-(2',4'-Dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1033760-58-5);
- 10 benfuresate, dimepiperate, EPTC, esprocarb, ethofumesate, molinate, orbencarb, prosulfocarb, thiobencarb and triallate;

b2) from the group of the ALS inhibitors:

- amidosulfuron, azimsulfuron, bensulfuron-methyl, bispyribac-sodium, chlorimuron-ethyl,
- 15 chlorsulfuron, cloransulam-methyl, cyclosulfamuron, diclosulam, ethametsulfuron-methyl, ethoxysulfuron, flazasulfuron, florasulam, flucarbazone-sodium, flucetosulfuron, flumetsulam, flupyrsulfuron-methyl-sodium, foramsulfuron, halosulfuron-methyl, imazamethabenz-methyl, imazamox, imazapic, imazapyr, imazaquin, imazethapyr, imazosulfuron, iodosulfuron, iodosulfuron-methyl-sodium, iofensulfuron, iofensulfuron-sodium, mesosulfuron,
- 20 metazosulfuron, metosulam, metsulfuron-methyl, nicosulfuron, orthosulfamuron, oxasulfuron, penoxsulam, primisulfuron-methyl, propoxycarbazone-sodium, propyrisulfuron, prosulfuron, pyrazosulfuron-ethyl, pyribenzoxim, pyrimisulfan, pyriftalid, pyriminobac-methyl, pyriithiobac-sodium, pyroxsulam, rimsulfuron, sulfometuron-methyl, sulfosulfuron, thiencarbazone-methyl, thifensulfuron-methyl, triasulfuron, tribenuron-methyl, trifloxysulfuron, triflusulfuron-methyl,
- 25 tritosulfuron and triafamone;

b3) from the group of the photosynthesis inhibitors:

- ametryn, amicarbazone, atrazine, bentazone, bentazone-sodium, bromoxynil and its salts and esters, chloridazone, chlorotoluron, cyanazine, desmedipham, diquat-dibromide, diuron,
- 30 fluometuron, hexazinone, ioxynil and its salts and esters, isoproturon, lenacil, linuron, metamitron, methabenzthiazuron, metribuzin, paraquat, paraquat-dichloride, phenmedipham, propanil, pyridate, simazine, terbutryn, terbuthylazine and thidiazuron;

b4) from the group of the protoporphyrinogen-IX oxidase inhibitors:

- 35 acifluorfen, acifluorfen-sodium, azafenidin, bencarbazone, benzfendizone, butafenacil, carfentrazone-ethyl, cinidon-ethyl, flufenpyr-ethyl, flumiclorac-pentyl, flumioxazin, fluoroglycofen-ethyl, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxadiazon, oxyfluorfen, pentoxazone, pyraflufen-ethyl, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]-acetate (CAS 353292-31-6; S-3100), N-ethyl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 452098-92-9), N-tetrahydrofurfuryl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 915396-43-9), N-ethyl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 452099-05-7), N-tetrahydrofurfuryl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-
- 40

- 1-carboxamide (CAS 452100-03-7), 3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-[1,3,5]triazinan-2,4-dione, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), 2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isoindole-1,3-dione ; 1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-1H-pyrimidine-2,4-dione, and 3-[7-Chloro-5-fluoro-2-(trifluoromethyl)-1H-benzimidazol-4-yl]-1-methyl-6-(trifluoromethyl)-1H-pyrimidine-2,4-dione (CAS 212754-02-4);
- 5
- 10 b5) from the group of the bleacher herbicides:
aclonifen, beflubutamid, benzobicyclon, clomazone, diflufenican, flurochloridone, flurtamone, isoxaflutole, mesotrione, norflurazon, picolinafen, pyrasulfotole, pyrazolynate, sulcotrione, tefuryltrione, tembotrione, topramezone, bicycloprrone, 4-(3-trifluoromethylphenoxy)-2-(4-trifluoromethylphenyl)pyrimidine (CAS 180608-33-7), amitrole and flumeturon;
- 15
- b6) from the group of the EPSP synthase inhibitors:
glyphosate, glyphosate-isopropylammonium, glyphosate-potassium and glyphosate-trimesium (sulfosate);
- 20 b7) from the group of the glutamine synthase inhibitors:
glufosinate, glufosinate-P, glufosinate-ammonium;
- b8) from the group of the DHP synthase inhibitors: asulam;
- 25 b9) from the group of the mitosis inhibitors:
benfluralin, dithiopyr, ethalfluralin, oryzalin, pendimethalin, thiazopyr and trifluralin;
- b10) from the group of the VLCFA inhibitors:
acetochlor, alachlor, anilofos, butachlor, cafenstrole, dimethenamid, dimethenamid-P,
- 30 fentrazamide, flufenacet, mefenacet, metazachlor, metolachlor, S-metolachlor, naproanilide, napropamide, pretilachlor, fenoxasulfone, ipfencarbazone, pyroxasulfone thenylchlor and isoxazoline-compounds of the formulae II.1, II.2, II.3, II.4, II.5, II.6, II.7, II.8 and II.9 as mentioned above;
- 35 b11) from the group of the cellulose biosynthesis inhibitors: dichlobenil, flupoxam, isoxaben and 1-Cyclohexyl-5-pentafluorophenyloxy-1⁴-[1,2,4,6]thiatriazin-3-ylamine;
- b13) from the group of the auxinic herbicides:
2,4-D and its salts and esters, aminocyclopyrachlor and its salts and esters, aminopyralid and its salts such as aminopyralid-tris(2-hydroxypropyl)ammonium and its esters, clopyralid and its salts and esters, dicamba and its salts and esters, dichlorprop-P and its salts and esters, fluroxypyr-meptyl, halauxifen and its salts and esters (CAS 943832-60-8), MCPA and its salts and esters, MCPB and its salts and esters, mecoprop-P and its salts and esters, picloram and its salts and esters, quinclorac, quinmerac and triclopyr and its salts and esters;
- 40

b14) from the group of the auxin transport inhibitors: diflufenzopyr and diflufenzopyr-sodium;

b15) from the group of the other herbicides: bromobutide, cinmethylin, cumyluron, cyclopyrimorate (CAS 499223-49-3) and its salts and esters, dalapon, difenzoquat, difenzoquat-metilsulfate, DSMA, dymron (= daimuron), flamprop, flamprop-isopropyl, flamprop-methyl, flamprop-M-isopropyl, flamprop-M-methyl, indanofan, indaziflam, metam, methylbromide, MSMA, oxaziclomefone, pyributicarb, triaziflam and tridiphane.

Particularly preferred herbicides B that can be used in combination with the PPO-inhibiting herbicides according to the present invention are:

10

b1) from the group of the lipid biosynthesis inhibitors: clodinafop-propargyl, cycloxydim, cyhalofop-butyl, fenoxaprop-P-ethyl, pinoxaden, profoxydim, tepraloxym, tralkoxydim, 4-(4'-Chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1312337-72-6); 4-(2',4'-Dichloro-4-cyclopropyl[1,1'-biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1312337-45-3); 4-(4'-Chloro-4-ethyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5-hydroxy-2,2,6,6-tetramethyl-2H-pyran-3(6H)-one (CAS 1033757-93-5); 4-(2',4'-Dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-2,2,6,6-tetramethyl-2H-pyran-3,5(4H,6H)-dione (CAS 1312340-84-3); 5-(Acetyloxy)-4-(4'-chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1312337-48-6); 5-(Acetyloxy)-4-(2',4'-dichloro-4-cyclopropyl- [1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one; 5-(Acetyloxy)-4-(4'-chloro-4-ethyl-2'-fluoro[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1312340-82-1); 5-(Acetyloxy)-4-(2',4'-dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-3,6-dihydro-2,2,6,6-tetramethyl-2H-pyran-3-one (CAS 1033760-55-2); 4-(4'-Chloro-4-cyclopropyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1312337-51-1); 4-(2',4'-Dichloro -4-cyclopropyl- [1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester; 4-(4'-Chloro-4-ethyl-2'-fluoro[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1312340-83-2); 4-(2',4'-Dichloro-4-ethyl[1,1'-biphenyl]-3-yl)-5,6-dihydro-2,2,6,6-tetramethyl-5-oxo-2H-pyran-3-yl carbonic acid methyl ester (CAS 1033760-58-5);

30 esprocarb, prosulfocarb, thiobencarb and triallate;

b2) from the group of the ALS inhibitors: bensulfuron-methyl, bispyribac-sodium, cyclosulfamuron, diclosulam, flumetsulam, flupyralsulfuron-methyl-sodium, foramsulfuron, imazamox, imazapic, imazapyr, imazaquin, imazethapyr, imazosulfuron, iodosulfuron, iodosulfuron-methyl-sodium, iofensulfuron, iofensulfuron-sodium, mesosulfuron, metazosulfuron, nicosulfuron, penoxsulam, propoxycarbazone-sodium, propyrisulfuron, pyrazosulfuron-ethyl, pyroxsulam, rimsulfuron, sulfosulfuron, thienencarbazone-methyl, tritosulfuron and triafamone;

35

b3) from the group of the photosynthesis inhibitors: ametryn, atrazine, diuron, fluometuron, hexazinone, isoproturon, linuron, metribuzin, paraquat, paraquat-dichloride, propanil, terbutryn and terbutylazine;

40

b4) from the group of the protoporphyrinogen-IX oxidase inhibitors: acifluorfen, acifluorfen-

sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-[1,3,5]triazinan-2,4-dione, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), and 2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isoindole-1,3-dione, and 1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-1H-pyrimidine-2,4-dione;

b5) from the group of the bleacher herbicides: clomazone, diflufenican, flurochloridone, isoxaflutole, mesotrione, picolinafen, sulcotrione, tefuryltrione, tembotrione, topramezone, bicyclopyrone, amitrole and flumeturon;

b6) from the group of the EPSP synthase inhibitors: glyphosate, glyphosate-isopropylammonium and glyphosate-trimesium (sulfosate);

b7) from the group of the glutamine synthase inhibitors: glufosinate, glufosinate-P and glufosinate-ammonium;

b9) from the group of the mitosis inhibitors: pendimethalin and trifluralin;

b10) from the group of the VLCFA inhibitors: acetochlor, cafenstrole, dimethenamid-P, fentrazamide, flufenacet, mefenacet, metazachlor, metolachlor, S-metolachlor, fenoxasulfone, ipfencarbazone and pyroxasulfone; likewise, preference is given to isoxazoline compounds of the formulae II.1, II.2, II.3, II.4, II.5, II.6, II.7, II.8 and II.9 as mentioned above;

b11) from the group of the cellulose biosynthesis inhibitors: isoxaben;

b13) from the group of the auxinic herbicides: 2,4-D and its salts and esters such as clacyfos, and aminocyclopyrachlor and its salts and esters, aminopyralid and its salts and its esters, clopyralid and its salts and esters, dicamba and its salts and esters, fluroxypyr-meptyl, quinclorac and quinmerac;

b14) from the group of the auxin transport inhibitors: diflufenzopyr and diflufenzopyr-sodium,

b15) from the group of the other herbicides: dymron (= daimuron), indanofan, indaziflam, oxaziclomefone and triaziflam.

Moreover, it may be useful to apply the PPO-inhibiting herbicides, when used in combination with a compound B described SUPRA, in combination with safeners. Safeners are chemical compounds which prevent or reduce damage on useful plants without having a major impact on the herbicidal action of herbicides towards unwanted plants. They can be applied either before

sowings (e.g. on seed treatments, shoots or seedlings) or in the pre-emergence application or post-emergence application of the useful plant.

Furthermore, the safeners C, the PPO-inhibiting herbicides and/or the herbicides B can be applied simultaneously or in succession.

Suitable safeners are e.g. (quinolin-8-oxy)acetic acids, 1-phenyl-5-haloalkyl-1H-1,2,4-triazol-3-carboxylic acids, 1-phenyl-4,5-dihydro-5-alkyl-1H-pyrazol-3,5-dicarboxylic acids, 4,5-dihydro-5,5-diaryl-3-isoxazol carboxylic acids, dichloroacetamides, alpha-oximinophenylacetoneitriles, acetophenonoximes, 4,6-dihalo-2-phenylpyrimidines, N-[[4-(aminocarbonyl)phenyl]sulfonyl]-2-benzoic amides, 1,8-naphthalic anhydride, 2-halo-4-(haloalkyl)-5-thiazol carboxylic acids, phosphorothiolates and N-alkyl-O-phenylcarbamates and their agriculturally acceptable salts and their agriculturally acceptable derivatives such amides, esters, and thioesters, provided they have an acid group.

Examples of preferred safeners C are benoxacor, cloquintocet, cyometrinil, cyprosulfamide, dichlormid, dicyclonon, dietholate, fenchlorazole, fenclorim, flurazole, fluxofenim, furilazole, isoxadifen, mefenpyr, mephenate, naphthalic anhydride, oxabetrinil, 4-(dichloroacetyl)-1-oxa-4-azaspiro[4.5]decane (MON4660, CAS 71526-07-3) and 2,2,5-trimethyl-3-(dichloroacetyl)-1,3-oxazolidine (R-29148, CAS 52836-31-4).

Especially preferred safeners C are benoxacor, cloquintocet, cyprosulfamide, dichlormid, fenchlorazole, fenclorim, flurazole, fluxofenim, furilazole, isoxadifen, mefenpyr, naphthalic anhydride, oxabetrinil, 4-(dichloroacetyl)-1-oxa-4-azaspiro[4.5]decane (MON4660, CAS 71526-07-3) and 2,2,5-trimethyl-3-(dichloroacetyl)-1,3-oxazolidine (R-29148, CAS 52836-31-4).

Particularly preferred safeners C are benoxacor, cloquintocet, cyprosulfamide, dichlormid, fenchlorazole, fenclorim, furilazole, isoxadifen, mefenpyr, naphthalic anhydride, 4-(dichloroacetyl)-1-oxa-4-azaspiro[4.5]decane (MON4660, CAS 71526-07-3), and 2,2,5-trimethyl-3-(dichloroacetyl)-1,3-oxazolidine (R-29148, CAS 52836-31-4).

Also preferred safeners C are benoxacor, cloquintocet, cyprosulfamide, dichlormid, fenchlorazole, fenclorim, furilazole, isoxadifen, mefenpyr, 4-(dichloroacetyl)-1-oxa-4-azaspiro[4.5]decane (MON4660, CAS 71526-07-3) and 2,2,5-trimethyl-3-(dichloroacetyl)-1,3-oxazolidine (R-29148, CAS 52836-31-4)..

Particularly preferred safeners C, which, as component C, are constituent of the composition according to the invention are the safeners C as defined above; in particular the safeners C.1 - C.12 listed below in table C:

Table C

	Safener C
C.1	benoxacor
C.2	cloquintocet

	C.3 cyprosulfamide
	C.4 dichlormid
	C.5 fenchlorazole
	C.6 fenclorim
	C.7 furilazole
	C.8 isoxadifen
	C.9 mefenpyr
	C.10 naphtalic acid anhydride
	C.11 4-(dichloroacetyl)-1-oxa-4-azaspiro[4.5]decane (MON4660, CAS 71526-07-3)
	C.12 2,2,5-trimethyl-3-(dichloro-acetyl)-1,3-oxazolidine (R-29148, CAS 52836-31-4)

The PPO-inhibiting herbicides (compounds A) and the active compounds B of groups b1) to b15) and the active compounds C are known herbicides and safeners, see, for example, The Compendium of Pesticide Common Names (<http://www.alanwood.net/pesticides/>); Farm

- 5 Chemicals Handbook 2000 volume 86, Meister Publishing Company, 2000; B. Hock, C. Fedtke, R. R. Schmidt, Herbicide [Herbicides], Georg Thieme Verlag, Stuttgart 1995; W. H. Ahrens, Herbicide Handbook, 7th edition, Weed Science Society of America, 1994; and K. K. Hatzios, Herbicide Handbook, Supplement for the 7th edition, Weed Science Society of America, 1998. 2,2,5-Trimethyl-3-(dichloroacetyl)-1,3-oxazolidine [CAS No. 52836-31-4] is also referred to as R-
10 29148. 4-(Dichloroacetyl)-1-oxa-4-azaspiro[4.5]decane [CAS No. 71526-07-3] is also referred to as AD-67 and MON 4660.

The assignment of the active compounds to the respective mechanisms of action is based on current knowledge. If several mechanisms of action apply to one active compound, this
15 substance was only assigned to one mechanism of action.

Active compounds B and C having a carboxyl group can be employed in the form of the acid, in the form of an agriculturally suitable salt as mentioned above or else in the form of an agriculturally acceptable derivative in the compositions according to the invention.

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In the case of dicamba, suitable salts include those, where the counterion is an agriculturally acceptable cation. For example, suitable salts of dicamba are dicamba-sodium, dicamba-potassium, dicamba-methylammonium, dicamba-dimethylammonium, dicamba-isopropylammonium, dicamba-diglycolamine, dicamba-olamine, dicamba-diolamine, dicamba-trolamine, dicamba-N,N-bis-(3-aminopropyl)methylamine and dicamba-diethylenetriamine. Examples of a suitable ester are dicamba-methyl and dicamba-butotyl.

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Suitable salts of 2,4-D are 2,4-D-ammonium, 2,4-D-dimethylammonium, 2,4-D-diethylammonium, 2,4-D-diethanolammonium (2,4-D-diolamine), 2,4-D-triethanolammonium, 2,4-D-isopropylammonium, 2,4-D-triisopropanolammonium, 2,4-D-heptylammonium, 2,4-D-dodecylammonium, 2,4-D-tetradecylammonium, 2,4-D-triethylammonium, 2,4-D-tris(2-hydroxypropyl)ammonium, 2,4-D-tris(isopropyl)ammonium, 2,4-D-trolamine, 2,4-D-lithium, 2,4-
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D-sodium. Examples of suitable esters of 2,4-D are 2,4-D-butotyl, 2,4-D-2-butoxypropyl, 2,4-D-3-butoxypropyl, 2,4-D-butyl, 2,4-D-ethyl, 2,4-D-ethylhexyl, 2,4-D-isobutyl, 2,4-D-isooctyl, 2,4-D-isopropyl, 2,4-D-meptyl, 2,4-D-methyl, 2,4-D-octyl, 2,4-D-pentyl, 2,4-D-propyl, 2,4-D-tefuryl and clacyfos.

5 Suitable salts of 2,4-DB are for example 2,4-DB-sodium, 2,4-DB-potassium and 2,4-DB-dimethylammonium. Suitable esters of 2,4-DB are for example 2,4-DB-butyl and 2,4-DB-isooctyl. Suitable salts of dichlorprop are for example dichlorprop-sodium, dichlorprop-potassium and dichlorprop-dimethylammonium. Examples of suitable esters of dichlorprop are dichlorprop-butotyl and dichlorprop-isooctyl.

10 Suitable salts and esters of MCPA include MCPA-butotyl, MCPA-butyl, MCPA-dimethylammonium, MCPA-diolamine, MCPA-ethyl, MCPA-thioethyl, MCPA-2-ethylhexyl, MCPA-isobutyl, MCPA-isooctyl, MCPA-isopropyl, MCPA-isopropylammonium, MCPA-methyl, MCPA-olamine, MCPA-potassium, MCPA-sodium and MCPA-trolamine.

A suitable salt of MCPB is MCPB sodium. A suitable ester of MCPB is MCPB-ethyl.

15 Suitable salts of clopyralid are clopyralid-potassium, clopyralid-olamine and clopyralid-tris-(2-hydroxypropyl)ammonium. Example of suitable esters of clopyralid is clopyralid-methyl. Examples of a suitable ester of fluroxypyr are fluroxypyr-meptyl and fluroxypyr-2-butoxy-1-methylethyl, wherein fluroxypyr-meptyl is preferred.

20 Suitable salts of picloram are picloram-dimethylammonium, picloram-potassium, picloram-triisopropanolammonium, picloram-triisopropylammonium and picloram-trolamine. A suitable ester of picloram is picloram-isooctyl.

A suitable salt of triclopyr is triclopyr-triethylammonium. Suitable esters of triclopyr are for example triclopyr-ethyl and triclopyr-butotyl.

25 Suitable salts and esters of chloramben include chloramben-ammonium, chloramben-diolamine, chloramben-methyl, chloramben-methylammonium and chloramben-sodium. Suitable salts and esters of 2,3,6-TBA include 2,3,6-TBA-dimethylammonium, 2,3,6-TBA-lithium, 2,3,6-TBA-potassium and 2,3,6-TBA-sodium.

Suitable salts and esters of aminopyralid include aminopyralid-potassium and aminopyralid-tris(2-hydroxypropyl)ammonium.

30 Suitable salts of glyphosate are for example glyphosate-ammonium, glyphosate-diammonium, glyphosate-dimethylammonium, glyphosate-isopropylammonium, glyphosate-potassium, glyphosate-sodium, glyphosate-trimesium as well as the ethanolamine and diethanolamine salts, preferably glyphosate-diammonium, glyphosate-isopropylammonium and glyphosate-trimesium (sulfosate).

35 A suitable salt of glufosinate is for example glufosinate-ammonium.

A suitable salt of glufosinate-P is for example glufosinate-P-ammonium.

Suitable salts and esters of bromoxynil are for example bromoxynil-butyrate, bromoxynil-heptanoate, bromoxynil-octanoate, bromoxynil-potassium and bromoxynil-sodium.

40 Suitable salts and esters of ioxonil are for example ioxonil-octanoate, ioxonil-potassium and ioxonil-sodium.

Suitable salts and esters of mecoprop include mecoprop-butotyl, mecoprop-dimethylammonium, mecoprop-diolamine, mecoprop-ethadyl, mecoprop-2-ethylhexyl, mecoprop-isooctyl, mecoprop-methyl, mecoprop-potassium, mecoprop-sodium and mecoprop-trolamine.

Suitable salts of mecoprop-P are for example mecoprop-P-butotyl, mecoprop-P-

dimethylammonium, mecoprop-P-2-ethylhexyl, mecoprop-P-isobutyl, mecoprop-P-potassium and mecoprop-P-sodium.

A suitable salt of diflufenzopyr is for example diflufenzopyr-sodium.

A suitable salt of naptalam is for example naptalam-sodium.

- 5 Suitable salts and esters of aminocyclopyrachlor are for example aminocyclopyrachlor-dimethylammonium, aminocyclopyrachlor-methyl, aminocyclopyrachlor-triisopropanolammonium, aminocyclopyrachlor-sodium and aminocyclopyrachlor-potassium.

A suitable salt of quinclorac is for example quinclorac-dimethylammonium.

A suitable salt of quinmerac is for example quinmerac-dimethylammonium.

- 10 A suitable salt of imazamox is for example imazamox-ammonium.

Suitable salts of imazapic are for example imazapic-ammonium and imazapic-isopropylammonium.

Suitable salts of imazapyr are for example imazapyr-ammonium and imazapyr-isopropylammonium.

- 15 A suitable salt of imazaquin is for example imazaquin-ammonium.

Suitable salts of imazethapyr are for example imazethapyr-ammonium and imazethapyr-isopropylammonium.

A suitable salt of topramezone is for example topramezone-sodium.

- 20 The preferred embodiments of the invention mentioned herein below have to be understood as being preferred either independently from each other or in combination with one another.

According to a preferred embodiment of the invention, the composition comprises as component B at least one, preferably exactly one herbicide B.

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According to another preferred embodiment of the invention, the composition comprises at least two, preferably exactly two, herbicides B different from each other.

According to another preferred embodiment of the invention, the composition comprises at least three, preferably exactly three, herbicides B different from each other.

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According to another preferred embodiment of the invention, the composition comprises as component A at least one, preferably exactly one PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)-phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100; 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), and as component B at least one, preferably exactly one, herbicide B.

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According to another preferred embodiment of the invention, the composition comprises as

component A at least one, preferably exactly preferably exactly one PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), and at least two, preferably exactly two, herbicides B different from each other.

According to another preferred embodiment of the invention, the composition comprises as component A at least one, preferably exactly preferably exactly one PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4) and at least three, preferably exactly three, herbicides B different from each other.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b1), in particular selected from the group consisting of clethodim, clodinafop-propargyl, cycloxydim, cyhalofop-butyl, fenoxaprop-P-ethyl, fluazifop, pinoxaden, profoxydim, quizalofop, sethoxydim, tepraloxydim, tralkoxydim, esprocarb, prosulfocarb, thiobencarb and triallate.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4) especially

preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b2), in particular selected from the group consisting of bensulfuron-methyl, bispyribac-sodium, cloransulam-methyl,

5 cyclosulfamuron, diclosulam, flumetsulam, flupyrsulfuron-methyl-sodium, foramsulfuron, halosulfuron-methyl, imazamox, imazapic, imazapyr, imazaquin, imazethapyr, imazosulfuron, iodosulfuron, iodosulfuron-methyl-sodium, mesosulfuron-methyl, metazosulfuron, nicosulfuron, penoxsulam, propoxycarbazon-sodium, pyrazosulfuron-ethyl, pyriithiobac-sodium, pyroxsulam,

10 rimsulfuron, sulfosulfuron, thiencazuron-methyl, thifensulfuron-methyl, trifloxysulfuron and tritosulfuron.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen,

15 oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-

20 2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b3), in particular selected from the group consisting of ametryn, atrazine, bentazon, bromoxynil, diuron, fluometuron, hexazinone, isoproturon, linuron, metribuzin, paraquat, paraquat-dichloride, prometryne, propanil, terbutryn and terbutylazine.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-

30 trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and

35 especially exactly one herbicidally active compound from group b4), in particular selected from the group consisting of acifluorfen, acifluorfen-sodium, azafenidin, bencarbazone, benzfendazole, bifenox, butafenacil, carfentrazone, carfentrazone-ethyl, chlormethoxyfen, cinidon-ethyl, fluazolate, flufenpyr, flufenpyr-ethyl, flumiclorac, flumiclorac-pentyl, flumioxazin, fluoroglycofen, fluoroglycofen-ethyl, fluthiacet, fluthiacet-methyl, fomesafen, halosafen, lactofen,

40 oxadiargyl, oxadiazon, oxyfluorfen, pentoxazone, proflumazone, pyraclonil, pyraflufen, pyraflufen-ethyl, saflufenacil, sulfentrazone, thidiazimin, tiafenacil, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), N-ethyl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 452098-92-9), N-tetrahydrofurfuryl-3-(2,6-dichloro-4-

trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 915396-43-9), N-ethyl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452099-05-7), N-tetrahydrofurfuryl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452100-03-7), 3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-[1,3,5]triazinan-2,4-dione, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), 2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isoindole-1,3-dione, 1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-1*H*-pyrimidine-2,4-dione, methyl (*E*)-4-[2-chloro-5-[4-chloro-5-(difluoromethoxy)-1*H*-methyl-pyrazol-3-yl]-4-fluorophenoxy]-3-methoxy-but-2-enoate [CAS 948893-00-3], 3-[7-Chloro-5-fluoro-2-(trifluoromethyl)-1*H*-benzimidazol-4-yl]-1-methyl-6-(trifluoromethyl)-1*H*-pyrimidine-2,4-dione (CAS 212754-02-4).

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b5), in particular selected from the group consisting of clomazone, diflufenican, flurochloridone, isoxaflutole, mesotrione, picolinafen, sulcotrione, tefuryltrione, tembotrione, topramezone, bicycloprrone, amitrole and flumeturon.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b6), in particular selected from the group consisting of glyphosate, glyphosate-isopropylammonium and glyphosate-trimesium (sulfosate).

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-

trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b7), in particular selected from the group consisting of glufosinate, glufosinate-P and glufosinate-ammonium.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4) especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b9), in particular selected from the group consisting of pendimethalin and trifluralin.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4)), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b10), in particular selected from the group consisting of acetochlor, cafenstrole, dimethenamid-P, fentrazamide, flufenacet, mefenacet, metazachlor, metolachlor, S-metolachlor, fenoxasulfone and pyroxasulfone. Likewise, preference is given to compositions comprising in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b10), in particular selected from the group consisting of isoxazoline compounds of the formulae II.1, II.2, II.3, II.4, II.5, II.6, II.7, II.8 and II.9, as defined above.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b13), in particular selected from the group consisting of 2,4-D and its salts and esters, aminocyclopyrachlor and its salts and esters, aminopyralid and its salts such as aminopyralid-tris(2-hydroxypropyl)ammonium and its esters, clopyralid and its salts and esters, dicamba and its salts and esters, fluroxypyr-meptyl, quinclorac and quinmerac.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b14), in particular selected from the group consisting of diflufenzopyr and diflufenzopyr-sodium.

According to another preferred embodiment of the invention, the composition comprises, in addition to a PPO-inhibiting herbicide, preferably acifluorfen, acifluorfen-sodium, butafenacil, cinidon-ethyl, carfentrazone-ethyl, flumioxazin, fluthiacet-methyl, fomesafen, lactofen, oxadiargyl, oxyfluorfen, saflufenacil, sulfentrazone, ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100), 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), especially preferred saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), at least one and especially exactly one herbicidally active compound from group b15), in particular selected from the group consisting of dymron (= daimuron), indanofan, indaziflam, oxaziclomefone and triaziflam.

Here and below, the term "binary compositions" includes compositions comprising one or more, for example 1, 2 or 3, active compounds of the PPO-inhibiting herbicide and either one or more, for example 1, 2 or 3, herbicides B.

In binary compositions comprising at least one PPO-inhibiting herbicide as component A and at least one herbicide B, the weight ratio of the active compounds A:B is generally in the range of from 1:1000 to 1000:1, preferably in the range of from 1:500 to 500:1, in particular in the range of from 1:250 to 250:1 and particularly preferably in the range of from 1:75 to 75:1.

5

Particularly preferred herbicides B are the herbicides B as defined above; in particular the herbicides B.1 - B.229 listed below in table B:

Table B:

	Herbicide B
B.1	clethodim
B.2	clodinafop-propargyl
B.3	cycloxydim
B.4	cyhalofop-butyl
B.5	fenoxaprop-ethyl
B.6	fenoxaprop-P-ethyl
B.7	fluazifop
B.8	metamifop
B.9	pinoxaden
B.10	profoxydim
B.11	quizalofop
B.12	sethoxydim
B.13	tepraloxydim
B.14	tralkoxydim
B.15	esprocarb
B.16	ethofumesate
B.17	molinate
B.18	prosulfocarb
B.19	thiobencarb
B.20	triallate
B.21	bensulfuron-methyl
B.22	bispyribac-sodium
B.23	cloransulam-methyl
B.24	chlorsulfuron
B.25	clorimuron
B.26	cyclosulfamuron
B.27	diclosulam
B.28	florasulam
B.29	flumetsulam
B.30	flupyrsulfuron-methyl-sodium
B.31	foramsulfuron
B.32	halosulfuron-methyl
B.33	imazamox

	Herbicide B
B.34	imazamox-ammonium
B.35	imazapic
B.36	imazapic-ammonium
B.37	imazapic-isopropylammonium
B.38	imazapyr
B.39	imazapyr-ammonium
B.40	imazapyr-isopropylammonium
B.41	imazaquin
B.42	imazaquin-ammonium
B.43	imazethapyr
B.44	imazethapyr-ammonium
B.45	imazethapyr-isopropylammonium
B.46	imazosulfuron
B.47	iodosulfuron-methyl-sodium
B.48	iofensulfuron
B.49	iofensulfuron-sodium
B.50	mesosulfuron-methyl
B.51	metazosulfuron
B.52	metsulfuron-methyl
B.53	metosulam
B.54	nicosulfuron
B.55	penoxsulam
B.56	propoxycarbazon-sodium
B.57	pyrazosulfuron-ethyl
B.58	pyribenzoxim
B.59	pyriftalid
B.60	pyrithiobac-sodium
B.61	pyroxsulam
B.62	propyrisulfuron
B.63	rimsulfuron
B.64	sulfosulfuron
B.65	thiencarbazone-methyl

	Herbicide B
B.66	thifensulfuron-methyl
B.67	tribenuron-methyl
B.68	trifloxysulfuron
B.69	tritosulfuron
B.70	triafamone
B.71	ametryne
B.72	atrazine
B.73	bentazon
B.74	bromoxynil
B.75	bromoxynil-octanoate
B.76	bromoxynil-heptanoate
B.77	bromoxynil-potassium
B.78	diuron
B.79	fluometuron
B.80	hexazinone
B.81	isoproturon
B.82	linuron
B.83	metamitron
B.84	metribuzin
B.85	prometryne
B.86	propanil
B.87	simazin
B.88	terbuthylazine
B.89	terbutryn
B.90	paraquat-dichloride
B.91	acifluorfen
B.92	acifluorfen-sodium
B.93	azafenidin
B.94	bencarbazone
B.95	benzfendizone
B.96	bifenox
B.97	butafenacil
B.98	carfentrazone
B.99	carfentrazone-ethyl
B.100	chlomethoxyfen
B.101	cinidon-ethyl
B.102	fluazolate
B.103	flufenpyr
B.104	flufenpyr-ethyl
B.105	flumiclorac
B.106	flumiclorac-pentyl

	Herbicide B
B.107	flumioxazin
B.108	fluoroglycofen
B.109	fluoroglycofen-ethyl
B.110	fluthiacet
B.111	fluthiacet-methyl
B.112	fomesafen
B.113	halosafen
B.114	lactofen
B.115	oxadiargyl
B.116	oxadiazon
B.117	oxyfluorfen
B.118	pentoxazone
B.119	profluazol
B.120	pyraclonil
B.121	pyraflufen
B.122	pyraflufen-ethyl
B.123	saflufenacil
B.124	sulfentrazone
B.125	thidiazimin
B.126	tiafenacil
B.127	ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6)
B.128	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4)
B.129	N-ethyl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 452098-92-9)
B.130	N-tetrahydrofurfuryl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1H-pyrazole-1-carboxamide (CAS 915396-43-9)

	Herbicide B
B.131	N-ethyl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1 <i>H</i> -pyrazole-1-carboxamide (CAS 452099-05-7)
B.132	N-tetrahydrofurfuryl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1 <i>H</i> -pyrazole-1-carboxamide (CAS 452100-03-7)
B.133	3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2 <i>H</i> -benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-[1,3,5]triazinan-2,4-dione
B.134	2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2 <i>H</i> -benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isoin-dole-1,3-dione
B.135	1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2 <i>H</i> -benzo[1,4]oxazin-6-yl)-1 <i>H</i> -pyrimidine-2,4-dione
B.136	methyl (<i>E</i>)-4-[2-chloro-5-[4-chloro-5-(difluoromethoxy)-1 <i>H</i> -methyl-pyrazol-3-yl]-4-fluorophenoxy]-3-methoxy-but-2-enoate [CAS 948893-00-3]
B.137	3-[7-Chloro-5-fluoro-2-(trifluoromethyl)-1 <i>H</i> -benzimidazol-4-yl]-1-methyl-6-(trifluoromethyl)-1 <i>H</i> -pyrimidine-2,4-dione (CAS 212754-02-4)
B.138	benzobicyclon
B.139	clomazone
B.140	diflufenican
B.141	flurochloridone
B.142	isoxaflutole
B.143	mesotrione
B.144	norflurazone
B.145	picolinafen

	Herbicide B
B.146	sulcotrione
B.147	tefuryltrione
B.148	tembotrione
B.149	topramezone
B.150	topramezone-sodium
B.151	bicyclopyrone
B.152	amitrole
B.153	fluometuron
B.154	glyphosate
B.155	glyphosate-ammonium
B.156	glyphosate-dimethylammonium
B.157	glyphosate-isopropylammonium
B.158	glyphosate-trimesium (sulfosate)
B.159	glyphosate-potassium
B.160	glufosinate
B.161	glufosinate-ammonium
B.162	glufosinate-P
B.163	glufosinate-P-ammonium
B.164	pendimethalin
B.165	trifluralin
B.166	acetochlor
B.167	butachlor
B.168	cafenstrole
B.169	dimethenamid-P
B.170	fentrazamide
B.171	flufenacet
B.172	mefenacet
B.173	metazachlor
B.174	metolachlor
B.175	S-metolachlor
B.176	pretilachlor
B.177	fenoxasulfone
B.178	isoxaben
B.179	ipfencarbazone
B.180	pyroxasulfone
B.181	2,4-D
B.182	2,4-D-isobutyl
B.183	2,4-D-dimethylammonium
B.184	2,4-D-N,N,N-trimethylethanolammonium

	Herbicide B
B.185	aminopyralid
B.186	aminopyralid-methyl
B.187	aminopyralid-tris(2-hydroxypropyl)ammonium
B.188	clopyralid
B.189	clopyralid-methyl
B.190	clopyralid-olamine
B.191	dicamba
B.192	dicamba-butotyl
B.193	dicamba-diglycolamine
B.194	dicamba-dimethylammonium
B.195	dicamba-diolamine
B.196	dicamba-isopropylammonium
B.197	dicamba-potassium
B.198	dicamba-sodium
B.199	dicamba-trolamine
B.200	dicamba-N,N-bis-(3-aminopropyl)methylamine
B.201	dicamba-diethylenetriamine
B.202	fluroxypyr
B.203	fluroxypyr-meptyl
B.204	MCPA
B.205	MCPA-2-ethylhexyl
B.206	MCPA-dimethylammonium

	Herbicide B
B.207	quinclorac
B.208	quinclorac-dimethylammonium
B.209	quinmerac
B.210	quinmerac-dimethylammonium
B.211	aminocyclopyrachlor
B.212	aminocyclopyrachlor-potassium
B.213	aminocyclopyrachlor-methyl
B.214	diflufenzopyr
B.215	diflufenzopyr-sodium
B.216	dymron
B.217	indanofan
B.218	indaziflam
B.219	oxaziclomefone
B.220	triaziflam
B.221	II.1
B.222	II.2
B.223	II.3
B.224	II.4
B.225	II.5
B.226	II.6
B.227	II.7
B.228	II.8
B.229	II.9

Particularly preferred are compositions 1.1 to 1.229, comprising acifluorfen and the substance(s) as defined in the respective row of table B-1:

5 **Table B-1**(compositions 1.1 to 1.229):

comp. no.	herbicide B
1.1	B.1
1.2	B.2
1.3	B.3
1.4	B.4
1.5	B.5
1.6	B.6
1.7	B.7
1.8	B.8
1.9	B.9
1.10	B.10

1.11	B.11
1.12	B.12
1.13	B.13
1.14	B.14
1.15	B.15
1.16	B.16
1.17	B.17
1.18	B.18
1.19	B.19
1.20	B.20

1.21	B.21
1.22	B.22
1.23	B.23
1.24	B.24
1.25	B.25
1.26	B.26
1.27	B.27
1.28	B.28
1.29	B.29
1.30	B.30

1.31	B.31
1.32	B.32
1.33	B.33
1.34	B.34
1.35	B.35
1.36	B.36
1.37	B.37
1.38	B.38
1.39	B.39
1.40	B.40
1.41	B.41
1.42	B.42
1.43	B.43
1.44	B.44
1.45	B.45
1.46	B.46
1.47	B.47
1.48	B.48
1.49	B.49
1.50	B.50
1.51	B.51
1.52	B.52
1.53	B.53
1.54	B.54
1.55	B.55
1.56	B.56
1.57	B.57
1.58	B.58.
1.59	B.59
1.60	B.60
1.61	B.61
1.62	B.62
1.63	B.63
1.64	B.64
1.65	B.65
1.66	B.66
1.67	B.67
1.68	B.68
1.69	B.69
1.70	B.70
1.71	B.71
1.72	B.72
1.73	B.73

1.74	B.74
1.75	B.75
1.76	B.76
1.77	B.77
1.78	B.78
1.79	B.79
1.80	B.80
1.81	B.81
1.82	B.82
1.83	B.83
1.84	B.84
1.85	B.85
1.86	B.86
1.87	B.87
1.88	B.88
1.89	B.89
1.90	B.90
1.91	B.91
1.92	B.92
1.93	B.93
1.94	B.94
1.95	B.95
1.96	B.96
1.97	B.97
1.98	B.98
1.99	B.99
1.100	B.100
1.101	B.101
1.102	B.102
1.103	B.103
1.104	B.104
1.105	B.105
1.106	B.106
1.107	B.107
1.108	B.108
1.109	B.109
1.110	B.110
1.111	B.111
1.112	B.112
1.113	B.113
1.114	B.114
1.115	B.115
1.116	B.116

1.117	B.117
1.118	B.118
1.119	B.119
1.120	B.120
1.121	B.121
1.122	B.122
1.123	B.123
1.124	B.124
1.125	B.125
1.126	B.126
1.127	B.127
1.128	B.128
1.129	B.129
1.130	B.130
1.131	B.131
1.132	B.132
1.133	B.133
1.134	B.134
1.135	B.135
1.136	B.136
1.137	B.137
1.138	B.138
1.139	B.139
1.140	B.140
1.141	B.141
1.142	B.142
1.143	B.143
1.144	B.144
1.145	B.145
1.146	B.146
1.147	B.147
1.148	B.148
1.149	B.149
1.150	B.150
1.151	B.151
1.152	B.152
1.153	B.153
1.154	B.154
1.155	B.155
1.156	B.156
1.157	B.157
1.158	B.158
1.159	B.159

1.160	B.160
1.161	B.161
1.162	B.162
1.163	B.163
1.164	B.164
1.165	B.165
1.166	B.166
1.167	B.167
1.168	B.168
1.169	B.169
1.170	B.170
1.171	B.171
1.172	B.172
1.173	B.173
1.174	B.174
1.175	B.175
1.176	B.176
1.177	B.177
1.178	B.178
1.179	B.179
1.180	B.180
1.181	B.181
1.182	B.182
1.183	B.183
1.184	B.184
1.185	B.185
1.186	B.186
1.187	B.187
1.188	B.188
1.189	B.189
1.190	B.190
1.191	B.191
1.192	B.192
1.193	B.193
1.194	B.194
1.195	B.195
1.196	B.196
1.197	B.197
1.198	B.198
1.199	B.199
1.200	B.200
1.201	B.201
1.202	B.202

1.203	B.203
1.204	B.204
1.205	B.205
1.206	B.206
1.207	B.207
1.208	B.208
1.209	B.209
1.210	B.210
1.211	B.211
1.212	B.212
1.213	B.213
1.214	B.214
1.215	B.215
1.216	B.216
1.217	B.217
1.218	B.218
1.219	B.219
1.220	B.220
1.221	B.221
1.222	B.222
1.223	B.223
1.224	B.224
1.225	B.225
1.226	B.226
1.227	B.227
1.228	B.228
1.229	B.229

Also especially preferred are compositions 2.1. to 2.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A acifluorfen-sodium.

5 Also especially preferred are compositions 3.1. to 3.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A azafenidin.

Also especially preferred are compositions 4.1. to 4.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A bencarbazone.

10 Also especially preferred are compositions 5.1. to 5.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A benzfendizone.

Also especially preferred are compositions 6.1. to 6.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A bifenox.
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Also especially preferred are compositions 7.1. to 7.229 which differ from the corresponding compositions 1.1 to 1.227 only in that they comprise as component A butafenacil.

Also especially preferred are compositions 8.1. to 8.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A carfentrazone.
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Also especially preferred are compositions 9.1. to 9.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A carfentrazone-ethyl.

25 Also especially preferred are compositions 10.1. to 10.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A chlormethoxyfen.

Also especially preferred are compositions 11.1. to 11.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A cinidon-ethyl.
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Also especially preferred are compositions 12.1. to 12.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A fluazolate.

Also especially preferred are compositions 13.1. to 13.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A flufenpyr.
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Also especially preferred are compositions 14.1. to 14.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A flufenpyr-ethyl.

40 Also especially preferred are compositions 15.1. to 15.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A flumiclorac.

Also especially preferred are compositions 16.1. to 16.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A flumiclorac-pentyl.

Also especially preferred are compositions 17.1. to 17.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A flumioxazin.

5 Also especially preferred are compositions 18.1. to 18.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A fluoroglycofen.

Also especially preferred are compositions 19.1. to 19.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A fluoroglycofen-ethyl.

10 Also especially preferred are compositions 20.1. to 20.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A fluthiacet.

Also especially preferred are compositions 21.1. to 21.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A fluthiacet-methyl.

15 Also especially preferred are compositions 22.1. to 22.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A fomesafen.

20 Also especially preferred are compositions 23.1. to 23.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A halosafen.

Also especially preferred are compositions 24.1. to 24.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A lactofen.

25 Also especially preferred are compositions 25.1. to 25.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A oxadiargyl.

Also especially preferred are compositions 26.1. to 26.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A oxadiazon.

30 Also especially preferred are compositions 27.1. to 27.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A oxyfluorfen.

35 Also especially preferred are compositions 28.1. to 28.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A pentoxazone.

Also especially preferred are compositions 29.1. to 29.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A profluazol.

40 Also especially preferred are compositions 30.1. to 30.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A pyraclonil.

Also especially preferred are compositions 31.1. to 31.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A pyraflufen.

Also especially preferred are compositions 32.1. to 32.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A pyraflufen-ethyl.

5 Also especially preferred are compositions 33.1. to 33.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A saflufenacil.

Also especially preferred are compositions 34.1. to 34.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A sulfentrazone.

10 Also especially preferred are compositions 35.1. to 35.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A thidiazimin.

Also especially preferred are compositions 36.1. to 36.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A tiafenacil.

15 Also especially preferred are compositions 37.1. to 37.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A ethyl [3-[2-chloro-4-fluoro-5-(1-methyl-6-trifluoromethyl-2,4-dioxo-1,2,3,4-tetrahydropyrimidin-3-yl)phenoxy]-2-pyridyloxy]acetate (CAS 353292-31-6; S-3100).

20 Also especially preferred are compositions 38.1. to 38.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4)

25 Also especially preferred are compositions 39.1. to 39.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A N-ethyl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452098-92-9).

30 Also especially preferred are compositions 40.1. to 40.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A N-tetrahydrofurfuryl-3-(2,6-dichloro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 915396-43-9).

35 Also especially preferred are compositions 41.1. to 41.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A N-ethyl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452099-05-7).

40 Also especially preferred are compositions 42.1. to 42.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A N-tetrahydrofurfuryl-3-(2-chloro-6-fluoro-4-trifluoromethylphenoxy)-5-methyl-1*H*-pyrazole-1-carboxamide (CAS 452100-03-7).

Also especially preferred are compositions 43.1. to 43.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A
3-[7-fluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl]-1,5-dimethyl-6-thioxo-
[1,3,5]triazinan-2,4-dione.

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Also especially preferred are compositions 44.1. to 44.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A methyl (*E*)-4-[2-chloro-5-[4-chloro-5-(difluoromethoxy)-1*H*-methyl-pyrazol-3-yl]-4-fluoro-phenoxy]-3-methoxy-but-2-enoate (CAS 948893-00-3).

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Also especially preferred are compositions 45.1. to 45.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A
3-[7-Chloro-5-fluoro-2-(trifluoromethyl)-1*H*-benzimidazol-4-yl]-1-methyl-6-(trifluoromethyl)-1*H*-pyrimidine-2,4-dione (CAS 212754-02-4).

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Also especially preferred are compositions 46.1. to 46.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A
2-(2,2,7-Trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-4,5,6,7-tetrahydro-isoindole-1,3-dione.

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Also especially preferred are compositions 47.1. to 47.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they comprise as component A
1-Methyl-6-trifluoromethyl-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-3,4-dihydro-2H-benzo[1,4]oxazin-6-yl)-1*H*-pyrimidine-2,4-dione

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Also especially preferred are compositions 48.1. to 48.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise benoxacor as safener C.

Also especially preferred are compositions 49.1. to 49.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise cloquintocet as safener C.

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Also especially preferred are compositions 50.1. to 50.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise cyprosulfamide as safener C.

Also especially preferred are compositions 51.1. to 51.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise dichlormid as safener C.

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Also especially preferred are compositions 52.1. to 52.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise fenclorazole as safener C.

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Also especially preferred are compositions 53.1. to 53.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise fenclorim as safener C.

Also especially preferred are compositions 54.1. to 54.229 which differ from the corresponding

compositions 1.1 to 1.229 only in that they additionally comprise furilazole as safener C.

Also especially preferred are compositions 55.1. to 55.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise isoxadifen as safener C.

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Also especially preferred are compositions 56.1. to 56.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise mefenpyr as safener C.

Also especially preferred are compositions 57.1. to 57.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise 4-(dichloroacetyl)-1-oxa-4-azaspiro[4.5]decane (MON4660, CAS 71526-07-3) as safener C.

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Also especially preferred are compositions 58.1. to 58.229 which differ from the corresponding compositions 1.1 to 1.229 only in that they additionally comprise 2,2,5-trimethyl-3-(dichloroacetyl)-1,3-oxazolidine (R-29148, CAS 52836-31-4) as safener C.

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It is generally preferred to use the compounds of the invention in combination with herbicides that are selective for the crop being treated and which complement the spectrum of weeds controlled by these compounds at the application rate employed. It is further generally preferred to apply the compounds of the invention and other complementary herbicides at the same time, either as a combination formulation or as a tank mix.

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It is recognized that the polynucleotide molecules and polypeptides of the invention encompass polynucleotide molecules and polypeptides comprising a nucleotide or an amino acid sequence that is sufficiently identical to nucleotide sequences set forth in SEQ ID Nos: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47, or to the amino acid sequences set forth in SEQ ID Nos: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48. The term "sufficiently identical" is used herein to refer to a first amino acid or nucleotide sequence that contains a sufficient or minimum number of identical or equivalent (e.g., with a similar side chain) amino acid residues or nucleotides to a second amino acid or nucleotide sequence such that the first and second amino acid or nucleotide sequences have a common structural domain and/or common functional activity.

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Generally, "sequence identity" refers to the extent to which two optimally aligned DNA or amino acid sequences are invariant throughout a window of alignment of components, e.g., nucleotides or amino acids. An "identity fraction" for aligned segments of a test sequence and a reference sequence is the number of identical components that are shared by the two aligned sequences divided by the total number of components in reference sequence segment, i.e., the entire reference sequence or a smaller defined part of the reference sequence. "Percent identity" is the identity fraction times 100. Optimal alignment of sequences for aligning a comparison window are well known to those skilled in the art and may be conducted by tools such as the local homology algorithm of Smith and Waterman, the homology alignment algorithm of Needleman and Wunsch, the search for similarity method of Pearson and Lipman, and preferably by computerized implementations of these algorithms such as GAP, BESTFIT, FASTA, and TFASTA available as

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part of the GCG. Wisconsin Package. (Accelrys Inc. Burlington, Mass.)

Polynucleotides and Oligonucleotides

5 By an "isolated polynucleotide", including DNA, RNA, or a combination of these, single or double stranded, in the sense or antisense orientation or a combination of both, dsRNA or otherwise, we mean a polynucleotide which is at least partially separated from the polynucleotide sequences with which it is associated or linked in its native state. That means other nucleic acid molecules are present in an amount less than 5% based on weight of the amount of the desired nucleic acid,
10 preferably less than 2% by weight, more preferably less than 1% by weight, most preferably less than 0.5% by weight. Preferably, an "isolated" nucleic acid is free of some of the sequences that naturally flank the nucleic acid (i.e., sequences located at the 5' and 3' ends of the nucleic acid) in the genomic DNA of the organism from which the nucleic acid is derived. For example, in various embodiments, the isolated herbicide resistance and/or tolerance related protein encoding nucleic
15 acid molecule can contain less than about 5 kb, 4 kb, 3 kb, 2 kb, 1 kb, 0.5 kb or 0.1 kb of nucleotide sequences which naturally flank the nucleic acid molecule in genomic DNA of the cell from which the nucleic acid is derived. Moreover, an "isolated" nucleic acid molecule, such as a cDNA molecule, can be free from some of the other cellular material with which it is naturally associated, or culture medium when produced by recombinant techniques, or chemical precursors
20 or other chemicals when chemically synthesized. Preferably, the isolated polynucleotide is at least 60% free, preferably at least 75% free, and most preferably at least 90% free from other components with which they are naturally associated. As the skilled addressee would be aware, an isolated polynucleotide can be an exogenous polynucleotide present in, for example, a transgenic organism which does not naturally comprise the polynucleotide.

25 Furthermore, the terms "polynucleotide(s)", "nucleic acid sequence(s)", "nucleotide sequence(s)", "nucleic acid(s)", "nucleic acid molecule" are used interchangeably herein and refer to nucleotides, either ribonucleotides or deoxyribonucleotides or a combination of both, in a polymeric unbranched form of any length.

30 The term "mutated PPO nucleic acid" refers to a PPO nucleic acid having a sequence that is mutated from a wild-type PPO nucleic acid and that confers increased PPO-inhibiting herbicide tolerance to a plant in which it is expressed. Furthermore, the term "mutated protoporphyrinogen oxidase (mutated PPO)" refers to the replacement of an amino acid of the wild-type primary
35 sequences SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, or a variant, a derivative, a homologue, an orthologue, or paralogue thereof, with another amino acid. The expression "mutated amino acid" will be used below to designate the amino acid which is replaced by another amino acid, thereby designating the site of the mutation in the primary sequence of the protein.

40 In a preferred embodiment, the PPO nucleotide sequence encoding a mutated PPO comprises the sequence of SEQ ID NO: 1, 3, 23, 29, 37, 45, or 47, or a variant or derivative thereof.

Furthermore, it will be understood by the person skilled in the art that the PPO nucleotide

sequences encompass homologues, paralogues and orthologues of SEQ ID NO: 1, 3, 23, 29, 37, 45, or 47, as defined hereinafter.

The term "variant" with respect to a sequence (e.g., a polypeptide or nucleic acid sequence such as – for example – a transcription regulating nucleotide sequence of the invention) is intended to mean substantially similar sequences. For nucleotide sequences comprising an open reading frame, variants include those sequences that, because of the degeneracy of the genetic code, encode the identical amino acid sequence of the native protein. Naturally occurring allelic variants such as these can be identified with the use of well-known molecular biology techniques, as, for example, with polymerase chain reaction (PCR) and hybridization techniques. Variant nucleotide sequences also include synthetically derived nucleotide sequences, such as those generated, for example, by using site-directed mutagenesis and for open reading frames, encode the native protein, as well as those that encode a polypeptide having amino acid substitutions relative to the native protein, e.g. the mutated PPO according to the present invention as disclosed herein.

Generally, nucleotide sequence variants of the invention will have at least 30, 40, 50, 60, to 70%, e.g., preferably 71%, 72%, 73%, 74%, 75%, 76%, 77%, 78%, to 79%, generally at least 80%, e.g., 81%-84%, at least 85%, e.g., 86%, 87%, 88%, 89%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, to 98% and 99% nucleotide "sequence identity" to the nucleotide sequence of SEQ ID NO: SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47. The % identity of a polynucleotide is determined by GAP (Needleman and Wunsch, 1970) analysis (GCG program) with a gap creation penalty=5, and a gap extension penalty=0.3. Unless stated otherwise, the query sequence is at least 45 nucleotides in length, and the GAP analysis aligns the two sequences over a region of at least 45 nucleotides. Preferably, the query sequence is at least 150 nucleotides in length, and the GAP analysis aligns the two sequences over a region of at least 150 nucleotides. More preferably, the query sequence is at least 300 nucleotides in length and the GAP analysis aligns the two sequences over a region of at least 300 nucleotides. Even more preferably, the GAP analysis aligns the two sequences over their entire length.

Polypeptides

By "substantially purified polypeptide" or "purified" a polypeptide is meant that has been separated from one or more lipids, nucleic acids, other polypeptides, or other contaminating molecules with which it is associated in its native state. It is preferred that the substantially purified polypeptide is at least 60% free, more preferably at least 75% free, and more preferably at least 90% free from other components with which it is naturally associated. As the skilled addressee will appreciate, the purified polypeptide can be a recombinantly produced polypeptide. The terms "polypeptide" and "protein" are generally used interchangeably and refer to a single polypeptide chain which may or may not be modified by addition of non-amino acid groups. It would be understood that such polypeptide chains may associate with other polypeptides or proteins or other molecules such as co-factors. The terms "proteins" and "polypeptides" as used herein also include variants, mutants, modifications, analogous and/or derivatives of the polypeptides of the invention as described herein.

The % identity of a polypeptide is determined by GAP (Needleman and Wunsch, 1970) analysis

(GCG program) with a gap creation penalty=5, and a gap extension penalty=0.3. The query sequence is at least 25 amino acids in length, and the GAP analysis aligns the two sequences over a region of at least 25 amino acids. More preferably, the query sequence is at least 50 amino acids in length, and the GAP analysis aligns the two sequences over a region of at least 50 amino acids.

- 5 More preferably, the query sequence is at least 100 amino acids in length and the GAP analysis aligns the two sequences over a region of at least 100 amino acids. Even more preferably, the query sequence is at least 250 amino acids in length and the GAP analysis aligns the two sequences over a region of at least 250 amino acids. Even more preferably, the GAP analysis aligns the two sequences over their entire length.

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With regard to a defined polypeptide, it will be appreciated that % identity figures higher than those provided above will encompass preferred embodiments. Thus, where applicable, in light of the minimum % identity figures, it is preferred that the PPO polypeptide of the invention comprises an amino acid sequence which is at least 40%, more preferably at least 45%, more preferably at least 50%, more preferably at least 55%, more preferably at least 60%, more preferably at least 65%, more preferably at least 70%, more preferably at least 75%, more preferably at least 80%, more preferably at least 85%, more preferably at least 90%, more preferably at least 91%, more preferably at least 92%, more preferably at least 93%, more preferably at least 94%, more preferably at least 95%, more preferably at least 96%, more preferably at least 97%, more preferably at least 98%, more preferably at least 99%, more preferably at least 99.1%, more preferably at least 99.2%, more preferably at least 99.3%, more preferably at least 99.4%, more preferably at least 99.5%, more preferably at least 99.6%, more preferably at least 99.7%, more preferably at least 99.8%, and even more preferably at least 99.9% identical to SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48.

25

By "variant" polypeptide is intended a polypeptide derived from the protein of SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48 by deletion (so-called truncation) or addition of one or more amino acids to the N-terminal and/or C-terminal end of the native protein; deletion or addition of one or more amino acids at one or more sites in the native protein; or substitution of one or more amino acids at one or more sites in the native protein. Such variants may result from, for example, genetic polymorphism or from human manipulation. Methods for such manipulations are generally known in the art.

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- 35 "Derivatives" of a protein encompass peptides, oligopeptides, polypeptides, proteins and enzymes having amino acid substitutions, deletions and/or insertions relative to the unmodified protein in question and having similar biological and functional activity as the unmodified protein from which they are derived.

- 40 "Homologues" of a protein encompass peptides, oligopeptides, polypeptides, proteins and enzymes having amino acid substitutions, deletions and/or insertions relative to the unmodified protein in question and having similar biological and functional activity as the unmodified protein from which they are derived.

A deletion refers to removal of one or more amino acids from a protein.

An insertion refers to one or more amino acid residues being introduced into a predetermined site in a protein. Insertions may comprise N-terminal and/or C-terminal fusions as well as intra-sequence insertions of single or multiple amino acids. Generally, insertions within the amino acid sequence will be smaller than N- or C-terminal fusions, of the order of about 1 to 10 residues.

Examples of N- or C-terminal fusion proteins or peptides include the binding domain or activation domain of a transcriptional activator as used in the yeast two-hybrid system, phage coat proteins, (histidine)-6-tag, glutathione S-transferase-tag, protein A, maltose-binding protein, dihydrofolate reductase, Tag•100 epitope, c-myc epitope, FLAG®-epitope, lacZ, CMP (calmodulin-binding peptide), HA epitope, protein C epitope and VSV epitope.

A substitution refers to replacement of amino acids of the protein with other amino acids having similar properties (such as similar hydrophobicity, hydrophilicity, antigenicity, propensity to form or break α -helical structures or β -sheet structures). Amino acid substitutions are typically of single residues, but may be clustered depending upon functional constraints placed upon the polypeptide and may range from 1 to 10 amino acids; insertions will usually be of the order of about 1 to 10 amino acid residues. The amino acid substitutions are preferably conservative amino acid substitutions. Conservative substitution tables are well known in the art (see for example Creighton (1984) Proteins. W.H. Freeman and Company (Eds).

Table 2: Examples of conserved amino acid substitutions

Residue	Conservative Substitutions	Residue	Conservative Substitutions
Ala	Ser	Leu	Ile; Val
Arg	Lys	Lys	Arg; Gln
Asn	Gln; His	Met	Leu; Ile
Asp	Glu	Phe	Met; Leu; Tyr
Gln	Asn	Ser	Thr; Gly
Cys	Ser	Thr	Ser; Val
Glu	Asp	Trp	Tyr
Gly	Pro	Tyr	Trp; Phe
His	Asn; Gln	Val	Ile; Leu
Ile	Leu, Val		

Amino acid substitutions, deletions and/or insertions may readily be made using peptide synthetic techniques well known in the art, such as solid phase peptide synthesis and the like, or by recombinant DNA manipulation. Methods for the manipulation of DNA sequences to produce substitution, insertion or deletion variants of a protein are well known in the art. For example, techniques for making substitution mutations at predetermined sites in DNA are well known to those skilled in the art and include M13 mutagenesis, T7-Gen in vitro mutagenesis (USB, Cleveland, OH), QuickChange Site Directed mutagenesis (Stratagene, San Diego, CA), PCR-mediated site-directed mutagenesis or other site-directed mutagenesis protocols.

“Derivatives” further include peptides, oligopeptides, polypeptides which may, compared to the amino acid sequence of the naturally-occurring form of the protein, such as the protein of interest, comprise substitutions of amino acids with non-naturally occurring amino acid residues, or

additions of non-naturally occurring amino acid residues. "Derivatives" of a protein also encompass peptides, oligopeptides, polypeptides which comprise naturally occurring altered (glycosylated, acylated, prenylated, phosphorylated, myristoylated, sulphated etc.) or non-naturally altered amino acid residues compared to the amino acid sequence of a naturally-occurring form of the

polypeptide. A derivative may also comprise one or more non-amino acid substituents or additions compared to the amino acid sequence from which it is derived, for example a reporter molecule or other ligand, covalently or non-covalently bound to the amino acid sequence, such as a reporter molecule which is bound to facilitate its detection, and non-naturally occurring amino acid residues relative to the amino acid sequence of a naturally-occurring protein. Furthermore, "derivatives" also include fusions of the naturally-occurring form of the protein with tagging peptides such as FLAG, HIS6 or thioredoxin (for a review of tagging peptides, see Terpe, *Appl. Microbiol. Biotechnol.* 60, 523-533, 2003).

"Orthologues" and "paralogues" encompass evolutionary concepts used to describe the ancestral relationships of genes. Paralogues are genes within the same species that have originated through duplication of an ancestral gene; orthologues are genes from different organisms that have originated through speciation, and are also derived from a common ancestral gene. A non-limiting list of examples of such orthologues are shown in Table 1.

It is well-known in the art that paralogues and orthologues may share distinct domains harboring suitable amino acid residues at given sites, such as binding pockets for particular substrates, compounds such as e.g. herbicides, or binding motifs for interaction with other proteins.

The term "domain" refers to a set of amino acids conserved at specific positions along an alignment of sequences of evolutionarily related proteins. While amino acids at other positions can vary between homologues, amino acids that are highly conserved at specific positions indicate amino acids that are likely essential in the structure, stability or function of a protein. Identified by their high degree of conservation in aligned sequences of a family of protein homologues, they can be used as identifiers to determine if any polypeptide in question belongs to a previously identified polypeptide family.

The term "motif" or "consensus sequence" refers to a short conserved region in the sequence of evolutionarily related proteins. Motifs are frequently highly conserved parts of domains, but may also include only part of the domain, or be located outside of conserved domain (if all of the amino acids of the motif fall outside of a defined domain).

Specialist databases exist for the identification of domains, for example, SMART (Schultz et al. (1998) *Proc. Natl. Acad. Sci. USA* 95, 5857-5864; Letunic et al. (2002) *Nucleic Acids Res* 30, 242-244), InterPro (Mulder et al., (2003) *Nucl. Acids. Res.* 31, 315-318), Prosite (Bucher and Bairoch (1994), A generalized profile syntax for biomolecular sequences motifs and its function in automatic sequence interpretation. (In) *ISMB-94; Proceedings 2nd International Conference on Intelligent Systems for Molecular Biology*. Altman R., Brutlag D., Karp P., Lathrop R., Searls D., Eds., pp53-61, AAAI Press, Menlo Park; Hulo et al., *Nucl. Acids. Res.* 32:D134-D137, (2004)), or Pfam (Bateman et al., *Nucleic Acids Research* 30(1): 276-280 (2002)). A set of tools for in silico

analysis of protein sequences is available on the ExPASy proteomics server (Swiss Institute of Bioinformatics (Gasteiger et al., ExPASy: the proteomics server for in-depth protein knowledge and analysis, *Nucleic Acids Res.* 31:3784-3788(2003)). Domains or motifs may also be identified using routine techniques, such as by sequence alignment.

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Methods for the alignment of sequences for comparison are well known in the art, such methods include GAP, BESTFIT, BLAST, FASTA and TFASTA. GAP uses the algorithm of Needleman and Wunsch ((1970) *J Mol Biol* 48: 443-453) to find the global (i.e. spanning the complete sequences) alignment of two sequences that maximizes the number of matches and minimizes the number of gaps. The BLAST algorithm (Altschul et al. (1990) *J Mol Biol* 215: 403-10) calculates percent sequence identity and performs a statistical analysis of the similarity between the two sequences. The software for performing BLAST analysis is publicly available through the National Centre for Biotechnology Information (NCBI). Homologues may readily be identified using, for example, the ClustalW multiple sequence alignment algorithm (version 1.83), with the default pairwise alignment parameters, and a scoring method in percentage. Global percentages of similarity and identity may also be determined using one of the methods available in the MatGAT software package (Campanella et al., *BMC Bioinformatics.* 2003 Jul 10;4:29. MatGAT: an application that generates similarity/identity matrices using protein or DNA sequences.). Minor manual editing may be performed to optimise alignment between conserved motifs, as would be apparent to a person skilled in the art. Furthermore, instead of using full-length sequences for the identification of homologues, specific domains may also be used. The sequence identity values may be determined over the entire nucleic acid or amino acid sequence or over selected domains or conserved motif(s), using the programs mentioned above using the default parameters. For local alignments, the Smith-Waterman algorithm is particularly useful (Smith TF, Waterman MS (1981) *J. Mol. Biol* 147(1);195-7).

The inventors of the present invention have found that by substituting one or more of the key amino acid residues, employing e.g. one of the above described methods to mutate the encoding nucleic acids, the herbicide tolerance or resistance could be remarkably increased as compared to the activity of the wild type PPO enzymes with SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48. Preferred substitutions of mutated PPO are those that increase the herbicide tolerance of the plant, but leave the biological activity of the oxidase activity substantially unaffected.

Accordingly, in another object of the present invention the key amino acid residues of a PPO enzyme comprising SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, a variant, derivative, orthologue, paralogue or homologue thereof, is substituted by any other amino acid.

In one embodiment, the key amino acid residues of a PPO enzyme, a variant, derivative, orthologue, paralogue or homologue thereof, is substituted by a conserved amino acid as depicted in Table 2.

It will be understood by the person skilled in the art that amino acids located in a close proximity to

the positions of amino acids mentioned below may also be substituted. Thus, in another embodiment the variant of SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, a variant, derivative, orthologue, paralogue or homologue thereof comprises a mutated PPO, wherein an amino acid ± 3 , ± 2 or ± 1 amino acid positions from a key amino acid is substituted by any other amino acid.

Based on techniques well-known in the art, a highly characteristic sequence pattern can be developed, by means of which further of mutated PPO candidates with the desired activity may be searched.

Searching for further mutated PPO candidates by applying a suitable sequence pattern would also be encompassed by the present invention. It will be understood by a skilled reader that the present sequence pattern is not limited by the exact distances between two adjacent amino acid residues of said pattern. Each of the distances between two neighbours in the above patterns may, for example, vary independently of each other by up to ± 10 , ± 5 , ± 3 , ± 2 or ± 1 amino acid positions without substantially affecting the desired activity.

Furthermore, by applying the method of site directed mutagenesis, in particular saturation mutagenesis (see e.g. Schenk et al., Biospektrum 03/2006, pages 277-279), the inventors of the present invention have identified and generated specific amino acid substitutions and combinations thereof, which - when introduced into a plant by transforming and expressing the respective mutated PPO encoding nucleic acid - confer increased herbicide resistance or tolerance to a PPO inhibiting herbicide to said plant.

Thus, in a particularly preferred embodiment, the variant or derivative of the mutated PPO refers to a polypeptide comprising SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 24, SEQ ID NO: 30, SEQ ID NO: 38, SEQ ID NO: 46, or SEQ ID NO: 48, comprising a single amino acid substitution of the following Table 3a. .

Table 3a: Single amino acid substitutions within SEQ ID NO: 2, SEQ ID NO: 4, SEQ ID NO: 24, SEQ ID NO: 30, SEQ ID NO: 38, SEQ ID NO: 46, SEQ ID NO: 48,

Mutation Number	SEQ ID NO:	Key amino acid position combination	Preferred Substitution
1	2	Arg128	Ala
2	2	Arg128	Leu
3	2	Arg128	Val
4	2	Arg128	Ile
5	2	Arg128	Met
6	2	Arg128	His
7	2	Arg128	Lys
8	2	Arg128	Asp
9	2	Arg128	Glu
10	2	Arg128	Ser

11	2	Arg128	Thr
12	2	Arg128	Asn
13	2	Arg128	Gln
14	2	Arg128	Cys
15	2	Arg128	Gly
16	2	Arg128	Pro
17	2	Arg128	Phe
18	2	Arg128	Tyr
19	2	Arg128	Trp
20	2	Phe420	Ala
21	2	Phe420	Leu
22	2	Phe420	Val
23	2	Phe420	Ile
24	2	Phe420	Met
25	2	Phe420	His
26	2	Phe420	Lys
27	2	Phe420	Asp
28	2	Phe420	Glu
29	2	Phe420	Ser
30	2	Phe420	Thr
31	2	Phe420	Asn
32	2	Phe420	Gln
33	2	Phe420	Cys
34	2	Phe420	Gly
35	2	Phe420	Pro
36	2	Phe420	Phe
37	2	Phe420	Tyr
38	2	Phe420	Trp
39	4	Arg128	Ala
40	4	Arg128	Leu
41	4	Arg128	Val
42	4	Arg128	Ile
43	4	Arg128	Met
44	4	Arg128	His
45	4	Arg128	Lys
46	4	Arg128	Asp
47	4	Arg128	Glu
48	4	Arg128	Ser
49	4	Arg128	Thr
50	4	Arg128	Asn
51	4	Arg128	Gln
52	4	Arg128	Cys
53	4	Arg128	Gly

54	4	Arg128	Pro
55	4	Arg128	Phe
56	4	Arg128	Tyr
57	4	Arg128	Trp
58	4	Phe420	Ala
59	4	Phe420	Leu
60	4	Phe420	Val
61	4	Phe420	Ile
62	4	Phe420	Met
63	4	Phe420	His
64	4	Phe420	Lys
65	4	Phe420	Asp
66	4	Phe420	Glu
67	4	Phe420	Ser
68	4	Phe420	Thr
69	4	Phe420	Asn
70	4	Phe420	Gln
71	4	Phe420	Cys
72	4	Phe420	Gly
73	4	Phe420	Pro
74	4	Phe420	Phe
75	4	Phe420	Tyr
76	4	Phe420	Trp
77	24	Arg130	Ala
78	24	Arg130	Leu
79	24	Arg130	Val
80	24	Arg130	Ile
81	24	Arg130	Met
82	24	Arg130	His
83	24	Arg130	Lys
84	24	Arg130	Asp
85	24	Arg130	Glu
86	24	Arg130	Ser
87	24	Arg130	Thr
88	24	Arg130	Asn
89	24	Arg130	Gln
90	24	Arg130	Cys
91	24	Arg130	Gly
92	24	Arg130	Pro
93	24	Arg130	Phe
94	24	Arg130	Tyr
95	24	Arg130	Trp
96	24	Phe433	Ala

97	24	Phe433	Leu
98	24	Phe433	Val
99	24	Phe433	Ile
100	24	Phe433	Met
101	24	Phe433	His
102	24	Phe433	Lys
103	24	Phe433	Asp
104	24	Phe433	Glu
105	24	Phe433	Ser
106	24	Phe433	Thr
107	24	Phe433	Asn
108	24	Phe433	Gln
109	24	Phe433	Cys
110	24	Phe433	Gly
111	24	Phe433	Pro
112	24	Phe433	Phe
113	24	Phe433	Tyr
114	24	Phe433	Trp
115	30	Arg130	Ala
116	30	Arg130	Leu
117	30	Arg130	Val
118	30	Arg130	Ile
119	30	Arg130	Met
120	30	Arg130	His
121	30	Arg130	Lys
122	30	Arg130	Asp
123	30	Arg130	Glu
124	30	Arg130	Ser
125	30	Arg130	Thr
126	30	Arg130	Asn
127	30	Arg130	Gln
128	30	Arg130	Cys
129	30	Arg130	Gly
130	30	Arg130	Pro
131	30	Arg130	Phe
132	30	Arg130	Tyr
133	30	Arg130	Trp
134	30	Phe433	Ala
135	30	Phe433	Leu
136	30	Phe433	Val
137	30	Phe433	Ile
138	30	Phe433	Met
139	30	Phe433	His

140	30	Phe433	Lys
141	30	Phe433	Asp
142	30	Phe433	Glu
143	30	Phe433	Ser
144	30	Phe433	Thr
145	30	Phe433	Asn
146	30	Phe433	Gln
147	30	Phe433	Cys
148	30	Phe433	Gly
149	30	Phe433	Pro
150	30	Phe433	Phe
151	30	Phe433	Tyr
152	30	Phe433	Trp
153	38	Arg98	Ala
154	38	Arg98	Leu
155	38	Arg98	Val
156	38	Arg98	Ile
157	38	Arg98	Met
158	38	Arg98	His
159	38	Arg98	Lys
160	38	Arg98	Asp
161	38	Arg98	Glu
162	38	Arg98	Ser
163	38	Arg98	Thr
164	38	Arg98	Asn
165	38	Arg98	Gln
166	38	Arg98	Cys
167	38	Arg98	Gly
168	38	Arg98	Pro
169	38	Arg98	Phe
170	38	Arg98	Tyr
171	38	Arg98	Trp
172	38	Phe392	Ala
173	38	Phe392	Leu
174	38	Phe392	Val
175	38	Phe392	Ile
176	38	Phe392	Met
177	38	Phe392	His
178	38	Phe392	Lys
179	38	Phe392	Asp
180	38	Phe392	Glu
181	38	Phe392	Ser
182	38	Phe392	Thr

183	38	Phe392	Asn
184	38	Phe392	Gln
185	38	Phe392	Cys
186	38	Phe392	Gly
187	38	Phe392	Pro
188	38	Phe392	Phe
189	38	Phe392	Tyr
190	38	Phe392	Trp
191	46	Arg139	Ala
192	46	Arg139	Leu
193	46	Arg139	Val
194	46	Arg139	Ile
195	46	Arg139	Met
196	46	Arg139	His
197	46	Arg139	Lys
198	46	Arg139	Asp
199	46	Arg139	Glu
200	46	Arg139	Ser
201	46	Arg139	Thr
202	46	Arg139	Asn
203	46	Arg139	Gln
204	46	Arg139	Cys
205	46	Arg139	Gly
206	46	Arg139	Pro
207	46	Arg139	Phe
208	46	Arg139	Tyr
209	46	Arg139	Trp
210	46	Phe465	Ala
211	46	Phe465	Leu
212	46	Phe465	Val
213	46	Phe465	Ile
214	46	Phe465	Met
215	46	Phe465	His
216	46	Phe465	Lys
217	46	Phe465	Asp
218	46	Phe465	Glu
219	46	Phe465	Ser
220	46	Phe465	Thr
221	46	Phe465	Asn
222	46	Phe465	Gln
223	46	Phe465	Cys
224	46	Phe465	Gly
225	46	Phe465	Pro

226	46	Phe465	Phe
227	46	Phe465	Tyr
228	46	Phe465	Trp
229	48	Arg157	Ala
230	48	Arg157	Leu
231	48	Arg157	Val
232	48	Arg157	Ile
233	48	Arg157	Met
234	48	Arg157	His
235	48	Arg157	Lys
236	48	Arg157	Asp
237	48	Arg157	Glu
238	48	Arg157	Ser
239	48	Arg157	Thr
240	48	Arg157	Asn
241	48	Arg157	Gln
242	48	Arg157	Cys
243	48	Arg157	Gly
244	48	Arg157	Pro
245	48	Arg157	Phe
246	48	Arg157	Tyr
247	48	Arg157	Trp
248	48	Tyr439	Ala
249	48	Tyr439	Leu
250	48	Tyr439	Val
251	48	Tyr439	Ile
252	48	Tyr439	Met
253	48	Tyr439	His
254	48	Tyr439	Lys
255	48	Tyr439	Asp
256	48	Tyr439	Glu
257	48	Tyr439	Ser
258	48	Tyr439	Thr
259	48	Tyr439	Asn
260	48	Tyr439	Gln
261	48	Tyr439	Cys
262	48	Tyr439	Gly
263	48	Tyr439	Pro
264	48	Tyr439	Phe
265	48	Tyr439	Tyr
266	48	Tyr439	Trp

In a further particularly preferred embodiment, the variant or derivative of the mutated PPO refers

to a polypeptide comprising SEQ ID NO: 2, SEQ ID NO:4, SEQ ID NO: 24, SEQ ID NO: 30, SEQ ID NO: 38, SEQ ID NO: 46, SEQ ID NO: 48, comprising a combination of amino acid substitutions selected from the following Table 3b.

5 **Table 3b:** SEQ ID NO: 2, SEQ ID NO:4, SEQ ID NO: 24, SEQ ID NO: 30, SEQ ID NO: 38, SEQ ID NO: 46, SEQ ID NO: 48, (combined amino acid substitutions)

Combination Number	SEQ ID NO:	Key amino acid position combination	Preferred Substitution
267	2 & 4	Arg128	Leu
		Phe420	Ala
268	2 & 4	Arg128	Leu
		Phe420	Leu
269	2 & 4	Arg128	Leu
		Phe420	Val
270	2 & 4	Arg128	Leu
		Phe420	Ile
271	2 & 4	Arg128	Leu
		Phe420	Met
272	2 & 4	Arg128	Ala
		Phe420	Ala
273	2 & 4	Arg128	Ala
		Phe420	Leu
274	2 & 4	Arg128	Ala
		Phe420	Val
275	2 & 4	Arg128	Ala
		Phe420	Ile
276	2 & 4	Arg128	Ala
		Phe420	Met
277	2 & 4	Arg128	Val
		Phe420	Ala
278	2 & 4	Arg128	Val
		Phe420	Leu
279	2 & 4	Arg128	Val
		Phe420	Val
280	2 & 4	Arg128	Val
		Phe420	Ile
281	2 & 4	Arg128	Val
		Phe420	Met
282	2 & 4	Arg128	Ile
		Phe420	Ala
283	2 & 4	Arg128	Ile
		Phe420	Leu

284	2 & 4	Arg128	Ile
		Phe420	Val
285	2 & 4	Arg128	Ile
		Phe420	Ile
286	2 & 4	Arg128	Ile
		Phe420	Met
287	2 & 4	Arg128	Met
		Phe420	Ala
288	2 & 4	Arg128	Met
		Phe420	Leu
289	2 & 4	Arg128	Met
		Phe420	Val
290	2 & 4	Arg128	Met
		Phe420	Ile
291	2 & 4	Arg128	Met
		Phe420	Met
292	2 & 4	Arg128	Tyr
		Phe420	Ala
293	2 & 4	Arg128	Tyr
		Phe420	Leu
294	2 & 4	Arg128	Tyr
		Phe420	Val
295	2 & 4	Arg128	Tyr
		Phe420	Ile
296	2 & 4	Arg128	Tyr
		Phe420	Met
297	2 & 4	Arg128	Gly
		Phe420	Ala
298	2 & 4	Arg128	Gly
		Phe420	Leu
299	2 & 4	Arg128	Gly
		Phe420	Val
300	2 & 4	Arg128	Gly
		Phe420	Ile
301	2 & 4	Arg128	Gly
		Phe420	Met
302	2 & 4	Arg128	Asn
		Phe420	Ala
303	2 & 4	Arg128	Asn
		Phe420	Leu
304	2 & 4	Arg128	Asn
		Phe420	Val
305	2 & 4	Arg128	Asn

		Phe420	Ile
306	2 & 4	Arg128	Asn
		Phe420	Met
307	2 & 4	Arg128	Cys
		Phe420	Ala
308	2 & 4	Arg128	Cys
		Phe420	Leu
309	2 & 4	Arg128	Cys
		Phe420	Val
310	2 & 4	Arg128	Cys
		Phe420	Ile
311	2 & 4	Arg128	Cys
		Phe420	Met
312	2 & 4	Arg128	Phe
		Phe420	Ala
313	2 & 4	Arg128	Phe
		Phe420	Leu
314	2 & 4	Arg128	Phe
		Phe420	Val
315	2 & 4	Arg128	Phe
		Phe420	Ile
316	2 & 4	Arg128	Phe
		Phe420	Met
317	2 & 4	Arg128	Ser
		Phe420	Ala
318	2 & 4	Arg128	Ser
		Phe420	Leu
319	2 & 4	Arg128	Ser
		Phe420	Val
320	2 & 4	Arg128	Ser
		Phe420	Ile
321	2 & 4	Arg128	Ser
		Phe420	Met
322	2 & 4	Arg128	Thr
		Phe420	Ala
323	2 & 4	Arg128	Thr
		Phe420	Leu
324	2 & 4	Arg128	Thr
		Phe420	Val
325	2 & 4	Arg128	Thr
		Phe420	Ile
326	2 & 4	Arg128	Thr
		Phe420	Met

327	2 & 4	Arg128	Gln
		Phe420	Ala
328	2 & 4	Arg128	Gln
		Phe420	Leu
329	2 & 4	Arg128	Gln
		Phe420	Val
330	2 & 4	Arg128	Gln
		Phe420	Ile
331	2 & 4	Arg128	Gln
		Phe420	Met
332	2 & 4	Arg128	His
		Phe420	Ala
333	2 & 4	Arg128	His
		Phe420	Leu
334	2 & 4	Arg128	His
		Phe420	Val
335	2 & 4	Arg128	His
		Phe420	Ile
336	2 & 4	Arg128	His
		Phe420	Met
337	24	Arg130	Leu
		Phe433	Ala
338	24	Arg130	Leu
		Phe433	Leu
339	24	Arg130	Leu
		Phe433	Val
340	24	Arg130	Leu
		Phe433	Ile
341	24	Arg130	Leu
		Phe433	Met
342	24	Arg130	Ala
		Phe433	Ala
343	24	Arg130	Ala
		Phe433	Leu
344	24	Arg130	Ala
		Phe433	Val
345	24	Arg130	Ala
		Phe433	Ile
346	24	Arg130	Ala
		Phe433	Met
347	24	Arg130	Val
		Phe433	Ala
348	24	Arg130	Val

		Phe433	Leu
349	24	Arg130	Val
		Phe433	Val
350	24	Arg130	Val
		Phe433	Ile
351	24	Arg130	Val
		Phe433	Met
352	24	Arg130	Ile
		Phe433	Ala
353	24	Arg130	Ile
		Phe433	Leu
354	24	Arg130	Ile
		Phe433	Val
355	24	Arg130	Ile
		Phe433	Ile
356	24	Arg130	Ile
		Phe433	Met
357	24	Arg130	Met
		Phe433	Ala
358	24	Arg130	Met
		Phe433	Leu
359	24	Arg130	Met
		Phe433	Val
360	24	Arg130	Met
		Phe433	Ile
361	24	Arg130	Met
		Phe433	Met
362	24	Arg130	Tyr
		Phe433	Ala
363	24	Arg130	Tyr
		Phe433	Leu
364	24	Arg130	Tyr
		Phe433	Val
365	24	Arg130	Tyr
		Phe433	Ile
366	24	Arg130	Tyr
		Phe433	Met
367	24	Arg130	Gly
		Phe433	Ala
368	24	Arg130	Gly
		Phe433	Leu
369	24	Arg130	Gly
		Phe433	Val

370	24	Arg130	Gly
		Phe433	Ile
371	24	Arg130	Gly
		Phe433	Met
372	24	Arg130	Asn
		Phe433	Ala
373	24	Arg130	Asn
		Phe433	Leu
374	24	Arg130	Asn
		Phe433	Val
375	24	Arg130	Asn
		Phe433	Ile
376	24	Arg130	Asn
		Phe433	Met
377	24	Arg130	Cys
		Phe433	Ala
378	24	Arg130	Cys
		Phe433	Leu
379	24	Arg130	Cys
		Phe433	Val
380	24	Arg130	Cys
		Phe433	Ile
381	24	Arg130	Cys
		Phe433	Met
382	24	Arg130	Phe
		Phe433	Ala
383	24	Arg130	Phe
		Phe433	Leu
384	24	Arg130	Phe
		Phe433	Val
385	24	Arg130	Phe
		Phe433	Ile
386	24	Arg130	Phe
		Phe433	Met
387	24	Arg130	Ser
		Phe433	Ala
388	24	Arg130	Ser
		Phe433	Leu
389	24	Arg130	Ser
		Phe433	Val
390	24	Arg130	Ser
		Phe433	Ile
391	24	Arg130	Ser

		Phe433	Met
392	24	Arg130	Thr
		Phe433	Ala
393	24	Arg130	Thr
		Phe433	Leu
394	24	Arg130	Thr
		Phe433	Val
395	24	Arg130	Thr
		Phe433	Ile
396	24	Arg130	Thr
		Phe433	Met
397	24	Arg130	Gln
		Phe433	Ala
398	24	Arg130	Gln
		Phe433	Leu
399	24	Arg130	Gln
		Phe433	Val
400	24	Arg130	Gln
		Phe433	Ile
401	24	Arg130	Gln
		Phe433	Met
402	24	Arg130	His
		Phe433	Ala
403	24	Arg130	His
		Phe433	Leu
404	24	Arg130	His
		Phe433	Val
405	24	Arg130	His
		Phe433	Ile
406	24	Arg130	His
		Phe433	Met
407	30	Arg130	Leu
		Phe433	Ala
408	30	Arg130	Leu
		Phe433	Leu
409	30	Arg130	Leu
		Phe433	Val
410	30	Arg130	Leu
		Phe433	Ile
411	30	Arg130	Leu
		Phe433	Met
412	30	Arg130	Ala
		Phe433	Ala

413	30	Arg130	Ala
		Phe433	Leu
414	30	Arg130	Ala
		Phe433	Val
415	30	Arg130	Ala
		Phe433	Ile
416	30	Arg130	Ala
		Phe433	Met
417	30	Arg130	Val
		Phe433	Ala
418	30	Arg130	Val
		Phe433	Leu
419	30	Arg130	Val
		Phe433	Val
420	30	Arg130	Val
		Phe433	Ile
421	30	Arg130	Val
		Phe433	Met
422	30	Arg130	Ile
		Phe433	Ala
423	30	Arg130	Ile
		Phe433	Leu
424	30	Arg130	Ile
		Phe433	Val
425	30	Arg130	Ile
		Phe433	Ile
426	30	Arg130	Ile
		Phe433	Met
427	30	Arg130	Met
		Phe433	Ala
428	30	Arg130	Met
		Phe433	Leu
429	30	Arg130	Met
		Phe433	Val
430	30	Arg130	Met
		Phe433	Ile
431	30	Arg130	Met
		Phe433	Met
432	30	Arg130	Tyr
		Phe433	Ala
433	30	Arg130	Tyr
		Phe433	Leu
434	30	Arg130	Tyr

		Phe433	Val
435	30	Arg130	Tyr
		Phe433	Ile
436	30	Arg130	Tyr
		Phe433	Met
437	30	Arg130	Gly
		Phe433	Ala
438	30	Arg130	Gly
		Phe433	Leu
439	30	Arg130	Gly
		Phe433	Val
440	30	Arg130	Gly
		Phe433	Ile
441	30	Arg130	Gly
		Phe433	Met
442	30	Arg130	Asn
		Phe433	Ala
443	30	Arg130	Asn
		Phe433	Leu
444	30	Arg130	Asn
		Phe433	Val
445	30	Arg130	Asn
		Phe433	Ile
446	30	Arg130	Asn
		Phe433	Met
447	30	Arg130	Cys
		Phe433	Ala
448	30	Arg130	Cys
		Phe433	Leu
449	30	Arg130	Cys
		Phe433	Val
450	30	Arg130	Cys
		Phe433	Ile
451	30	Arg130	Cys
		Phe433	Met
452	30	Arg130	Phe
		Phe433	Ala
453	30	Arg130	Phe
		Phe433	Leu
454	30	Arg130	Phe
		Phe433	Val
455	30	Arg130	Phe
		Phe433	Ile

456	30	Arg130	Phe
		Phe433	Met
457	30	Arg130	Ser
		Phe433	Ala
458	30	Arg130	Ser
		Phe433	Leu
459	30	Arg130	Ser
		Phe433	Val
460	30	Arg130	Ser
		Phe433	Ile
461	30	Arg130	Ser
		Phe433	Met
462	30	Arg130	Thr
		Phe433	Ala
463	30	Arg130	Thr
		Phe433	Leu
464	30	Arg130	Thr
		Phe433	Val
465	30	Arg130	Thr
		Phe433	Ile
466	30	Arg130	Thr
		Phe433	Met
467	30	Arg130	Gln
		Phe433	Ala
468	30	Arg130	Gln
		Phe433	Leu
469	30	Arg130	Gln
		Phe433	Val
470	30	Arg130	Gln
		Phe433	Ile
471	30	Arg130	Gln
		Phe433	Met
472	30	Arg130	His
		Phe433	Ala
473	30	Arg130	His
		Phe433	Leu
474	30	Arg130	His
		Phe433	Val
475	30	Arg130	His
		Phe433	Ile
476	30	Arg130	His
		Phe433	Met
477	38	Arg98	Leu

		Phe392	Ala
478	38	Arg98	Leu
		Phe392	Leu
479	38	Arg98	Leu
		Phe392	Val
480	38	Arg98	Leu
		Phe392	Ile
481	38	Arg98	Leu
		Phe392	Met
482	38	Arg98	Ala
		Phe392	Ala
483	38	Arg98	Ala
		Phe392	Leu
484	38	Arg98	Ala
		Phe392	Val
485	38	Arg98	Ala
		Phe392	Ile
486	38	Arg98	Ala
		Phe392	Met
487	38	Arg98	Val
		Phe392	Ala
488	38	Arg98	Val
		Phe392	Leu
489	38	Arg98	Val
		Phe392	Val
490	38	Arg98	Val
		Phe392	Ile
491	38	Arg98	Val
		Phe392	Met
492	38	Arg98	Ile
		Phe392	Ala
493	38	Arg98	Ile
		Phe392	Leu
494	38	Arg98	Ile
		Phe392	Val
495	38	Arg98	Ile
		Phe392	Ile
496	38	Arg98	Ile
		Phe392	Met
497	38	Arg98	Met
		Phe392	Ala
498	38	Arg98	Met
		Phe392	Leu

499	38	Arg98	Met
		Phe392	Val
500	38	Arg98	Met
		Phe392	Ile
501	38	Arg98	Met
		Phe392	Met
502	38	Arg98	Tyr
		Phe392	Ala
503	38	Arg98	Tyr
		Phe392	Leu
504	38	Arg98	Tyr
		Phe392	Val
505	38	Arg98	Tyr
		Phe392	Ile
506	38	Arg98	Tyr
		Phe392	Met
507	38	Arg98	Gly
		Phe392	Ala
508	38	Arg98	Gly
		Phe392	Leu
509	38	Arg98	Gly
		Phe392	Val
510	38	Arg98	Gly
		Phe392	Ile
511	38	Arg98	Gly
		Phe392	Met
512	38	Arg98	Asn
		Phe392	Ala
513	38	Arg98	Asn
		Phe392	Leu
514	38	Arg98	Asn
		Phe392	Val
515	38	Arg98	Asn
		Phe392	Ile
516	38	Arg98	Asn
		Phe392	Met
517	38	Arg98	Cys
		Phe392	Ala
518	38	Arg98	Cys
		Phe392	Leu
519	38	Arg98	Cys
		Phe392	Val
520	38	Arg98	Cys

		Phe392	Ile
521	38	Arg98	Cys
		Phe392	Met
522	38	Arg98	Phe
		Phe392	Ala
523	38	Arg98	Phe
		Phe392	Leu
524	38	Arg98	Phe
		Phe392	Val
525	38	Arg98	Phe
		Phe392	Ile
526	38	Arg98	Phe
		Phe392	Met
527	38	Arg98	Ser
		Phe392	Ala
528	38	Arg98	Ser
		Phe392	Leu
529	38	Arg98	Ser
		Phe392	Val
530	38	Arg98	Ser
		Phe392	Ile
531	38	Arg98	Ser
		Phe392	Met
532	38	Arg98	Thr
		Phe392	Ala
533	38	Arg98	Thr
		Phe392	Leu
534	38	Arg98	Thr
		Phe392	Val
535	38	Arg98	Thr
		Phe392	Ile
536	38	Arg98	Thr
		Phe392	Met
537	38	Arg98	Gln
		Phe392	Ala
538	38	Arg98	Gln
		Phe392	Leu
539	38	Arg98	Gln
		Phe392	Val
540	38	Arg98	Gln
		Phe392	Ile
541	38	Arg98	Gln
		Phe392	Met

542	38	Arg98	His
		Phe392	Ala
543	38	Arg98	His
		Phe392	Leu
544	38	Arg98	His
		Phe392	Val
545	38	Arg98	His
		Phe392	Ile
546	38	Arg98	His
		Phe392	Met
547	46	Arg139	Leu
		Phe465	Ala
548	46	Arg139	Leu
		Phe465	Leu
549	46	Arg139	Leu
		Phe465	Val
550	46	Arg139	Leu
		Phe465	Ile
551	46	Arg139	Leu
		Phe465	Met
552	46	Arg139	Ala
		Phe465	Ala
553	46	Arg139	Ala
		Phe465	Leu
554	46	Arg139	Ala
		Phe465	Val
555	46	Arg139	Ala
		Phe465	Ile
556	46	Arg139	Ala
		Phe465	Met
557	46	Arg139	Val
		Phe465	Ala
558	46	Arg139	Val
		Phe465	Leu
559	46	Arg139	Val
		Phe465	Val
560	46	Arg139	Val
		Phe465	Ile
561	46	Arg139	Val
		Phe465	Met
562	46	Arg139	Ile
		Phe465	Ala
563	46	Arg139	Ile

		Phe465	Leu
564	46	Arg139	Ile
		Phe465	Val
565	46	Arg139	Ile
		Phe465	Ile
566	46	Arg139	Ile
		Phe465	Met
567	46	Arg139	Met
		Phe465	Ala
568	46	Arg139	Met
		Phe465	Leu
569	46	Arg139	Met
		Phe465	Val
570	46	Arg139	Met
		Phe465	Ile
571	46	Arg139	Met
		Phe465	Met
572	46	Arg139	Tyr
		Phe465	Ala
573	46	Arg139	Tyr
		Phe465	Leu
574	46	Arg139	Tyr
		Phe465	Val
575	46	Arg139	Tyr
		Phe465	Ile
576	46	Arg139	Tyr
		Phe465	Met
577	46	Arg139	Gly
		Phe465	Ala
578	46	Arg139	Gly
		Phe465	Leu
579	46	Arg139	Gly
		Phe465	Val
580	46	Arg139	Gly
		Phe465	Ile
581	46	Arg139	Gly
		Phe465	Met
582	46	Arg139	Asn
		Phe465	Ala
583	46	Arg139	Asn
		Phe465	Leu
584	46	Arg139	Asn
		Phe465	Val

585	46	Arg139	Asn
		Phe465	Ile
586	46	Arg139	Asn
		Phe465	Met
587	46	Arg139	Cys
		Phe465	Ala
588	46	Arg139	Cys
		Phe465	Leu
589	46	Arg139	Cys
		Phe465	Val
590	46	Arg139	Cys
		Phe465	Ile
591	46	Arg139	Cys
		Phe465	Met
592	46	Arg139	Phe
		Phe465	Ala
593	46	Arg139	Phe
		Phe465	Leu
594	46	Arg139	Phe
		Phe465	Val
595	46	Arg139	Phe
		Phe465	Ile
596	46	Arg139	Phe
		Phe465	Met
597	46	Arg139	Ser
		Phe465	Ala
598	46	Arg139	Ser
		Phe465	Leu
599	46	Arg139	Ser
		Phe465	Val
600	46	Arg139	Ser
		Phe465	Ile
601	46	Arg139	Ser
		Phe465	Met
602	46	Arg139	Thr
		Phe465	Ala
603	46	Arg139	Thr
		Phe465	Leu
604	46	Arg139	Thr
		Phe465	Val
605	46	Arg139	Thr
		Phe465	Ile
606	46	Arg139	Thr

		Phe465	Met
607	46	Arg139	Gln
		Phe465	Ala
608	46	Arg139	Gln
		Phe465	Leu
609	46	Arg139	Gln
		Phe465	Val
610	46	Arg139	Gln
		Phe465	Ile
611	46	Arg139	Gln
		Phe465	Met
612	46	Arg139	His
		Phe465	Ala
613	46	Arg139	His
		Phe465	Leu
614	46	Arg139	His
		Phe465	Val
615	46	Arg139	His
		Phe465	Ile
616	46	Arg139	His
		Phe465	Met
617	48	Arg157	Leu
		Tyr439	Ala
618	48	Arg157	Leu
		Tyr439	Leu
619	48	Arg157	Leu
		Tyr439	Val
620	48	Arg157	Leu
		Tyr439	Ile
621	48	Arg157	Leu
		Tyr439	Met
622	48	Arg157	Ala
		Tyr439	Ala
623	48	Arg157	Ala
		Tyr439	Leu
624	48	Arg157	Ala
		Tyr439	Val
625	48	Arg157	Ala
		Tyr439	Ile
626	48	Arg157	Ala
		Tyr439	Met
627	48	Arg157	Val
		Tyr439	Ala

628	48	Arg157	Val
		Tyr439	Leu
629	48	Arg157	Val
		Tyr439	Val
630	48	Arg157	Val
		Tyr439	Ile
631	48	Arg157	Val
		Tyr439	Met
632	48	Arg157	Ile
		Tyr439	Ala
633	48	Arg157	Ile
		Tyr439	Leu
634	48	Arg157	Ile
		Tyr439	Val
635	48	Arg157	Ile
		Tyr439	Ile
636	48	Arg157	Ile
		Tyr439	Met
637	48	Arg157	Met
		Tyr439	Ala
638	48	Arg157	Met
		Tyr439	Leu
639	48	Arg157	Met
		Tyr439	Val
640	48	Arg157	Met
		Tyr439	Ile
641	48	Arg157	Met
		Tyr439	Met
642	48	Arg157	Tyr
		Tyr439	Ala
643	48	Arg157	Tyr
		Tyr439	Leu
644	48	Arg157	Tyr
		Tyr439	Val
645	48	Arg157	Tyr
		Tyr439	Ile
646	48	Arg157	Tyr
		Tyr439	Met
647	48	Arg157	Gly
		Tyr439	Ala
648	48	Arg157	Gly
		Tyr439	Leu
649	48	Arg157	Gly

		Tyr439	Val
650	48	Arg157	Gly
		Tyr439	Ile
651	48	Arg157	Gly
		Tyr439	Met
652	48	Arg157	Asn
		Tyr439	Ala
653	48	Arg157	Asn
		Tyr439	Leu
654	48	Arg157	Asn
		Tyr439	Val
655	48	Arg157	Asn
		Tyr439	Ile
656	48	Arg157	Asn
		Tyr439	Met
657	48	Arg157	Cys
		Tyr439	Ala
658	48	Arg157	Cys
		Tyr439	Leu
659	48	Arg157	Cys
		Tyr439	Val
660	48	Arg157	Cys
		Tyr439	Ile
661	48	Arg157	Cys
		Tyr439	Met
662	48	Arg157	Phe
		Tyr439	Ala
663	48	Arg157	Phe
		Tyr439	Leu
664	48	Arg157	Phe
		Tyr439	Val
665	48	Arg157	Phe
		Tyr439	Ile
666	48	Arg157	Phe
		Tyr439	Met
667	48	Arg157	Ser
		Tyr439	Ala
668	48	Arg157	Ser
		Tyr439	Leu
669	48	Arg157	Ser
		Tyr439	Val
670	48	Arg157	Ser
		Tyr439	Ile

671	48	Arg157	Ser
		Tyr439	Met
672	48	Arg157	Thr
		Tyr439	Ala
673	48	Arg157	Thr
		Tyr439	Leu
674	48	Arg157	Thr
		Tyr439	Val
675	48	Arg157	Thr
		Tyr439	Ile
676	48	Arg157	Thr
		Tyr439	Met
677	48	Arg157	Gln
		Tyr439	Ala
678	48	Arg157	Gln
		Tyr439	Leu
679	48	Arg157	Gln
		Tyr439	Val
680	48	Arg157	Gln
		Tyr439	Ile
681	48	Arg157	Gln
		Tyr439	Met
682	48	Arg157	His
		Tyr439	Ala
683	48	Arg157	His
		Tyr439	Leu
684	48	Arg157	His
		Tyr439	Val
685	48	Arg157	His
		Tyr439	Ile
686	48	Arg157	His
		Tyr439	Met

It is to be understood that any amino acid besides the ones mentioned in the above tables 3 could be used as a substituent. Assays to test for the functionality of such mutants are readily available in the art, and respectively, described in the Example section of the present invention.

5

In a preferred embodiment, the mutated PPO refers to a polypeptide of SEQ ID NO: 2 or SEQ ID NO: 4 in which the amino acid sequence differs from an amino acid sequence of SEQ ID NO: 2 or SEQ ID NO: 4 at position 128, and/or position 420.

10 Examples of differences at these amino acid positions include, but are not limited to, one or more of the following:

the amino acid at or corresponding to position 128 of SEQ ID NO:2 is other than Arginine;
the amino acid at or corresponding to position 420 of SEQ ID NO:2 is other than Phenylalanine,

5 In some embodiments, the mutated PPO enzyme of SEQ ID NO: 2 or SEQ ID NO: 4 comprises one or more of the following:

the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, Ala, Val, or Ile;
the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val, Met, Ala, Ile, or Leu;

10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, or His, and/or the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala, Leu, Val, Ile, or Met.

15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala, Leu, Val, Ile, Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or

corresponding to position 420 of SEQ ID NO:2 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 5 the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 10 the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 15 the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 20 the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 25 the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 30 the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 35 the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 40 the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Val, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ile, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a

variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Met, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

- 5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.
- 10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.
- 15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.
- 20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.
- 25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Tyr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.
- 30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.
- 35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.
- 40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gly, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Asn, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Cys, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Phe, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ser, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Thr, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Gln, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

5

In a particularly preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, or SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Met.

10

In another particularly preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, or SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ile.

15

In another particularly preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, or SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Leu.

20

In an especially preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 2, or SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Ala, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

25

In another especially preferred embodiment, the the mutated PPO comprises a sequence of SEQ ID NO: 2, or SEQ ID NO: 4, a variant, derivative, orthologue, paralogue or homologue thereof, in which the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Val.

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In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 130 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, or His, and the amino acid at or corresponding to position 433 is Ala, Leu, Val, Ile, or Met.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Ala.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Leu.

5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Val.

10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Ile.

15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Met.

20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Ala.

25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Leu.

30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Val.

35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Ile.

40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a

variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Ala.

- 5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Leu.
- 10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Val.
- 15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Ile.
- 20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Met.
- 25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Ala.
- 30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Leu.
- 35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Val.
- 40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Met.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Ala.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Leu.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Val.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Ile.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Met.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Ala.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Leu.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Ile.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Met.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Ala.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Leu.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Val.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Ile.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Met.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding to position 433 is Ala.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding to position 433 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding
5 to position 433 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding
10 to position 433 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding
15 to position 433 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding
20 to position 433 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding
25 to position 433 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding
30 to position 433 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding
35 to position 433 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding
40 to position 433 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding

to position 433 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 5 the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 10 the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 15 the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 20 the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 25 the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 30 the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 35 the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 40 the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Met.

5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Ala.

10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Leu.

15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Val.

20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Ile.

25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Met.

30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Ala.

35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Leu.

40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a

variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Ile.

- 5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Met.
- 10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Ala.
- 15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Leu.
- 20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Val.
- 25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Ile.
- 30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 24, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Met.
- 35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, His, and the amino acid at or corresponding to position 433 is Ala, Leu, Val, Ile, Met.
- 40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Leu.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Val.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Ile.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Leu, and the amino acid at or corresponding to position 433 is Met.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Ala.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Leu.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Val.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Ile.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ala, and the amino acid at or corresponding to position 433 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Ala.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Leu.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Val.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Ile.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Val, and the amino acid at or corresponding to position 433 is Met.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Ala.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Leu.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Val.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ile, and the amino acid at or corresponding to position 433 is Met.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Ala.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Leu.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Val.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Ile.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Met, and the amino acid at or corresponding to position 433 is Met.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Ala.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Leu.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Ile.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Tyr, and the amino acid at or corresponding to position 433 is Met.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Ala.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Leu.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Val.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Ile.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gly, and the amino acid at or corresponding to position 433 is Met.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding to position 433 is Ala.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding to position 433 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding to position 433 is Val.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding to position 433 is Ile.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Asn, and the amino acid at or corresponding to position 433 is Met.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding to position 433 is Ala.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding to position 433 is Leu.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding to position 433 is Val.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding to position 433 is Ile.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Cys, and the amino acid at or corresponding to position 433 is Met.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Leu.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Val.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Ile.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Phe, and the amino acid at or corresponding to position 433 is Met.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Ala.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Leu.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Val.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Ile.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Ser, and the amino acid at or corresponding to position 433 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Ala.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Leu.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Val.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Ile.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Thr, and the amino acid at or corresponding to position 433 is Met.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Ala.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Leu.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Val.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is Gln, and the amino acid at or corresponding to position 433 is Met.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Ala.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Leu.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Val.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Ile.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 30, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 130 is His, and the amino acid at or corresponding to position 433 is Met.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, His, and the amino acid at or corresponding to position 392 is Ala, Leu, Val, Ile, Met.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Leu, and the amino acid at or corresponding to position 392 is Ala.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Leu, and the amino acid at or corresponding to position 392 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Leu, and the amino acid at or corresponding to position 392 is Val.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Leu, and the amino acid at or corresponding to position 392 is Ile.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Leu, and the amino acid at or corresponding to position 392 is Met.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ala, and the amino acid at or corresponding to position 392 is Ala.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ala, and the amino acid at or corresponding to position 392 is Leu.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ala, and the amino acid at or corresponding to position 392 is Val.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ala, and the amino acid at or corresponding to position 392 is Ile.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ala, and the amino acid at or corresponding to position 392 is Met.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Val, and the amino acid at or corresponding to position 392 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Val, and the amino acid at or corresponding to position 392 is Leu.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Val, and the amino acid at or corresponding to position 392 is Val.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Val, and the amino acid at or corresponding to position 392 is Ile.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Val, and the amino acid at or corresponding to position 392 is Met.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ile, and the amino acid at or corresponding to position 392 is Ala.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ile, and the amino acid at or corresponding to position 392 is Leu.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ile, and the amino acid at or corresponding to position 392 is Val.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ile, and the amino acid at or corresponding to position 392 is Ile.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Ile, and the amino acid at or corresponding to position 392 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Met, and the amino acid at or corresponding to position 392 is Ala.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Met, and the amino acid at or corresponding to position 392 is Leu.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Met, and the amino acid at or corresponding to position 392 is Val.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Met, and the amino acid at or corresponding to position 392 is Ile.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Met, and the amino acid at or corresponding to position 392 is Met.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Tyr, and the amino acid at or corresponding to position 392 is Ala.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Tyr, and the amino acid at or corresponding to position 392 is Leu.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Tyr, and the amino acid at or corresponding to position 392 is Val.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Tyr, and the amino acid at or corresponding to position 392 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Tyr, and the amino acid at or corresponding to position 392 is Met.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gly, and the amino acid at or corresponding to position 392 is Ala.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gly, and the amino acid at or corresponding to position 392 is Leu.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gly, and the amino acid at or corresponding to position 392 is Val.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gly, and the amino acid at or corresponding to position 392 is Ile.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gly, and the amino acid at or corresponding to position 392 is Met.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Asn, and the amino acid at or corresponding to position 392 is Ala.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Asn, and the amino acid at or corresponding to position 392 is Leu.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Asn, and the amino acid at or corresponding to position 392 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Asn, and the amino acid at or corresponding to position 392 is Ile.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Asn, and the amino acid at or corresponding to position 392 is Met.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Cys, and the amino acid at or corresponding to position 392 is Ala.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Cys, and the amino acid at or corresponding to position 392 is Leu.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Cys, and the amino acid at or corresponding to position 392 is Val.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Cys, and the amino acid at or corresponding to position 392 is Ile.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Cys, and the amino acid at or corresponding to position 392 is Met.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Phe, and the amino acid at or corresponding to position 392 is Ala.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Phe, and the amino acid at or corresponding to position 392 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Phe, and the amino acid at or corresponding to position 392 is Val.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Phe, and the amino acid at or corresponding to position 392 is Ile.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Phe, and the amino acid at or corresponding to position 392 is Met.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Ser, and the amino acid at or corresponding to position 392 is Ala.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Ser, and the amino acid at or corresponding to position 392 is Leu.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Ser, and the amino acid at or corresponding to position 392 is Val.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Ser, and the amino acid at or corresponding to position 392 is Ile.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Ser, and the amino acid at or corresponding to position 392 is Met.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 98 is Thr, and the amino acid at or corresponding to position 392 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Thr, and the amino acid at or corresponding to position 392 is Leu.

- 5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Thr, and the amino acid at or corresponding to position 392 is Val.
- 10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Thr, and the amino acid at or corresponding to position 392 is Ile.
- 15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Thr, and the amino acid at or corresponding to position 392 is Met.
- 20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gln, and the amino acid at or corresponding to position 392 is Ala.
- 25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gln, and the amino acid at or corresponding to position 392 is Leu.
- 30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gln, and the amino acid at or corresponding to position 392 is Val.
- 35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gln, and the amino acid at or corresponding to position 392 is Ile.
- 40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is Gln, and the amino acid at or corresponding to position 392 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is His, and the amino acid at or corresponding to position 392 is Ala.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is His, and the amino acid at or corresponding to position 392 is Leu.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is His, and the amino acid at or corresponding to position 392 is Val.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is His, and the amino acid at or corresponding to position 392 is Ile.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 38, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 98 is His, and the amino acid at or corresponding to position 392 is Met.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, His, and the amino acid at or corresponding to position 465 is Ala, Leu, Val, Ile, Met.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Leu, and the amino acid at or corresponding to position 465 is Ala.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Leu, and the amino acid at or corresponding to position 465 is Leu.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Leu, and the amino acid at or corresponding

to position 465 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 5 the amino acid at or corresponding to position 139 is Leu, and the amino acid at or corresponding to position 465 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 10 the amino acid at or corresponding to position 139 is Leu, and the amino acid at or corresponding to position 465 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 15 the amino acid at or corresponding to position 139 is Ala, and the amino acid at or corresponding to position 465 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 20 the amino acid at or corresponding to position 139 is Ala, and the amino acid at or corresponding to position 465 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 25 the amino acid at or corresponding to position 139 is Ala, and the amino acid at or corresponding to position 465 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 30 the amino acid at or corresponding to position 139 is Ala, and the amino acid at or corresponding to position 465 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 35 the amino acid at or corresponding to position 139 is Ala, and the amino acid at or corresponding to position 465 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

- 40 the amino acid at or corresponding to position 139 is Val, and the amino acid at or corresponding to position 465 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 139 is Val, and the amino acid at or corresponding to position 465 is Leu.

5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Val, and the amino acid at or corresponding to position 465 is Val.

10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Val, and the amino acid at or corresponding to position 465 is Ile.

15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Val, and the amino acid at or corresponding to position 465 is Met.

20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ile, and the amino acid at or corresponding to position 465 is Ala.

25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ile, and the amino acid at or corresponding to position 465 is Leu.

30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ile, and the amino acid at or corresponding to position 465 is Val.

35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ile, and the amino acid at or corresponding to position 465 is Ile.

40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ile, and the amino acid at or corresponding to position 465 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a

variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Met, and the amino acid at or corresponding to position 465 is Ala.

- 5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Met, and the amino acid at or corresponding to position 465 is Leu.
- 10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Met, and the amino acid at or corresponding to position 465 is Val.
- 15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Met, and the amino acid at or corresponding to position 465 is Ile.
- 20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Met, and the amino acid at or corresponding to position 465 is Met.
- 25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Tyr, and the amino acid at or corresponding to position 465 is Ala.
- 30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Tyr, and the amino acid at or corresponding to position 465 is Leu.
- 35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Tyr, and the amino acid at or corresponding to position 465 is Val.
- 40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Tyr, and the amino acid at or corresponding to position 465 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Tyr, and the amino acid at or corresponding to position 465 is Met.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gly, and the amino acid at or corresponding to position 465 is Ala.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gly, and the amino acid at or corresponding to position 465 is Leu.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gly, and the amino acid at or corresponding to position 465 is Val.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gly, and the amino acid at or corresponding to position 465 is Ile.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gly, and the amino acid at or corresponding to position 465 is Met.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Asn, and the amino acid at or corresponding to position 465 is Ala.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Asn, and the amino acid at or corresponding to position 465 is Leu.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Asn, and the amino acid at or corresponding to position 465 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Asn, and the amino acid at or corresponding to position 465 is Ile.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Asn, and the amino acid at or corresponding to position 465 is Met.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Cys, and the amino acid at or corresponding to position 465 is Ala.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Cys, and the amino acid at or corresponding to position 465 is Leu.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Cys, and the amino acid at or corresponding to position 465 is Val.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Cys, and the amino acid at or corresponding to position 465 is Ile.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Cys, and the amino acid at or corresponding to position 465 is Met.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Phe, and the amino acid at or corresponding to position 465 is Ala.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Phe, and the amino acid at or corresponding to position 465 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Phe, and the amino acid at or corresponding to position 465 is Val.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Phe, and the amino acid at or corresponding to position 465 is Ile.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Phe, and the amino acid at or corresponding to position 465 is Met.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ser, and the amino acid at or corresponding to position 465 is Ala.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ser, and the amino acid at or corresponding to position 465 is Leu.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ser, and the amino acid at or corresponding to position 465 is Val.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ser, and the amino acid at or corresponding to position 465 is Ile.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Ser, and the amino acid at or corresponding to position 465 is Met.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Thr, and the amino acid at or corresponding to position 465 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Thr, and the amino acid at or corresponding to position 465 is Leu.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Thr, and the amino acid at or corresponding to position 465 is Val.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Thr, and the amino acid at or corresponding to position 465 is Ile.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Thr, and the amino acid at or corresponding to position 465 is Met.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gln, and the amino acid at or corresponding to position 465 is Ala.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gln, and the amino acid at or corresponding to position 465 is Leu.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gln, and the amino acid at or corresponding to position 465 is Val.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gln, and the amino acid at or corresponding to position 465 is Ile.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is Gln, and the amino acid at or corresponding to position 465 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is His, and the amino acid at or corresponding to position 465 is Ala.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is His, and the amino acid at or corresponding to position 465 is Leu.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is His, and the amino acid at or corresponding to position 465 is Val.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is His, and the amino acid at or corresponding to position 465 is Ile.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 46, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 139 is His, and the amino acid at or corresponding to position 465 is Met.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, His, and the amino acid at or corresponding to position 439 is Ala, Leu, Val, Ile, Met.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Leu, and the amino acid at or corresponding to position 439 is Ala.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Leu, and the amino acid at or corresponding to position 439 is Leu.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 157 is Leu, and the amino acid at or corresponding to position 439 is Val.

5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Leu, and the amino acid at or corresponding to position 439 is Ile.

10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Leu, and the amino acid at or corresponding to position 439 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
15 the amino acid at or corresponding to position 157 is Ala, and the amino acid at or corresponding to position 439 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
20 the amino acid at or corresponding to position 157 is Ala, and the amino acid at or corresponding to position 439 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
25 the amino acid at or corresponding to position 157 is Ala, and the amino acid at or corresponding to position 439 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
30 the amino acid at or corresponding to position 157 is Ala, and the amino acid at or corresponding to position 439 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
35 the amino acid at or corresponding to position 157 is Ala, and the amino acid at or corresponding to position 439 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
40 the amino acid at or corresponding to position 157 is Val, and the amino acid at or corresponding to position 439 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:

the amino acid at or corresponding to position 157 is Val, and the amino acid at or corresponding to position 439 is Leu.

5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Val, and the amino acid at or corresponding to position 439 is Val.

10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Val, and the amino acid at or corresponding to position 439 is Ile.

15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Val, and the amino acid at or corresponding to position 439 is Met.

20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ile, and the amino acid at or corresponding to position 439 is Ala.

25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ile, and the amino acid at or corresponding to position 439 is Leu.

30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ile, and the amino acid at or corresponding to position 439 is Val.

35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ile, and the amino acid at or corresponding to position 439 is Ile.

40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ile, and the amino acid at or corresponding to position 439 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a

variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Met, and the amino acid at or corresponding to position 439 is Ala.

- 5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Met, and the amino acid at or corresponding to position 439 is Leu.
- 10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Met, and the amino acid at or corresponding to position 439 is Val.
- 15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Met, and the amino acid at or corresponding to position 439 is Ile.
- 20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Met, and the amino acid at or corresponding to position 439 is Met.
- 25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Tyr, and the amino acid at or corresponding to position 439 is Ala.
- 30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Tyr, and the amino acid at or corresponding to position 439 is Leu.
- 35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Tyr, and the amino acid at or corresponding to position 439 is Val.
- 40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Tyr, and the amino acid at or corresponding to position 439 is Ile.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Tyr, and the amino acid at or corresponding to position 439 is Met.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gly, and the amino acid at or corresponding to position 439 is Ala.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gly, and the amino acid at or corresponding to position 439 is Leu.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gly, and the amino acid at or corresponding to position 439 is Val.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gly, and the amino acid at or corresponding to position 439 is Ile.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gly, and the amino acid at or corresponding to position 439 is Met.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Asn, and the amino acid at or corresponding to position 439 is Ala.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Asn, and the amino acid at or corresponding to position 439 is Leu.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Asn, and the amino acid at or corresponding to position 439 is Val.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Asn, and the amino acid at or corresponding to position 439 is Ile.

- 5 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Asn, and the amino acid at or corresponding to position 439 is Met.
- 10 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Cys, and the amino acid at or corresponding to position 439 is Ala.
- 15 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Cys, and the amino acid at or corresponding to position 439 is Leu.
- 20 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Cys, and the amino acid at or corresponding to position 439 is Val.
- 25 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Cys, and the amino acid at or corresponding to position 439 is Ile.
- 30 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Cys, and the amino acid at or corresponding to position 439 is Met.
- 35 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Phe, and the amino acid at or corresponding to position 439 is Ala.
- 40 In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Phe, and the amino acid at or corresponding to position 439 is Leu.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Phe, and the amino acid at or corresponding to position 439 is Val.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Phe, and the amino acid at or corresponding to position 439 is Ile.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Phe, and the amino acid at or corresponding to position 439 is Met.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ser, and the amino acid at or corresponding to position 439 is Ala.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ser, and the amino acid at or corresponding to position 439 is Leu.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ser, and the amino acid at or corresponding to position 439 is Val.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ser, and the amino acid at or corresponding to position 439 is Ile.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Ser, and the amino acid at or corresponding to position 439 is Met.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Thr, and the amino acid at or corresponding to position 439 is Ala.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Thr, and the amino acid at or corresponding to position 439 is Leu.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Thr, and the amino acid at or corresponding to position 439 is Val.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Thr, and the amino acid at or corresponding to position 439 is Ile.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Thr, and the amino acid at or corresponding to position 439 is Met.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gln, and the amino acid at or corresponding to position 439 is Ala.

25

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gln, and the amino acid at or corresponding to position 439 is Leu.

30

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gln, and the amino acid at or corresponding to position 439 is Val.

35

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gln, and the amino acid at or corresponding to position 439 is Ile.

40

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which:
the amino acid at or corresponding to position 157 is Gln, and the amino acid at or corresponding to position 439 is Met.

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 157 is His, and the amino acid at or corresponding to position 439 is Ala.

5

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 157 is His, and the amino acid at or corresponding to position 439 is Leu.

10

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 157 is His, and the amino acid at or corresponding to position 439 is Val.

15

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 157 is His, and the amino acid at or corresponding to position 439 is Ile.

20

In another preferred embodiment, the mutated PPO comprises a sequence of SEQ ID NO: 48, a variant, derivative, orthologue, paralogue or homologue thereof, in which: the amino acid at or corresponding to position 157 is His, and the amino acid at or corresponding to position 439 is Met.

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It will be within the knowledge of the skilled artisan to identify conserved regions and motifs shared between the homologues, orthologues and paralogues encoded by SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47, such as those depicted in Table 1. Having identified such conserved regions that may represent suitable binding motifs, amino acids corresponding to the amino acids listed in Table 3a and 3b, can be chosen to be substituted by any other amino acid, for example by conserved amino acids as shown in table 2, preferably by the amino acids of tables 3a and 3b.

30

Table 3c shows an overview of preferred mutation sites that are shared between homologues, orthologues and paralogues listed in Table 1.

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Table 3c

SEQ ID NO	Pos 1	Pos 2	Pos 3	Pos 4	Pos 5	Pos 6	Pos 7	Pos 8	Pos 9	Pos 10	Pos 11	Pos 12	Pos 13	Pos 14	Pos 15
2	N126	K127	R128	Y129	I130	A131	S149	I151	A154	P164	K169	E182	S183	E189	F196
4	N126	K127	R128	Y129	I130	A131	S149	I151	A154	P164	K169	E182	S183	E189	F196
6	N126	K127	R128	Y129	I130	A131	S149	I151	A154	P164	K169	E182	S183	E189	F196
8	N126	K127	R128	Y129	I130	A131	S149	I151	A154	P164	K169	E182	S183	E189	F196
10	K145	K146	R147	Y148	I149	V150	S168	V170	T173	P183	K188	E200	S201	Q207	V214
12	A153	P154	R155	F156	V157	L158	F176	L178	I181	F189	-	E203	S204	R210	V217
14	A160	P161	R162	F163	V164	L165	F183	L185	F188	F196	-	E210	S211	R217	V224
16	S167	P168	R169	F170	V171	L172	F190	L192	F195	L203	-	E217	S218	R224	V231
18	N125	K126	R127	Y128	I129	A130	S148	I150	A153	P163	K168	E181	S182	E188	F195
20	A162	P163	R164	F165	V166	L167	F185	L187	I190	F198	-	E212	S213	R219	V226
22	A140	P141	R142	F143	V144	L145	F163	L165	I168	L176	-	E190	S191	R197	V204
24	H128	K129	R130	Y131	I132	V133	S151	V153	T156	P166	T174	E187	S188	E194	V201
26	A165	P166	R167	F168	V169	W170	F187	L189	I192	L200	-	E215	S216	R222	V229
28	L128	P129	R130	W131	I132	L133	-	L152	T155	V165	-	E180	S181	R187	I194
30	H128	K129	R130	Y131	I132	V133	S151	V153	T156	P166	T174	E187	S188	E194	V201
32	A141	P142	R143	F144	V145	L146	F164	L166	I169	L177	-	E191	S192	R198	V205
34	N96	K97	R98	Y99	I100	A101	S119	I121	A124	P134	K139	E152	S153	E159	F166
36	A142	P143	R144	F145	V146	L147	F165	L167	I170	F178	-	E192	S193	R199	V206
38	N96	K97	R98	Y99	I100	A101	S119	F121	T124	P134	N139	E150	S151	Q157	V164
40	H96	K97	R98	Y99	I100	V101	S119	L121	A124	P134	R139	E152	S153	E159	V166
42	A28	P29	R30	F31	V32	L33	F51	L53	I56	F64	-	E78	S79	R85	V92
44	H93	K94	R95	Y96	I97	V98	S116	V118	T121	P131	R139	E152	S153	C158	V165
46	H137	K138	R139	Y140	I141	V142	S160	V162	T165	P175	R183	E196	S197	E203	V210
48	A155	P156	R157	F158	V159	L160	F178	L180	F183	L191	-	E205	S206	R212	V219

Table 3c continued

SEQ ID NO	Pos 16	Pos 17	Pos 18	Pos 19	Pos 20	Pos 21	Pos 22	Pos 23	Pos 24	Pos 25	Pos 26	Pos 27	Pos 28	Pos 29	Pos 30
2	D202	C209	G210	G211	L216	M218	H219	H220	N227	S234	S246	K259	P260	R261	L295
4	D202	C209	G210	G211	L216	M218	H219	H220	N227	S234	S246	K259	P260	R261	L295
6	D202	C209	G210	-	L215	M217	Y218	H219	N226	S233	S245	K258	P259	R260	L294
8	D202	C209	G210	-	L215	M217	H218	H219	N226	S233	S245	K258	P259	R260	L294
10	D220	S227	A228	A229	L234	M236	K237	H238	N245	S249	A261	K276	K277	G278	L312
12	E223	Y230	A231	G232	L237	M239	K240	A241	K248	G254	E266	K281	P282	K283	S316
14	E230	Y237	A238	G239	L244	M246	K247	A248	N255	G261	D273	K288	P289	K290	T323
16	E237	Y244	A245	G246	L251	M253	K254	A255	V262	G268	E280	K295	P296	K297	S330
18	D201	S208	G209	G210	L215	M217	R218	H219	N226	S233	S245	K259	P260	R261	L295
20	E232	Y239	A240	G241	L246	M248	K249	A250	K257	G263	E275	T290	P291	K292	S325
22	E210	Y217	A218	G219	L224	M226	K227	A228	R235	G241	E253	K268	P269	K270	T303
24	D207	S214	A215	G216	L221	I223	R224	H225	N232	S239	A251	R266	R267	N268	L302
26	E235	Y242	A243	G244	L249	M251	K252	A253	I260	G266	E278	K294	P295	K296	V329
28	E200	Y207	A208	G209	L214	M216	R217	A218	E225	G232	N244	S271	S272	S273	V306
30	D207	S214	A215	G216	L221	I223	C224	H225	N232	S239	A251	R266	R267	N268	L302
32	E211	Y218	A219	G220	L225	M227	K228	A229	R236	G242	E254	T269	P270	K271	T304
34	D172	C179	G180	G181	L186	M188	H189	H190	N197	S204	S216	K230	P231	R232	L266
36	E212	Y219	A220	G221	L226	M228	K229	A230	K237	G243	E255	K270	P271	Q272	S305
38	D170	C177	G178	G179	L184	M186	H187	H188	N195	S202	P214	K229	K230	R231	L265
40	D172	S179	A180	A181	L186	M188	R189	H190	N197	S204	A216	N231	K232	H233	L267
42	E98	Y105	A106	G107	L112	M114	K115	A116	R123	G129	E141	K156	P157	K158	S191
44	D171	S178	G179	G180	L185	I187	R188	H189	N196	S203	T215	G230	R231	N232	L266
46	D216	S223	G224	G225	L230	I232	R233	H234	N241	S248	T260	G275	R276	N277	L311
48	E225	Y232	A233	G234	L239	M241	K242	A243	T250	G256	E268	K283	P284	K285	S318

Table 3c continued

SEQ ID NO	Pos 31	Pos 32	Pos 33	Pos 34	Pos 35	Pos 36	Pos 37	Pos 38	Pos 39	Pos 40	Pos 41	Pos 42	Pos 43	Pos 44	Pos 45
2	Q301	G308	S324	R335	G346	F349	L351	D352	T358	L384	L397	F417	T418	T419	F420
4	Q301	G308	S324	R335	G346	F349	L351	D352	T358	L384	L397	F417	T418	T419	F420
6	Q300	G307	S323	R334	G345	F348	L350	D351	T357	L383	L396	F416	T417	T418	F419
8	Q300	G307	S323	R334	G345	F348	L350	D351	T357	L383	L396	F416	T417	T418	F419
10	S318	E323	R337	C348	G359	F362	L364	N365	N371	L397	L410	Y430	T431	T432	F433
12	E322	-	Q340	Y351	A365	L368	N370	F371	G377	L404	L414	L434	L435	N436	Y437
14	E329	-	Q347	Y358	A372	L375	K377	F378	A384	L411	L421	L441	L442	N443	Y444
16	S336	-	R354	Y365	A379	L382	K384	F385	A391	L418	L428	I448	L449	N450	Y451
18	H301	E308	P324	N335	E346	F349	L351	D352	S358	L384	L397	Y417	T418	T419	F420
20	E331	-	R349	Y360	A374	L377	S379	F380	A386	L413	L423	L443	L444	N445	Y446
22	D309	-	Q327	Y338	A352	L355	R357	F358	A364	L391	L401	L421	L422	N423	Y424
24	F308	G315	T336	S347	G358	V361	L363	D364	D370	L396	L410	Y430	T431	T432	F433
26	A335	-	F353	Y364	A378	L381	S383	F384	G390	L418	L428	L448	L449	N450	Y451
28	Q312	A319	V362	F373	A388	L391	E393	V394	A400	L430	L440	L460	L461	N462	F463
30	L308	G315	T336	S347	G358	F361	L363	D364	D370	L396	L410	Y430	T431	T432	F433
32	D310	-	Q328	Y339	A353	L356	I358	F359	A365	L392	L402	L422	L423	N424	Y425
34	Q272	G279	S295	R306	G317	F320	L322	D323	S329	L355	L368	F388	T389	T390	F391
36	E311	-	Q329	H340	A354	L357	K359	L360	A366	L393	L403	L423	L424	N425	Y426
38	C271	D278	S296	C307	G318	F321	L323	N324	D330	L356	L369	Y389	T390	T391	F392
40	H273	Q280	D294	Y305	G316	F319	L321	N322	S328	L354	L367	Y387	T388	T389	F390
42	D197	-	L215	Y226	A240	L243	K245	F246	A252	L279	L289	L309	L310	N311	Y312
44	C272	G279	S300	S311	G322	F325	L327	D328	D334	L360	L374	Y394	T395	S396	F397

46		C317	G324	S345	S356	G367	F370	L372	D373	D379	L405	L419	Y462	T463	S464	F465
48		L324	-	R342	Y353	A367	L370	K372	F373	A379	L406	L416	I436	L437	S438	Y439

Table 3c continued

SEQ ID NO	Pos 46	Pos 47	Pos 48	Pos 49	Pos 50	Pos 51	Pos 52	Pos 53	Pos 54	Pos 55	Pos 56	Pos 57	Pos 58	Pos 59
2	A432	T434	K438	L449	T451	F462	Y470	S476	V477	D482	Y493	K498	E515	K528
4	A432	T434	K438	L449	T451	F462	Y470	S476	V477	D482	Y493	K498	E515	K528
6	A431	T433	K437	L448	T450	F461	Y469	S475	V476	D481	Y492	K497	E514	K527
8	A431	T433	K437	L448	T450	F461	Y469	C475	V476	D481	Y492	K497	E514	K527
10	A445	T447	K451	L462	V464	Y475	Y483	S489	V490	D495	Y506	R511	D528	K541
12	K449	E451	V455	K468	K470	V481	F489	D495	T496	K501	L514	V519	S536	-
14	K456	E458	V462	R475	D477	V488	F496	D502	I503	K508	L521	V526	A543	-
16	K463	K465	A469	N482	N484	V495	F503	D509	L510	K515	L528	V533	A550	-
18	A432	T434	K438	L449	T451	Y462	Y470	S476	V477	E482	Y493	K498	E515	K525
20	K458	E460	V464	K477	K479	V490	F498	D504	T505	K510	L523	V528	S545	-
22	K436	E438	V442	N455	T457	V468	F476	D482	L483	K488	L501	V506	S523	-
24	A445	T447	K451	L462	V464	Y475	Y483	S489	V490	E495	Y506	K511	D528	N541
26	Q463	T465	V469	K482	D484	V495	F503	E509	Q510	R515	L528	V533	A550	A563
28	A475	P477	A481	R495	G497	V508	F516	D522	R523	K528	L545	V550	E567	-
30	A445	T447	K451	L462	V464	Y475	Y483	S489	V490	E495	Y506	K511	D528	N541
32	K437	E439	V443	N456	K458	V469	F477	D483	H484	K489	L502	V507	S524	-
34	A403	T405	K409	L420	T422	F433	Y441	S447	V448	D453	Y464	K469	E486	K499
36	K438	E440	V444	-	-	-	F447	D453	I454	K459	L472	V477	I494	-
38	A404	R406	K410	L421	A423	Y434	Y442	S448	V449	D454	Y465	R470	D487	-
40	A402	T404	R408	L419	A421	Y432	Y440	S446	V447	D452	F463	K468	D485	T498
42	Q324	E326	I330	N343	N345	V356	F364	D370	V371	K376	L389	V394	-	-
44	A409	T411	K415	L426	V428	H439	Y447	L453	V454	A459	Y470	K475	D492	D505

46	A477	T479	K483	L494	V496	H507	Y515	L521	V522	A527	Y538	K543	D560	D573
48	K451	E453	A457	N470	N472	V483	F491	D497	V498	K503	L516	V521	S538	-

In addition, the present invention refers to a method for identifying a PPO-inhibiting herbicide by using a mutated PPO encoded by a nucleic acid which comprises the nucleotide sequence of SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47, or a variant or derivative thereof.

5

Said method comprises the steps of:

- a) generating a transgenic cell or plant comprising a nucleic acid encoding a mutated PPO, wherein the mutated PPO is expressed;
- b) applying a PPO-inhibiting herbicide to the transgenic cell or plant of a) and to a control cell or plant of the same variety;
- c) determining the growth or the viability of the transgenic cell or plant and the control cell or plant after application of said PPO-inhibiting herbicide, and
- d) selecting "PPO-inhibiting herbicides" which confer reduced growth to the control cell or plant as compared to the growth of the transgenic cell or plant.

15

By "control cell" or "similar, wild-type, plant, plant tissue, plant cell or host cell" is intended a plant, plant tissue, plant cell, or host cell, respectively, that lacks the herbicide-resistance characteristics and/or particular polynucleotide of the invention that are disclosed herein. The use of the term "wild-type" is not, therefore, intended to imply that a plant, plant tissue, plant cell, or other host cell lacks recombinant DNA in its genome, and/or does not possess herbicide-resistant characteristics that are different from those disclosed herein.

20

Another object refers to a method of identifying a nucleotide sequence encoding a mutated PPO which is resistant or tolerant to a PPO-inhibiting herbicide, the method comprising:

- a) generating a library of mutated PPO-encoding nucleic acids,
- b) screening a population of the resulting mutated PPO-encoding nucleic acids by expressing each of said nucleic acids in a cell or plant and treating said cell or plant with a PPO-inhibiting herbicide,
- c) comparing the PPO-inhibiting herbicide-tolerance levels provided by said population of mutated PPO encoding nucleic acids with the PPO-inhibiting herbicide-tolerance level provided by a control PPO-encoding nucleic acid,
- d) selecting at least one mutated PPO-encoding nucleic acid that provides a significantly increased level of tolerance to a PPO-inhibiting herbicide as compared to that provided by the control PPO-encoding nucleic acid.

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In a preferred embodiment, the mutated PPO-encoding nucleic acid selected in step d) provides at least 2-fold as much resistance or tolerance of a cell or plant to a PPO-inhibiting herbicide as compared to that provided by the control PPO-encoding nucleic acid.

In a further preferred embodiment, the mutated PPO-encoding nucleic acid selected in step d) provides at least 2-fold, at least 5-fold, at least 10-fold, at least 20-fold, at least 50-fold, at least 100-fold, at least 500-fold, as much resistance or tolerance of a cell or plant to a PPO-inhibiting herbicide as compared to that provided by the control PPO-encoding nucleic acid.

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The resistance or tolerance can be determined by generating a transgenic plant or host cell, preferably a plant cell, comprising a nucleic acid sequence of the library of step a) and comparing said transgenic plant with a control plant or host cell, preferably a plant cell.

- 5 Another object refers to a method of identifying a plant or algae containing a nucleic acid comprising a nucleotide sequence encoding a wild-type or mutated PPO which is resistant or tolerant to a PPO-inhibiting herbicide, the method comprising:
- a) identifying an effective amount of a PPO-inhibiting herbicide in a culture of plant cells or green algae that leads to death of said cells.
 - 10 b) treating said plant cells or green algae with a mutagenizing agent,
 - c) contacting said mutagenized cells population with an effective amount of PPO-inhibiting herbicide, identified in a),
 - d) selecting at least one cell surviving these test conditions,
 - e) PCR-amplification and sequencing of PPO genes from cells selected in d) and comparing
15 such sequences to wild-type PPO gene sequences, respectively.

In a preferred embodiment, said mutagenizing agent is ethylmethanesulfonate (EMS).

- 20 Many methods well known to the skilled artisan are available for obtaining suitable candidate nucleic acids for identifying a nucleotide sequence encoding a mutated PPO from a variety of different potential source organisms including microbes, plants, fungi, algae, mixed cultures etc. as well as environmental sources of DNA such as soil. These methods include inter alia the preparation of cDNA or genomic DNA libraries, the use of suitably degenerate oligonucleotide primers, the use of probes based upon known sequences or complementation assays (for
25 example, for growth upon tyrosine) as well as the use of mutagenesis and shuffling in order to provide recombined or shuffled mutated PPO-encoding sequences.

- Nucleic acids comprising candidate and control PPO encoding sequences can be expressed in yeast, in a bacterial host strain, in an alga or in a higher plant such as tobacco or Arabidopsis and
30 the relative levels of inherent tolerance of the PPO encoding sequences screened according to a visible indicator phenotype of the transformed strain or plant in the presence of different concentrations of the selected PPO-inhibiting herbicide. Dose responses and relative shifts in dose responses associated with these indicator phenotypes (formation of brown color, growth inhibition, herbicidal effect etc) are conveniently expressed in terms, for example, of GR50 (concentration for
35 50% reduction of growth) or MIC (minimum inhibitory concentration) values where increases in values correspond to increases in inherent tolerance of the expressed PPO. For example, in a relatively rapid assay system based upon transformation of a bacterium such as E. coli, each mutated PPO encoding sequence may be expressed, for example, as a DNA sequence under expression control of a controllable promoter such as the lacZ promoter and taking suitable
40 account, for example by the use of synthetic DNA, of such issues as codon usage in order to obtain as comparable a level of expression as possible of different PPO sequences. Such strains expressing nucleic acids comprising alternative candidate PPO sequences may be plated out on different concentrations of the selected PPO-inhibiting herbicide in, optionally, a tyrosine supplemented medium and the relative levels of inherent tolerance of the expressed PPO enzymes

estimated on the basis of the extent and MIC for inhibition of the formation of the brown, ochronotic pigment.

5 In another embodiment, candidate nucleic acids are transformed into plant material to generate a transgenic plant, regenerated into morphologically normal fertile plants which are then measured for differential tolerance to selected PPO-inhibiting herbicides as described in the Example section hereinafter. Many suitable methods for transformation using suitable selection markers such as kanamycin, binary vectors such as from *Agrobacterium* and plant regeneration as, for example, from tobacco leaf discs are well known in the art. Optionally, a control population of plants is
10 likewise transformed with a nucleic acid expressing the control PPO. Alternatively, an untransformed dicot plant such as *Arabidopsis* or Tobacco can be used as a control since this, in any case, expresses its own endogenous PPO. The average, and distribution, of herbicide tolerance levels of a range of primary plant transformation events or their progeny to PPO-inhibiting herbicides described supra are evaluated in the normal manner based upon plant
15 damage, meristematic bleaching symptoms etc. at a range of different concentrations of herbicides. These data can be expressed in terms of, for example, GR50 values derived from dose/response curves having "dose" plotted on the x-axis and "percentage kill", "herbicidal effect", "numbers of emerging green plants" etc. plotted on the y-axis where increased GR50 values correspond to increased levels of inherent tolerance of the expressed PPO. Herbicides can
20 suitably be applied pre-emergence or post-emergence.

Another object of the present invention refers to an isolated nucleic acid encoding a mutated PPO as disclosed SUPRA, wherein the nucleic acid comprises the nucleotide sequence of SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45 or 47, or a variant
25 or derivative thereof.

In one embodiment, the nucleic acid is identifiable by a method as defined above.

In a preferred embodiment, the encoded mutated PPO is a variant of SEQ ID NO: 2 or SEQ ID NO: 4, or an orthologue thereof, which includes one or more of the following: the amino acid at or
30 corresponding to position 128 of SEQ ID NO:2 is other than Arginine; and/or the amino acid at or corresponding to position 420 of SEQ ID NO:2 is other than Phenylalanine.

In another embodiment, the invention refers to a plant cell transformed by a nucleic acid encoding
35 a mutated PPO polypeptide according to the present invention or to a plant cell which has been mutated to obtain a plant expressing a nucleic acid encoding a mutated PPO polypeptide according to the present invention, wherein expression of the nucleic acid in the plant cell results in increased resistance or tolerance to a PPO-inhibiting herbicide as compared to a wild type variety of the plant cell. Preferably, the mutated PPO polypeptide encoding nucleic acid comprises a
40 polynucleotide sequence selected from the group consisting of: a) a polynucleotide as shown in SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45 or 47, or a variant or derivative thereof; b) a polynucleotide encoding a polypeptide as shown in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, or a variant or derivative thereof; c) a polynucleotide comprising at least 60 consecutive nucleotides of

any of a) or b); and d) a polynucleotide complementary to the polynucleotide of any of a) through c).

The term “expression/expressing” or “gene expression” means the transcription of a specific gene or specific genes or specific genetic construct. The term “expression” or “gene expression” in particular means the transcription of a gene or genes or genetic construct into structural RNA (rRNA, tRNA) or mRNA with or without subsequent translation of the latter into a protein. The process includes transcription of DNA and processing of the resulting mRNA product.

To obtain the desired effect, i.e. plants that are tolerant or resistant to the PPO-inhibiting herbicide derivative herbicide of the present invention, it will be understood that the at least one nucleic acid is “over-expressed” by methods and means known to the person skilled in the art.

The term “increased expression” or “overexpression” as used herein means any form of expression that is additional to the original wild-type expression level. Methods for increasing expression of genes or gene products are well documented in the art and include, for example, overexpression driven by appropriate promoters, the use of transcription enhancers or translation enhancers. Isolated nucleic acids which serve as promoter or enhancer elements may be introduced in an appropriate position (typically upstream) of a non-heterologous form of a polynucleotide so as to upregulate expression of a nucleic acid encoding the polypeptide of interest. For example, endogenous promoters may be altered in vivo by mutation, deletion, and/or substitution (see, Kmiec, US 5,565,350; Zarling et al., WO9322443), or isolated promoters may be introduced into a plant cell in the proper orientation and distance from a gene of the present invention so as to control the expression of the gene.

If polypeptide expression is desired, it is generally desirable to include a polyadenylation region at the 3'-end of a polynucleotide coding region. The polyadenylation region can be derived from the natural gene, from a variety of other plant genes, or from T-DNA. The 3' end sequence to be added may be derived from, for example, the nopaline synthase or octopine synthase genes, or alternatively from another plant gene, or less preferably from any other eukaryotic gene.

An intron sequence may also be added to the 5' untranslated region (UTR) or the coding sequence of the partial coding sequence to increase the amount of the mature message that accumulates in the cytosol. Inclusion of a spliceable intron in the transcription unit in both plant and animal expression constructs has been shown to increase gene expression at both the mRNA and protein levels up to 1000-fold (Buchman and Berg (1988) Mol. Cell Biol. 8: 4395-4405; Callis et al. (1987) Genes Dev 1:1183-1200). Such intron enhancement of gene expression is typically greatest when placed near the 5' end of the transcription unit. Use of the maize introns Adh1-S intron 1, 2, and 6, the Bronze-1 intron are known in the art. For general information see: The Maize Handbook, Chapter 116, Freeling and Walbot, Eds., Springer, N.Y. (1994)

The term “introduction” or “transformation” as referred to herein encompasses the transfer of an exogenous polynucleotide into a host cell, irrespective of the method used for transfer. Plant tissue capable of subsequent clonal propagation, whether by organogenesis or embryogenesis, may be

transformed with a genetic construct of the present invention and a whole plant regenerated there from. The particular tissue chosen will vary depending on the clonal propagation systems available for, and best suited to, the particular species being transformed. Exemplary tissue targets include leaf disks, pollen, embryos, cotyledons, hypocotyls, megagametophytes, callus tissue, existing
5 meristematic tissue (e.g., apical meristem, axillary buds, and root meristems), and induced meristem tissue (e.g., cotyledon meristem and hypocotyl meristem). The polynucleotide may be transiently or stably introduced into a host cell and may be maintained non-integrated, for example, as a plasmid. Alternatively, it may be integrated into the host genome. The resulting transformed plant cell may then be used to regenerate a transformed plant in a manner known to persons
10 skilled in the art.

The transfer of foreign genes into the genome of a plant is called transformation. Transformation of plant species is now a fairly routine technique. Advantageously, any of several transformation methods may be used to introduce the gene of interest into a suitable ancestor cell. The methods
15 described for the transformation and regeneration of plants from plant tissues or plant cells may be utilized for transient or for stable transformation. Transformation methods include the use of liposomes, electroporation, chemicals that increase free DNA uptake, injection of the DNA directly into the plant, particle gun bombardment, transformation using viruses or pollen and microprojection. Methods may be selected from the calcium/polyethylene glycol method for
20 protoplasts (Krens, F.A. et al., (1982) *Nature* 296, 72-74; Negrutiu I et al. (1987) *Plant Mol Biol* 8: 363-373); electroporation of protoplasts (Shillito R.D. et al. (1985) *Bio/Technol* 3, 1099-1102); microinjection into plant material (Crossway A et al., (1986) *Mol. Gen Genet* 202: 179-185); DNA or RNA-coated particle bombardment (Klein TM et al., (1987) *Nature* 327: 70) infection with (non-integrative) viruses and the like. Transgenic plants, including transgenic crop plants, are preferably
25 produced via *Agrobacterium*-mediated transformation. An advantageous transformation method is the transformation in planta. To this end, it is possible, for example, to allow the *agrobacteria* to act on plant seeds or to inoculate the plant meristem with *agrobacteria*. It has proved particularly expedient in accordance with the invention to allow a suspension of transformed *agrobacteria* to act on the intact plant or at least on the flower primordia. The plant is subsequently grown on until
30 the seeds of the treated plant are obtained (Clough and Bent, *Plant J.* (1998) 16, 735-743). Methods for *Agrobacterium*-mediated transformation of rice include well known methods for rice transformation, such as those described in any of the following: European patent application EP 1198985 A1, Aldemita and Hodges (*Planta* 199: 612-617, 1996); Chan et al. (*Plant Mol Biol* 22 (3): 491-506, 1993), Hiei et al. (*Plant J* 6 (2): 271-282, 1994), which disclosures are incorporated by
35 reference herein as if fully set forth. In the case of corn transformation, the preferred method is as described in either Ishida et al. (*Nat. Biotechnol* 14(6): 745-50, 1996) or Frame et al. (*Plant Physiol* 129(1): 13-22, 2002), which disclosures are incorporated by reference herein as if fully set forth. Said methods are further described by way of example in B. Jenes et al., *Techniques for Gene Transfer*, in: *Transgenic Plants*, Vol. 1, Engineering and Utilization, eds. S.D. Kung and R. Wu, Academic Press (1993) 128-143 and in Potrykus *Annu. Rev. Plant Physiol. Plant Molec. Biol.* 42
40 (1991) 205-225). The nucleic acids or the construct to be expressed is preferably cloned into a vector, which is suitable for transforming *Agrobacterium tumefaciens*, for example pBin19 (Bevan et al., *Nucl. Acids Res.* 12 (1984) 8711). *Agrobacteria* transformed by such a vector can then be used in known manner for the transformation of plants, such as plants used as a model, like

Arabidopsis (*Arabidopsis thaliana* is within the scope of the present invention not considered as a crop plant), or crop plants such as, by way of example, tobacco plants, for example by immersing bruised leaves or chopped leaves in an agrobacterial solution and then culturing them in suitable media. The transformation of plants by means of *Agrobacterium tumefaciens* is described, for example, by Höfgen and Willmitzer in Nucl. Acid Res. (1988) 16, 9877 or is known inter alia from F.F. White, Vectors for Gene Transfer in Higher Plants; in Transgenic Plants, Vol. 1, Engineering and Utilization, eds. S.D. Kung and R. Wu, Academic Press, 1993, pp. 15-38.

In addition to the transformation of somatic cells, which then have to be regenerated into intact plants, it is also possible to transform the cells of plant meristems and in particular those cells which develop into gametes. In this case, the transformed gametes follow the natural plant development, giving rise to transgenic plants. Thus, for example, seeds of *Arabidopsis* are treated with agrobacteria and seeds are obtained from the developing plants of which a certain proportion is transformed and thus transgenic [Feldman, KA and Marks MD (1987). Mol Gen Genet 208:274-289; Feldmann K (1992). In: C Koncz, N-H Chua and J Shell, eds, Methods in *Arabidopsis* Research. Word Scientific, Singapore, pp. 274-289]. Alternative methods are based on the repeated removal of the inflorescences and incubation of the excision site in the center of the rosette with transformed agrobacteria, whereby transformed seeds can likewise be obtained at a later point in time (Chang (1994). Plant J. 5: 551-558; Katavic (1994). Mol Gen Genet, 245: 363-370). However, an especially effective method is the vacuum infiltration method with its modifications such as the "floral dip" method. In the case of vacuum infiltration of *Arabidopsis*, intact plants under reduced pressure are treated with an agrobacterial suspension [Bechthold, N (1993). C R Acad Sci Paris Life Sci, 316: 1194-1199], while in the case of the "floral dip" method the developing floral tissue is incubated briefly with a surfactant-treated agrobacterial suspension [Clough, SJ and Bent AF (1998) The Plant J. 16, 735-743]. A certain proportion of transgenic seeds are harvested in both cases, and these seeds can be distinguished from non-transgenic seeds by growing under the above-described selective conditions. In addition the stable transformation of plastids is of advantages because plastids are inherited maternally in most crops reducing or eliminating the risk of transgene flow through pollen. The transformation of the chloroplast genome is generally achieved by a process which has been schematically displayed in Klaus et al., 2004 [Nature Biotechnology 22 (2), 225-229]. Briefly the sequences to be transformed are cloned together with a selectable marker gene between flanking sequences homologous to the chloroplast genome. These homologous flanking sequences direct site specific integration into the plastome. Plastid transformation has been described for many different plant species and an overview is given in Bock (2001) Transgenic plastids in basic research and plant biotechnology. J Mol Biol. 2001 Sep 21; 312 (3):425-38 or Maliga, P (2003) Progress towards commercialization of plastid transformation technology. Trends Biotechnol. 21, 20-28. Further biotechnological progress has recently been reported in form of marker free plastid transformants, which can be produced by a transient co-integrated marker gene (Klaus et al., 2004, Nature Biotechnology 22(2), 225-229). The genetically modified plant cells can be regenerated via all methods with which the skilled worker is familiar. Suitable methods can be found in the abovementioned publications by S.D. Kung and R. Wu, Potrykus or Höfgen and Willmitzer.

Generally after transformation, plant cells or cell groupings are selected for the presence of one or

more markers which are encoded by plant-expressible genes co-transferred with the gene of interest, following which the transformed material is regenerated into a whole plant. To select transformed plants, the plant material obtained in the transformation is, as a rule, subjected to selective conditions so that transformed plants can be distinguished from untransformed plants.

For example, the seeds obtained in the above-described manner can be planted and, after an initial growing period, subjected to a suitable selection by spraying. A further possibility consists in growing the seeds, if appropriate after sterilization, on agar plates using a suitable selection agent so that only the transformed seeds can grow into plants. Alternatively, the transformed plants are screened for the presence of a selectable marker such as the ones described above.

Following DNA transfer and regeneration, putatively transformed plants may also be evaluated, for instance using Southern analysis, for the presence of the gene of interest, copy number and/or genomic organisation. Alternatively or additionally, expression levels of the newly introduced DNA may be monitored using Northern and/or Western analysis, both techniques being well known to persons having ordinary skill in the art.

The generated transformed plants may be propagated by a variety of means, such as by clonal propagation or classical breeding techniques. For example, a first generation (or T1) transformed plant may be selfed and homozygous second-generation (or T2) transformants selected, and the T2 plants may then further be propagated through classical breeding techniques. The generated transformed organisms may take a variety of forms. For example, they may be chimeras of transformed cells and non-transformed cells; clonal transformants (e.g., all cells transformed to contain the expression cassette); grafts of transformed and untransformed tissues (e.g., in plants, a transformed rootstock grafted to an untransformed scion).

Preferably, the wild-type or mutated PPO nucleic acid comprises a polynucleotide sequence selected from the group consisting of : a) a polynucleotide as shown in SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47, or a variant or derivative thereof; b) a polynucleotide encoding a polypeptide as shown in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, or a variant or derivative thereof; c) a polynucleotide comprising at least 60 consecutive nucleotides of any of a) or b); and d) a polynucleotide complementary to the polynucleotide of any of a) through c).

Preferably, the expression of the nucleic acid in the plant results in the plant's increased resistance to PPO-inhibiting herbicide as compared to a wild type variety of the plant.

In another embodiment, the invention refers to a plant, preferably a transgenic plant, comprising a plant cell according to the present invention, wherein expression of the nucleic acid in the plant results in the plant's increased resistance to PPO-inhibiting herbicide as compared to a wild type variety of the plant.

The plants described herein can be either transgenic crop plants or non-transgenic plants.

For the purposes of the invention, "transgenic", "transgene" or "recombinant" means with regard to,

for example, a nucleic acid sequence, an expression cassette, gene construct or a vector comprising the nucleic acid sequence or an organism transformed with the nucleic acid sequences, expression cassettes or vectors according to the invention, all those constructions brought about by recombinant methods in which either

- 5 (a) the nucleic acid sequences encoding proteins useful in the methods of the invention, or
- (b) genetic control sequence(s) which is operably linked with the nucleic acid sequence according to the invention, for example a promoter, or
- (c) a) and b)

are not located in their natural genetic environment or have been modified by recombinant
10 methods, it being possible for the modification to take the form of, for example, a substitution, addition, deletion, inversion or insertion of one or more nucleotide residues in order to allow for the expression of the mutated PPO of the present invention. The natural genetic environment is understood as meaning the natural genomic or chromosomal locus in the original plant or the presence in a genomic library. In the case of a genomic library, the natural genetic environment of
15 the nucleic acid sequence is preferably retained, at least in part. The environment flanks the nucleic acid sequence at least on one side and has a sequence length of at least 50 bp, preferably at least 500 bp, especially preferably at least 1000 bp, most preferably at least 5000 bp. A naturally occurring expression cassette – for example the naturally occurring combination of the natural promoter of the nucleic acid sequences with the corresponding nucleic acid sequence encoding a
20 polypeptide useful in the methods of the present invention, as defined above – becomes a transgenic expression cassette when this expression cassette is modified by non-natural, synthetic ("artificial") methods such as, for example, mutagenic treatment. Suitable methods are described, for example, in US 5,565,350 or WO 00/15815.

25 A transgenic plant for the purposes of the invention is thus understood as meaning, as above, that the nucleic acids of the invention are not at their natural locus in the genome of said plant, it being possible for the nucleic acids to be expressed homologously or heterologously. However, as mentioned, transgenic also means that, while the nucleic acids according to the invention or used in the inventive method are at their natural position in the genome of a plant, the sequence has
30 been modified with regard to the natural sequence, and/or that the regulatory sequences of the natural sequences have been modified. Transgenic is preferably understood as meaning the expression of the nucleic acids according to the invention at an unnatural locus in the genome, i.e. homologous or, preferably, heterologous expression of the nucleic acids takes place. Preferred transgenic plants are mentioned herein. Furthermore, the term "transgenic" refers to any plant,
35 plant cell, callus, plant tissue, or plant part, that contains all or part of at least one recombinant polynucleotide. In many cases, all or part of the recombinant polynucleotide is stably integrated into a chromosome or stable extra-chromosomal element, so that it is passed on to successive generations. For the purposes of the invention, the term "recombinant polynucleotide" refers to a polynucleotide that has been altered, rearranged, or modified by genetic engineering. Examples
40 include any cloned polynucleotide, or polynucleotides, that are linked or joined to heterologous sequences. The term "recombinant" does not refer to alterations of polynucleotides that result from naturally occurring events, such as spontaneous mutations, or from non-spontaneous mutagenesis followed by selective breeding.

Plants containing mutations arising due to non-spontaneous mutagenesis and selective breeding are referred to herein as non-transgenic plants and are included in the present invention. In embodiments wherein the plant is transgenic and comprises multiple mutated PPO nucleic acids, the nucleic acids can be derived from different genomes or from the same genome. Alternatively, in embodiments wherein the plant is non-transgenic and comprises multiple mutated PPO nucleic acids, the nucleic acids are located on different genomes or on the same genome. As used herein, "mutagenized" refers to an organism or DNA thereof having alteration(s) in the biomolecular sequence of its native genetic material as compared to the sequence of the genetic material of a corresponding wild-type organism or DNA, wherein the alteration(s) in genetic material were induce and/or selected by human action. Methods of inducing mutations can induce mutations in random positions in the genetic material or can induce mutations in specific locations in the genetic material (i.e., can be directed mutagenesis techniques), such as by use of a genoplasty technique.

In certain embodiments, the present invention involves herbicide-resistant plants that are produced by mutation breeding. Such plants comprise a polynucleotide encoding a mutated PPO and are tolerant to one or more PPO-inhibiting herbicides. Such methods can involve, for example, exposing the plants or seeds to a mutagen, particularly a chemical mutagen such as, for example, ethyl methanesulfonate (EMS) and selecting for plants that have enhanced tolerance to at least one or more PPO-inhibiting herbicide.

However, the present invention is not limited to herbicide-tolerant plants that are produced by a mutagenesis method involving the chemical mutagen EMS. Any mutagenesis method known in the art may be used to produce the herbicide-resistant plants of the present invention. Such mutagenesis methods can involve, for example, the use of any one or more of the following mutagens: radiation, such as X-rays, Gamma rays (e.g., cobalt 60 or cesium 137), neutrons, (e.g., product of nuclear fission by uranium 235 in an atomic reactor), Beta radiation (e.g., emitted from radioisotopes such as phosphorus 32 or carbon 14), and ultraviolet radiation (preferably from 2500 to 2900 nm), and chemical mutagens such as base analogues (e.g., 5-bromo-uracil), related compounds (e.g., 8-ethoxy caffeine), antibiotics (e.g., streptonigrin), alkylating agents (e.g., sulfur mustards, nitrogen mustards, epoxides, ethylenamines, sulfates, sulfonates, sulfones, lactones), azide, hydroxylamine, nitrous acid, or acridines. Herbicide-resistant plants can also be produced by using tissue culture methods to select for plant cells comprising herbicide-resistance mutations and then regenerating herbicide-resistant plants therefrom. See, for example, U.S. Patent Nos. 5,773,702 and 5,859,348, both of which are herein incorporated in their entirety by reference. Further details of mutation breeding can be found in "Principals of Cultivar Development" Fehr, 1993 Macmillan Publishing Company the disclosure of which is incorporated herein by reference

In addition to the definition above, the term "plant" is intended to encompass crop plants at any stage of maturity or development, as well as any tissues or organs (plant parts) taken or derived from any such plant unless otherwise clearly indicated by context. Plant parts include, but are not limited to, stems, roots, flowers, ovules, stamens, leaves, embryos, meristematic regions, callus tissue, anther cultures, gametophytes, sporophytes, pollen, microspores, protoplasts, and the like.

The plant of the present invention comprises at least one mutated PPO nucleic acid or over-

expressed wild-type PPO nucleic acid, and has increased tolerance to a PPO-inhibiting herbicide as compared to a wild-type variety of the plant. It is possible for the plants of the present invention to have multiple wild-type or mutated PPO nucleic acids from different genomes since these plants can contain more than one genome. For example, a plant contains two genomes, usually referred to as the A and B genomes. Because PPO is a required metabolic enzyme, it is assumed that each genome has at least one gene coding for the PPO enzyme (i.e. at least one PPO gene). As used herein, the term "PPO gene locus" refers to the position of an PPO gene on a genome, and the terms "PPO gene" and "PPO nucleic acid" refer to a nucleic acid encoding the PPO enzyme. The PPO nucleic acid on each genome differs in its nucleotide sequence from an PPO nucleic acid on another genome. One of skill in the art can determine the genome of origin of each PPO nucleic acid through genetic crossing and/or either sequencing methods or exonuclease digestion methods known to those of skill in the art.

The present invention includes plants comprising one, two, three, or more mutated PPO alleles, wherein the plant has increased tolerance to a PPO-inhibiting herbicide as compared to a wild-type variety of the plant. The mutated PPO alleles can comprise a nucleotide sequence selected from the group consisting of a polynucleotide as defined in SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47, or a variant or derivative thereof, a polynucleotide encoding a polypeptide as defined in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, or a variant or derivative, homologue, orthologue, paralogue thereof, a polynucleotide comprising at least 60 consecutive nucleotides of any of the aforementioned polynucleotides; and a polynucleotide complementary to any of the aforementioned polynucleotides.

"Alleles" or "allelic variants" are alternative forms of a given gene, located at the same chromosomal position. Allelic variants encompass Single Nucleotide Polymorphisms (SNPs), as well as Small Insertion/Deletion Polymorphisms (INDELs). The size of INDELs is usually less than 100 bp. SNPs and INDELs form the largest set of sequence variants in naturally occurring polymorphic strains of most organisms

The term "variety" refers to a group of plants within a species defined by the sharing of a common set of characteristics or traits accepted by those skilled in the art as sufficient to distinguish one cultivar or variety from another cultivar or variety. There is no implication in either term that all plants of any given cultivar or variety will be genetically identical at either the whole gene or molecular level or that any given plant will be homozygous at all loci. A cultivar or variety is considered "true breeding" for a particular trait if, when the true-breeding cultivar or variety is self-pollinated, all of the progeny contain the trait. The terms "breeding line" or "line" refer to a group of plants within a cultivar defined by the sharing of a common set of characteristics or traits accepted by those skilled in the art as sufficient to distinguish one breeding line or line from another breeding line or line. There is no implication in either term that all plants of any given breeding line or line will be genetically identical at either the whole gene or molecular level or that any given plant will be homozygous at all loci. A breeding line or line is considered "true breeding" for a particular trait if, when the true-breeding line or breeding line is self-pollinated, all of the progeny contain the trait. In the present invention, the trait arises from a mutation in a PPO gene of the plant or seed.

In some embodiments, traditional plant breeding is employed whereby the PPO-inhibiting herbicides-tolerant trait is introduced in the progeny plant resulting therefrom. In one embodiment, the present invention provides a method for producing a PPO-inhibiting herbicides-tolerant progeny plant, the method comprising: crossing a parent plant with a PPO-inhibiting herbicides-tolerant
5 plant to introduce the PPO-inhibiting herbicides-tolerance characteristics of the PPO-inhibiting herbicides-tolerant plant into the germplasm of the progeny plant, wherein the progeny plant has increased tolerance to the PPO-inhibiting herbicides relative to the parent plant. In other
10 embodiments, the method further comprises the step of introgressing the PPO-inhibiting herbicides-tolerance characteristics through traditional plant breeding techniques to obtain a descendent plant having the PPO-inhibiting herbicides-tolerance characteristics

The herbicide-resistant plants of the invention that comprise polynucleotides encoding mutated PPO polypeptides also find use in methods for increasing the herbicide-resistance of a plant through conventional plant breeding involving sexual reproduction. The methods comprise crossing
15 a first plant that is a herbicide-resistant plant of the invention to a second plant that may or may not be resistant to the same herbicide or herbicides as the first plant or may be resistant to different herbicide or herbicides than the first plant. The second plant can be any plant that is capable of producing viable progeny plants (i.e., seeds) when crossed with the first plant. Typically, but not necessarily, the first and second plants are of the same species. The methods can optionally
20 involve selecting for progeny plants that comprise the mutated PPO polypeptides of the first plant and the herbicide resistance characteristics of the second plant. The progeny plants produced by this method of the present invention have increased resistance to a herbicide when compared to either the first or second plant or both. When the first and second plants are resistant to different herbicides, the progeny plants will have the combined herbicide tolerance characteristics of the first
25 and second plants. The methods of the invention can further involve one or more generations of backcrossing the progeny plants of the first cross to a plant of the same line or genotype as either the first or second plant. Alternatively, the progeny of the first cross or any subsequent cross can be crossed to a third plant that is of a different line or genotype than either the first or second plant. The present invention also provides plants, plant organs, plant tissues, plant cells, seeds, and non-
30 human host cells that are transformed with the at least one polynucleotide molecule, expression cassette, or transformation vector of the invention. Such transformed plants, plant organs, plant tissues, plant cells, seeds, and non-human host cells have enhanced tolerance or resistance to at least one herbicide, at levels of the herbicide that kill or inhibit the growth of an untransformed plant, plant tissue, plant cell, or non-human host cell, respectively. Preferably, the transformed
35 plants, plant tissues, plant cells, and seeds of the invention are *Arabidopsis thaliana* and crop plants.

In other aspects, plants of the invention include those plants which, in addition to being tolerant to PPO-inhibiting herbicides, have been subjected to further genetic modifications by breeding,
40 mutagenesis or genetic engineering, e.g. have been rendered tolerant to applications of specific other classes of herbicides, such as AHAS inhibitors; auxinic herbicides; bleaching herbicides such as hydroxyphenylpyruvate dioxygenase (HPPD) inhibitors or phytoene desaturase (PDS) inhibitors; EPSPS inhibitors such as glyphosate; glutamine synthetase (GS) inhibitors such as glufosinate; lipid biosynthesis inhibitors such as acetyl CoA carboxylase (ACCase) inhibitors; or

oxynil (i.e. bromoxynil or ioxynil) herbicides as a result of conventional methods of breeding or genetic engineering. Thus, PPO-inhibiting herbicides-tolerant plants of the invention can be made resistant to multiple classes of herbicides through multiple genetic modifications, such as resistance to both glyphosate and glufosinate or to both glyphosate and a herbicide from another class such as HPPD inhibitors, AHAS inhibitors, or ACCase inhibitors. These herbicide resistance technologies are, for example, described in Pest Management Science (at volume, year, page): 61, 2005, 246; 61, 2005, 258; 61, 2005, 277; 61, 2005, 269; 61, 2005, 286; 64, 2008, 326; 64, 2008, 332; Weed Science 57, 2009, 108; Australian Journal of Agricultural Research 58, 2007, 708; Science 316, 2007, 1185; and references quoted therein. For example, PPO-inhibiting herbicides-tolerant plants of the invention, in some embodiments, may be tolerant to ACCase inhibitors, such as "dms" (e.g., cycloxydim, sethoxydim, clethodim, or tepraloxydim), "fops" (e.g., clodinafop, diclofop, fluazifop, haloxyfop, or quizalofop), and "dens" (such as pinoxaden); to auxinic herbicides, such as dicamba; to EPSPS inhibitors, such as glyphosate; to other PPO inhibitors; and to GS inhibitors, such as glufosinate.

In addition to these classes of inhibitors, PPO-inhibiting herbicides-tolerant plants of the invention may also be tolerant to herbicides having other modes of action, for example, chlorophyll/carotenoid pigment inhibitors, cell membrane disrupters, photosynthesis inhibitors, cell division inhibitors, root inhibitors, shoot inhibitors, and combinations thereof.

Such tolerance traits may be expressed, e.g. : as mutant or wildtype PPO proteins, as mutant AHASL proteins, mutant ACCase proteins, mutant EPSPS proteins, or mutant glutamine synthetase proteins; or as mutant native, inbred, or transgenic aryloxyalkanoate dioxygenase (AAD or DHT), haloarylnitrilase (BXN), 2,2-dichloropropionic acid dehalogenase (DEH), glyphosate-N-acetyltransferase (GAT), glyphosate decarboxylase (GDC), glyphosate oxidoreductase (GOX), glutathione-S-transferase (GST), phosphinothricin acetyltransferase (PAT or bar), or CYP450s proteins having an herbicide-degrading activity.

PPO-inhibiting herbicides-tolerant plants hereof can also be stacked with other traits including, but not limited to, pesticidal traits such as Bt Cry and other proteins having pesticidal activity toward coleopteran, lepidopteran, nematode, or other pests; nutrition or nutraceutical traits such as modified oil content or oil profile traits, high protein or high amino acid concentration traits, and other trait types known in the art.

Furthermore, in other embodiments, PPO-inhibiting herbicides-tolerant plants are also covered which are, by the use of recombinant DNA techniques and/or by breeding and/or otherwise selected for such characteristics, rendered able to synthesize one or more insecticidal proteins, especially those known from the bacterial genus *Bacillus*, particularly from *Bacillus thuringiensis*, such as [delta]-endotoxins, e.g. CryIA(b), CryIA(c), CryIF, CryIF(a2), CryIIA(b), CryIIIA, CryIIIB(bI) or Cry9c; vegetative insecticidal proteins (VIP), e.g. VIP1, VIP2, VIP3 or VIP3A; insecticidal proteins of bacteria colonizing nematodes, e.g. *Photorhabdus* spp. or *Xenorhabdus* spp.; toxins produced by animals, such as scorpion toxins, arachnid toxins, wasp toxins, or other insect-specific neurotoxins; toxins produced by fungi, such as streptomycete toxins; plant lectins, such as pea or barley lectins; agglutinins; proteinase inhibitors, such as trypsin inhibitors, serine protease

inhibitors, patatin, cystatin or papain inhibitors; ribosome-inactivating proteins (RIP), such as ricin, maize-RIP, abrin, luffin, saporin or bryodin; steroid metabolism enzymes, such as 3-hydroxy-steroid oxidase, ecdysteroid-IDP-glycosyl-transferase, cholesterol oxidases, ecdysone inhibitors or HMG- CoA-reductase; ion channel blockers, such as blockers of sodium or calcium channels; juvenile hormone esterase; diuretic hormone receptors (helicokinin receptors); stilben synthase, bibenzyl synthase, chitinases or glucanases. In the context of the present invention these insecticidal proteins or toxins are to be understood expressly also as pre-toxins, hybrid proteins, truncated or otherwise modified proteins. Hybrid proteins are characterized by a new combination of protein domains, (see, e.g. WO 02/015701). Further examples of such toxins or genetically modified plants capable of synthesizing such toxins are disclosed, e.g., in EP-A 374 753, WO 93/007278, WO 95/34656, EP-A 427 529, EP-A 451 878, WO 03/18810 und WO 03/52073. The methods for producing such genetically modified plants are generally known to the person skilled in the art and are described, e.g. in the publications mentioned above. These insecticidal proteins contained in the genetically modified plants impart to the plants producing these proteins tolerance to harmful pests from all taxonomic groups of arthropods, especially to beetles (Coleoptera), two-winged insects (Diptera), and moths (Lepidoptera) and to nematodes (Nematoda).

In some embodiments, expression of one or more protein toxins (e.g., insecticidal proteins) in the PPO-inhibiting herbicides-tolerant plants is effective for controlling organisms that include, for example, members of the classes and orders: Coleoptera such as the American bean weevil *Acanthoscelides obtectus*; the leaf beetle *Agelastica alni*; click beetles (*Agriotes lineatus*, *Agriotes obscurus*, *Agriotes bicolor*); the grain beetle *Ahasverus advena*; the summer schafer *Amphimallon solstitialis*; the furniture beetle *Anobium punctatum*; *Anthonomus* spp. (weevils); the Pygmy mangold beetle *Atomaria linearis*; carpet beetles (*Anthrenus* spp., *Attagenus* spp.); the cowpea weevil *Callosobruchus maculatus*; the fried fruit beetle *Carpophilus hemipterus*; the cabbage seedpod weevil *Ceutorhynchus assimilis*; the rape winter stem weevil *Ceutorhynchus pictaris*; the wireworms *Conoderus vespertinus* and *Conoderus falli*; the banana weevil *Cosmopolites sordidus*; the New Zealand grass grub *Costelytra zealandica*; the June beetle *Cotinis nitida*; the sunflower stem weevil *Cylindrocopturus adspersus*; the larder beetle *Dermestes lardarius*; the corn rootworms *Diabrotica virgifera*, *Diabrotica virgifera virgifera*, and *Diabrotica barberi*; the Mexican bean beetle *Epilachna varivestis*; the old house borer *Hylotropes bajulus*; the lucerne weevil *Hypera postica*; the shiny spider beetle *Gibbium psyllodes*; the cigarette beetle *Lasioderma serricornis*; the Colorado potato beetle *Leptinotarsa decemlineata*; *Lyctus* beetles {*Lyctus* spp. , the pollen beetle *Meligethes aeneus*; the common cockshafer *Melolontha melolontha*; the American spider beetle *Mezium americanum*; the golden spider beetle *Niptus hololeucus*; the grain beetles *Oryzaephilus surinamensis* and *Oryzaephilus Mercator*; the black vine weevil *Otiorynchus sulcatus*; the mustard beetle *Phaedon cochleariae*, the crucifer flea beetle *Phyllotreta cruciferae*; the striped flea beetle *Phyllotreta striolata*; the cabbage steam flea beetle *Psylliodes chrysocephala*; *Ptinus* spp. (spider beetles); the lesser grain borer *Rhizopertha dominica*; the pea and bean weevil *Sitona lineatus*; the rice and granary beetles *Sitophilus oryzae* and *Sitophilus granaries*; the red sunflower seed weevil *Smicronyx fulvus*; the drugstore beetle *Stegobium paniceum*; the yellow mealworm beetle *Tenebrio molitor*, the flour beetles *Tribolium castaneum* and *Tribolium confusum*; warehouse and cabinet beetles {*Trogoderma* spp.}; the sunflower beetle *Zygogramma exclamationis*; Dermaptera (earwigs) such as the European earwig *Forficula*

auricularia and the striped earwig *Labidura riparia*; Dictyoptera such as the oriental cockroach *Blatta orientalis*; the greenhouse millipede *Oxidus gracilis*; the beet fly *Pegomyia betae*; the frit fly *Oscinella frit*; fruitflies (*Dacus* spp., *Drosophila* spp.); Isoptera (termites) including species from the families Hodotermitidae, Kalotermitidae, Mastotermitidae, Rhinotermitidae, Serritermitidae, 5 Termitidae, Termopsidae; the tarnished plant bug *Lygus lineolaris*; the black bean aphid *Aphis fabae*; the cotton or melon aphid *Aphis gossypii*; the green apple aphid *Aphis pomi*; the citrus spiny whitefly *Aleurocanthus spiniferus*; the sweet potato whitefly *Bemisia tabaci*; the cabbage aphid *Brevicoryne brassicae*; the pear psylla *Cacopsylla pyricola*; the currant aphid *Cryptomyzus ribis*; the grape phylloxera *Daktulosphaira vitifoliae*; the citrus psylla *Diaphorina citri*; the potato 10 leafhopper *Empoasca fabae*; the bean leafhopper *Empoasca Solana*; the vine leafhopper *Empoasca vitis*; the woolly aphid *Eriosoma lanigerum*; the European fruit scale *Eulecanium corni*; the mealy plum aphid *Hyalopterus arundinis*; the small brown planthopper *Laodelphax striatellus*; the potato aphid *Macrosiphum euphorbiae*; the green peach aphid *Myzus persicae*; the green rice leafhopper *Nephotettix cincticeps*; the brown planthopper *Nilaparvata lugens*; the hop aphid 15 *Phorodon humuli*; the bird-cherry aphid *Rhopalosiphum padi*; the grain aphid *Sitobion avenae*; Lepidoptera such as *Adoxophyes orana* (summer fruit tortrix moth); *Archips podana* (fruit tree tortrix moth); *Bucculatrix pyrivorella* (pear leafminer); *Bucculatrix thurberiella* (cotton leaf perforator); *Bupalus piniarius* (pine looper); *Carpocapsa pomonella* (codling moth); *Chilo suppressalis* (striped rice borer); *Choristoneura fumiferana* (eastern spruce budworm); *Cochylis* 20 *hospes* (banded sunflower moth); *Diatraea grandiosella* (southwestern corn borer); *Eupoecilia ambiguella* (European grape berry moth); *Helicoverpa armigera* (cotton bollworm); *Helicoverpa zea* (cotton bollworm); *Heliothis virescens* (tobacco budworm); *Homeosoma electellum* (sunflower moth); *Homona magnanima* (oriental tea tree tortrix moth); *Lithocolletis blattellata* (spotted tentiform leafminer); *Lymantria dispar* (gypsy moth); *Malacosoma neustria* (tent caterpillar); 25 *Mamestra brassicae* (cabbage armyworm); *Mamestra configurata* (Bertha armyworm); *Operophtera brumata* (winter moth); *Ostrinia nubilalis* (European corn borer); *Panolis flammea* (pine beauty moth); *Phyllocnistis citrella* (citrus leafminer); *Pieris brassicae* (cabbage white butterfly); *Rachiplusia ni* (soybean looper); *Spodoptera exigua* (beet armyworm); *Spodoptera littoralis* (cotton leafworm); *Sylepta derogata* (cotton leaf roller); *Trichoplusia ni* (cabbage looper); 30 Orthoptera such as the common cricket *Acheta domesticus*, tree locusts (*Anacridium* spp.), the migratory locust *Locusta migratoria*, the two-striped grasshopper *Melanoplus bivittatus*, the differential grasshopper *Melanoplus differentialis*, the redlegged grasshopper *Melanoplus femurrubrum*, the migratory grasshopper *Melanoplus sanguinipes*, the northern mole cricket *Neocurtilla hexadactyla*, the red locust *Nomadacris septemfasciata*, the shortwinged mole cricket 35 *Scapteriscus abbreviatus*, the southern mole cricket *Scapteriscus borellii*, the tawny mole cricket *Scapteriscus vicinus*, and the desert locust *Schistocerca gregaria*; Symphyla such as the garden symphytan *Scutigerella immaculata*; Thysanoptera such as the tobacco thrips *Frankliniella fusca*, the flower thrips *Frankliniella intonsa*, the western flower thrips *Frankliniella occidentalis*, the cotton bud thrips *Frankliniella schultzei*, the banded greenhouse thrips *Hercinothrips femoralis*, the 40 soybean thrips *Neohydatothrips variabilis*, Kelly's citrus thrips *Pezothrips kellyanus*, the avocado thrips *Scirtothrips perseae*, the melon thrips *Thrips palmi*, and the onion thrips *Thrips tabaci*; and the like, and combinations comprising one or more of the foregoing organisms.

In some embodiments, expression of one or more protein toxins (e.g., insecticidal proteins) in the

PPO-inhibiting herbicides-tolerant plants is effective for controlling flea beetles, i.e. members of the flea beetle tribe of family Chrysomelidae, preferably against *Phyllotreta* spp., such as *Phyllotreta cruciferae* and/or *Phyllotreta triolata*. In other embodiments, expression of one or more protein toxins {e.g., insecticidal proteins} in the PPO-inhibiting herbicides- tolerant plants is effective for controlling cabbage seedpod weevil, the Bertha armyworm, *Lygus* bugs, or the diamondback moth. Furthermore, in one embodiment, PPO-inhibiting herbicides-tolerant plants are also covered which are, e.g. by the use of recombinant DNA techniques and/or by breeding and/or otherwise selected for such traits, rendered able to synthesize one or more proteins to increase the resistance or tolerance of those plants to bacterial, viral or fungal pathogens. The methods for producing such genetically modified plants are generally known to the person skilled in the art.

Furthermore, in another embodiment, PPO-inhibiting herbicides-tolerant plants are also covered which are, e.g. by the use of recombinant DNA techniques and/or by breeding and/or otherwise selected for such traits, rendered able to synthesize one or more proteins to increase the productivity (e.g. oil content), tolerance to drought, salinity or other growth-limiting environmental factors or tolerance to pests and fungal, bacterial or viral pathogens of those plants.

Furthermore, in other embodiments, PPO-inhibiting herbicides-tolerant plants are also covered which are, e.g. by the use of recombinant DNA techniques and/or by breeding and/or otherwise selected for such traits, altered to contain a modified amount of one or more substances or new substances, for example, to improve human or animal nutrition, e.g. oil crops that produce health-promoting long-chain omega-3 fatty acids or unsaturated omega-9 fatty acids (e.g. Nexera(R) rape, Dow Agro Sciences, Canada).

Furthermore, in some embodiments, PPO-inhibiting herbicides-tolerant plants are also covered which are, e.g. by the use of recombinant DNA techniques and/or by breeding and/or otherwise selected for such traits, altered to contain increased amounts of vitamins and/or minerals, and/or improved profiles of nutraceutical compounds.

In one embodiment, PPO-inhibiting herbicides-tolerant plants of the present invention, relative to a wild-type plant, comprise an increased amount of, or an improved profile of, a compound selected from the group consisting of: glucosinolates (e.g., glucoraphanin (4-methylsulfinylbutyl-glucosinolate), sulforaphane, 3-indolylmethyl-glucosinolate (glucobrassicin), 1-methoxy-3-indolylmethyl-glucosinolate (neoglucobrassicin)); phenolics (e.g., flavonoids (e.g., quercetin, kaempferol), hydroxycinnamoyl derivatives (e.g., 1,2,2'-trisinapoylgentiobiose, 1,2-diferuloylgentiobiose, 1,2'-disinapoyl-2-feruloylgentiobiose, 3-O-caffeoyl-quinic (neochlorogenic acid)); and vitamins and minerals (e.g., vitamin C, vitamin E, carotene, folic acid, niacin, riboflavin, thiamine, calcium, iron, magnesium, potassium, selenium, and zinc).

In another embodiment, PPO-inhibiting herbicides-tolerant plants of the present invention, relative to a wild-type plant, comprise an increased amount of, or an improved profile of, a

- compound selected from the group consisting of: progoitrin; isothiocyanates; indoles (products of glucosinolate hydrolysis); glutathione; carotenoids such as beta-carotene, lycopene, and the xanthophyll carotenoids such as lutein and zeaxanthin; phenolics comprising the flavonoids such as the flavonols (e.g. quercetin, rutin), the flavans/tannins (such as the procyanidins
- 5 comprising coumarin, proanthocyanidins, catechins, and anthocyanins); flavones; phytoestrogens such as coumestans, lignans, resveratrol, isoflavones e.g. genistein, daidzein, and glycitein; resorcylic acid lactones; organosulphur compounds; phytosterols; terpenoids such as carnosol, rosmarinic acid, glycyrrhizin and saponins; chlorophyll; chlorophyllin, sugars, anthocyanins, and vanilla. In other embodiments, PPO-inhibiting herbicides-tolerant plants of
- 10 the present invention, relative to a wild-type plant, comprise an increased amount of, or an improved profile of, a compound selected from the group consisting of: vincristine, vinblastine, taxanes (e.g., taxol (paclitaxel), baccatin III, 10-desacetylbaccatin III, 10-desacetyl taxol, xylosyl taxol, 7- epitaxol, 7-epibaccatin III, 10-desacetylcephalomannine, 7- epicephalomannine, taxotere, cephalomannine, xylosyl cephalomannine, taxagifine, 8-
- 15 benxoyloxy taxagifine, 9-acetyloxy taxusin, 9-hydroxy taxusin, taiwanxam, taxane Ia, taxane Ib, taxane Ic, taxane Id, GMP paclitaxel, 9-dihydro 13-acetylbaccatin III, 10-desacetyl-7-epitaxol, tetrahydrocannabinol (THC), cannabidiol (CBD), genistein, diadzein, codeine, morphine, quinine, shikonin, ajmalacine, serpentine, and the like.
- 20 It is to be understood that the plant of the present invention can comprise a wild type PPO nucleic acid in addition to a mutated PPO nucleic acid. It is contemplated that the PPO-inhibiting herbicide tolerant lines may contain a mutation in only one of multiple PPO isoenzymes. Therefore, the present invention includes a plant comprising one or more mutated PPO nucleic acids in addition to one or more wild type PPO nucleic acids.
- 25 In another embodiment, the invention refers to a seed produced by a transgenic plant comprising a plant cell of the present invention, wherein the seed is true breeding for an increased resistance to a PPO-inhibiting herbicide as compared to a wild type variety of the seed.
- 30 In another embodiment, the invention refers to a method of producing a transgenic plant cell with an increased resistance to a PPO-inhibiting herbicide as compared to a wild type variety of the plant cell comprising, transforming the plant cell with an expression cassette comprising a mutated PPO nucleic acid.
- 35 In another embodiment, the invention refers to a method of producing a transgenic plant comprising, (a) transforming a plant cell with an expression cassette comprising a mutated PPO nucleic acid, and (b) generating a plant with an increased resistance to PPO-inhibiting herbicide from the plant cell.
- 40 Consequently, mutated PPO nucleic acids of the invention are provided in expression cassettes for expression in the plant of interest. The cassette will include regulatory sequences operably linked to a mutated PPO nucleic acid sequence of the invention. The term "regulatory element" as used herein refers to a polynucleotide that is capable of regulating the transcription of an operably linked polynucleotide. It includes, but not limited to, promoters, enhancers, introns, 5' UTRs, and 3' UTRs.

By "operably linked" is intended a functional linkage between a promoter and a second sequence, wherein the promoter sequence initiates and mediates transcription of the DNA sequence corresponding to the second sequence. Generally, operably linked means that the nucleic acid sequences being linked are contiguous and, where necessary to join two protein coding regions, contiguous and in the same reading frame. The cassette may additionally contain at least one additional gene to be cotransformed into the organism. Alternatively, the additional gene(s) can be provided on multiple expression cassettes.

Such an expression cassette is provided with a plurality of restriction sites for insertion of the mutated PPO nucleic acid sequence to be under the transcriptional regulation of the regulatory regions. The expression cassette may additionally contain selectable marker genes.

The expression cassette of the present invention will include in the 5'-3' direction of transcription, a transcriptional and translational initiation region (i.e., a promoter), a mutated PPO encoding nucleic acid sequence of the invention, and a transcriptional and translational termination region (i.e., termination region) functional in plants. The promoter may be native or analogous, or foreign or heterologous, to the plant host and/or to the mutated PPO nucleic acid sequence of the invention. Additionally, the promoter may be the natural sequence or alternatively a synthetic sequence. Where the promoter is "foreign" or "heterologous" to the plant host, it is intended that the promoter is not found in the native plant into which the promoter is introduced. Where the promoter is "foreign" or "heterologous" to the mutated PPO nucleic acid sequence of the invention, it is intended that the promoter is not the native or naturally occurring promoter for the operably linked mutated PPO nucleic acid sequence of the invention. As used herein, a chimeric gene comprises a coding sequence operably linked to a transcription initiation region that is heterologous to the coding sequence.

While it may be preferable to express the mutated PPO nucleic acids of the invention using heterologous promoters, the native promoter sequences may be used. Such constructs would change expression levels of the mutated PPO protein in the plant or plant cell. Thus, the phenotype of the plant or plant cell is altered.

The termination region may be native with the transcriptional initiation region, may be native with the operably linked mutated PPO sequence of interest, may be native with the plant host, or may be derived from another source (i.e., foreign or heterologous to the promoter, the mutated PPO nucleic acid sequence of interest, the plant host, or any combination thereof). Convenient termination regions are available from the Ti-plasmid of *A. tumefaciens*, such as the octopine synthase and nopaline synthase termination regions. See also Guerineau et al. (1991) Mol. Gen. Genet. 262: 141-144; Proudfoot (1991) Cell 64:671-674; Sanfacon et al. (1991) Genes Dev. 5: 141-149; Mogen et al. (1990) Plant Cell 2: 1261-1272; Munroe et al. (1990) Gene 91: 151-158; Ballas et al. (1989) Nucleic Acids Res. 17:7891-7903; and Joshi et al. (1987) Nucleic Acid Res. 15:9627-9639. Where appropriate, the gene(s) may be optimized for increased expression in the transformed plant. That is, the genes can be synthesized using plant-preferred codons for improved expression. See, for example, Campbell and Gowri (1990) Plant Physiol. 92: 1-11 for a discussion of host-preferred codon usage. Methods are available in the art for synthesizing plant-

preferred genes. See, for example, U.S. Patent Nos. 5,380,831, and 5,436,391, and Murray et al. (1989) *Nucleic Acids Res.* 17:477-498, herein incorporated by reference.

Additional sequence modifications are known to enhance gene expression in a cellular host. These include elimination of sequences encoding spurious polyadenylation signals, exon-intron splice site signals, transposon-like repeats, and other such well-characterized sequences that may be deleterious to gene expression. The G-C content of the sequence may be adjusted to levels average for a given cellular host, as calculated by reference to known genes expressed in the host cell. When possible, the sequence is modified to avoid predicted hairpin secondary mRNA structures. Nucleotide sequences for enhancing gene expression can also be used in the plant expression vectors. These include the introns of the maize *Adhl*, *intron1* gene (Callis et al. *Genes and Development* 1: 1183-1200, 1987), and leader sequences, (W- sequence) from the Tobacco Mosaic virus (TMV), Maize Chlorotic Mottle Virus and Alfalfa Mosaic Virus (Gallie et al. *Nucleic Acid Res.* 15:8693-8711, 1987 and Skuzeski et al. *Plant Mol. Biol.* 15:65-79, 1990). The first intron from the *shrunken-1* locus of maize, has been shown to increase expression of genes in chimeric gene constructs. U.S. Pat. Nos. 5,424,412 and 5,593,874 disclose the use of specific introns in gene expression constructs, and Gallie et al. (*Plant Physiol.* 106:929-939, 1994) also have shown that introns are useful for regulating gene expression on a tissue specific basis. To further enhance or to optimize mutated PPO gene expression, the plant expression vectors of the invention may also contain DNA sequences containing matrix attachment regions (MARs). Plant cells transformed with such modified expression systems, then, may exhibit overexpression or constitutive expression of a nucleotide sequence of the invention.

The expression cassettes of the present invention may additionally contain 5' leader sequences in the expression cassette construct. Such leader sequences can act to enhance translation. Translation leaders are known in the art and include: picornavirus leaders, for example, EMCV leader (Encephalomyocarditis 5' noncoding region) (Elroy-Stein et al. (1989) *Proc. Natl. Acad. Sci. USA* 86:6126-6130); potyvirus leaders, for example, TEV leader (Tobacco Etch Virus) (Gallie et al. (1995) *Gene* 165(2):233-238), MDMV leader (Maize Dwarf Mosaic Virus) (*Virology* 154:9-20), and human immunoglobulin heavy-chain binding protein (BiP) (Macejak et al. (1991) *Nature* 353:90-94); untranslated leader from the coat protein mRNA of alfalfa mosaic virus (AMV RNA 4) (Jobling et al. (1987) *Nature* 325:622-625); tobacco mosaic virus leader (TMV) (Gallie et al. (1989) in *Molecular Biology of RNA*, ed. Cech (Liss, New York), pp. 237-256); and maize chlorotic mottle virus leader (MCMV) (Lommel et al. (1991) *Virology* 81:382-385). See also, Della-Cioppa et al. (1987) *Plant Physiol.* 84:965-968. Other methods known to enhance translation can also be utilized, for example, introns, and the like.

In preparing the expression cassette, the various DNA fragments may be manipulated, so as to provide for the DNA sequences in the proper orientation and, as appropriate, in the proper reading frame. Toward this end, adapters or linkers may be employed to join the DNA fragments or other manipulations may be involved to provide for convenient restriction sites, removal of superfluous DNA, removal of restriction sites, or the like. For this purpose, *in vitro* mutagenesis, primer repair, restriction, annealing, resubstitutions, e.g., transitions and trans versions, may be involved.

A number of promoters can be used in the practice of the invention. The promoters can be selected based on the desired outcome. The nucleic acids can be combined with constitutive, tissue -preferred, or other promoters for expression in plants. Such constitutive promoters include, for example, the core promoter of the Rsyn7 promoter and other constitutive promoters disclosed in WO 99/43838 and U.S. Patent No. 6,072,050; the core CaMV 35S promoter (Odell et al. (1985) Nature 313:810-812); rice actin (McElroy et al. (1990) Plant Cell 2: 163-171); ubiquitin (Christensen et al. (1989) Plant Mol. Biol. 12:619-632 and Christensen et al. (1992) Plant Mol. Biol. 18:675-689); pEMU (Last et al. (1991) Theor. Appl. Genet. 81:581- 588); MAS (Velten et al. (1984) EMBO J. 3:2723-2730); ALS promoter (U.S. Patent No. 5,659,026), and the like. Other constitutive promoters include, for example, U.S. Patent Nos. 5,608,149; 5,608,144; 5,604,121; 5,569,597; 5,466,785; 5,399,680; 5,268,463; 5,608,142; and 6,177,611.

Tissue-preferred promoters can be utilized to target enhanced mutated PPO expression within a particular plant tissue. Such tissue-preferred promoters include, but are not limited to, leaf -preferred promoters, root-preferred promoters, seed- preferred promoters, and stem-preferred promoters. Tissue-preferred promoters include Yamamoto et al. (1997) Plant J. 12(2):255-265; Kawamata et al. (1997) Plant Cell Physiol. 38(7):792-803; Hansen et al. (1997) Mol. Gen Genet. 254(3):337-343; Russell et al. (1997) Transgenic Res. 6(2): 157-168; Rinehart et al. (1996) Plant Physiol. 112(3): 1331-1341; Van Camp et al. (1996) Plant Physiol. 112(2):525-535; Canevascini et al. (1996) Plant Physiol. 112(2):513-524; Yamamoto et al. (1994) Plant Cell Physiol. 35(5):773-778; Lam (1994) Results Probl. Cell Differ. 20: 181- 196; Orozco et al. (1993) Plant Mol Biol. 23(6): 1129-1138; Matsuoka et al. (1993) Proc Natl. Acad. Sci. USA 90(20):9586-9590; and Guevara-Garcia et al. (1993) Plant J. 4(3):495-505. Such promoters can be modified, if necessary, for weak expression. In one embodiment, the nucleic acids of interest are targeted to the chloroplast for expression.

In this manner, where the nucleic acid of interest is not directly inserted into the chloroplast, the expression cassette will additionally contain a chloroplast-targeting sequence comprising a nucleotide sequence that encodes a chloroplast transit peptide to direct the gene product of interest to the chloroplasts. Such transit peptides are known in the art. With respect to chloroplast-targeting sequences, "operably linked" means that the nucleic acid sequence encoding a transit peptide (i.e., the chloroplast-targeting sequence) is linked to the mutated PPO nucleic acid of the invention such that the two sequences are contiguous and in the same reading frame. See, for example, Von Heijne et al. (1991) Plant Mol. Biol. Rep. 9: 104-126; Clark et al. (1989) J. Biol. Chem. 264:17544-17550; Della-Cioppa et al. (1987) Plant Physiol. 84:965-968; Romer et al. (1993) Biochem. Biophys. Res. Commun. 196:1414-1421; and Shah et al. (1986) Science 233:478-481. While the mutated PPO proteins of the invention include a native chloroplast transit peptide, any chloroplast transit peptide known in the art can be fused to the amino acid sequence of a mature mutated PPO protein of the invention by operably linking a chloroplast-targeting sequence to the 5'-end of a nucleotide sequence encoding a mature mutated PPO protein of the invention. Chloroplast targeting sequences are known in the art and include the chloroplast small subunit of ribulose-1,5-bisphosphate carboxylase (Rubisco) (de Castro Silva Filho et al. (1996) Plant Mol. Biol. 30:769-780; Schnell et al. (1991) J. Biol. Chem. 266(5):3335-3342); 5 - (enolpyruvyl)shikimate-3 -phosphate synthase (EPSPS) (Archer et al. (1990) J. Bioenerg.

Biomemb. 22(6):789-810); tryptophan synthase (Zhao et al. (1995) J. Biol. Chem. 270(11):6081-6087); plastocyanin(Lawrence et al. (1997) J. Biol. Chem. 272(33):20357-20363); chorismate synthase (Schmidt et al. (1993) J. Biol. Chem. 268(36):27447-27457); and the light harvesting chlorophyll a/b binding protein (LHBP) (Lamppa et al. (1988) J. Biol. Chem. 263: 14996-14999).

5 See also Von Heijne et al. (1991) Plant Mol. Biol. Rep. 9: 104- 126; Clark et al. (1989) J. Biol. Chem. 264:17544-17550; Della-Cioppa et al. (1987) Plant Physiol. 84:965-968; Romer et al. (1993) Biochem. Biophys. Res. Commun. 196: 1414-1421; and Shah et al. (1986) Science 233:478-481.

10 In a preferred embodiment, the targeting sequence comprises a nucleotide sequence that encodes a transit peptide comprising the amino acid sequence of SEQ ID NO: 49, 50, 51, 52, or 53 (Ferredoxin transit peptide Fdxt). Preferably, the transit peptide encoding nucleic acid is operably linked such that the transit peptide is fused to the valine at position 46 in SEQ ID NO: 2 or 4.

15 In another preferred embodiment, the transit peptide encoding nucleic acid is operably linked such that the transit peptide is fused to the aspartic acid at position 71 in SEQ ID NO: 48.

In a particularly preferred embodiment, the nucleic acid sequence encoding a transit peptide comprises the sequence of SEQ ID NO: 54 (for expression in corn codon-optimized nucleic acid
20 encoding the Ferredoxin transit peptide of *Silene pratensis*) or SEQ ID NO: 55 (for expression in soy codon-optimized nucleic acid encoding the Ferredoxin transit peptide of *Silene pratensis*).

Methods for transformation of chloroplasts are known in the art. See, for example, Svab et al. (1990) Proc. Natl. Acad. Sci. USA 87:8526-8530; Svab and Maliga (1993) Proc. Natl. Acad. Sci.
25 USA 90:913-917; Svab and Maliga (1993) EMBO J. 12:601-606. The method relies on particle gun delivery of DNA containing a selectable marker and targeting of the DNA to the plastid genome through homologous recombination. Additionally, plastid transformation can be accomplished by transactivation of a silent plastid-borne transgene by tissue-preferred expression of a nuclear-encoded and plastid-directed RNA polymerase. Such a system has been reported in McBride et al.
30 (1994) Proc. Natl. Acad. Sci. USA 91:7301-7305. The nucleic acids of interest to be targeted to the chloroplast may be optimized for expression in the chloroplast to account for differences in codon usage between the plant nucleus and this organelle. In this manner, the nucleic acids of interest may be synthesized using chloroplast-preferred codons. See, for example, U.S. Patent No. 5,380,831, herein incorporated by reference.

35 In a preferred embodiment, the mutated PPO nucleic acid comprises a polynucleotide sequence selected from the group consisting of: a) a polynucleotide as shown in SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, or 47, or a variant or derivative thereof; b) a polynucleotide encoding a polypeptide as shown in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14,
40 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, or a variant or derivative thereof; c) a polynucleotide comprising at least 60 consecutive nucleotides of any of a) or c); and d) a polynucleotide complementary to the polynucleotide of any of a) through c)

Preferably, the expression cassette of the present invention further comprises a transcription

initiation regulatory region and a translation initiation regulatory region that are functional in the plant.

While the polynucleotides of the invention find use as selectable marker genes for plant transformation, the expression cassettes of the invention can include another selectable marker gene for the selection of transformed cells. Selectable marker genes, including those of the present invention, are utilized for the selection of transformed cells or tissues. Marker genes include, but are not limited to, genes encoding antibiotic resistance, such as those encoding neomycin phosphotransferase II (NEO) and hygromycin phosphotransferase (HPT), as well as genes conferring resistance to herbicidal compounds, such as glufosinate ammonium, bromoxynil, imidazolinones, and 2,4-dichlorophenoxyacetate (2,4-D). See generally, Yarranton (1992) *Curr. Opin. Biotech.* 3 :506-511 ; Christophers on et al (1992) *Proc. Natl. Acad. Sci USA* 89:6314-6318; Yao et al. (1992) *Cell* 71:63-72; Reznikoff (1992) *Mol Microbiol* 6:2419-2422; Barkley et al (1980) in *The Operon*, pp. 177-220; Hu et al (1987) *Cell* 48:555-566; Brown et al (1987) *Cell* 49:603-612; Figge et al (1988) *Cell* 52:713-722; Deuschle et al (1989) *Proc. Natl Acad. Sci USA* 86:5400-5404; Fuerst et al (1989) *Proc. Natl Acad. Sci USA* 86:2549-2553; Deuschle et al (1990) *Science* 248:480-483; Gossen (1993) Ph.D. Thesis, University of Heidelberg; Reines et al (1993) *Proc. Natl Acad. Sci USA* 90: 1917-1921; Labow et al (1990) *Mol Cell Biol* 10:3343-3356; Zambretti et al (1992) *Proc. Natl Acad. Sci USA* 89:3952-3956; Bairn et al (1991) *Proc. Natl Acad. Sci USA* 88:5072-5076; Wyborski et al (1991) *Nucleic Acids Res.* 19:4647-4653; Hillenand-Wissman (1989) *Topics Mol Struc. Biol* 10: 143- 162; Degenkolb et al (1991) *Antimicrob. Agents Chemother.* 35: 1591-1595; Kleinschmidt et al (1988) *Biochemistry* 27: 1094-1104; Bonin (1993) Ph.D. Thesis, University of Heidelberg; Gossen et al (1992) *Proc. Natl Acad. Sci USA* 89:5547- 5551; Oliva et al (1992) *Antimicrob. Agents Chemother.* 36:913-919; Hlavka et al (1985) *Handbook of Experimental Pharmacology*, Vol. 78 (Springer-Verlag, Berlin); Gill et al (1988) *Nature* 334:721-724. Such disclosures are herein incorporated by reference. The above list of selectable marker genes is not meant to be limiting. Any selectable marker gene can be used in the present invention.

The invention further provides an isolated recombinant expression vector comprising the expression cassette containing a mutated PPO nucleic acid as described above, wherein expression of the vector in a host cell results in increased tolerance to a PPO-inhibiting herbicide as compared to a wild type variety of the host cell. As used herein, the term "vector" refers to a nucleic acid molecule capable of transporting another nucleic acid to which it has been linked. One type of vector is a "plasmid," which refers to a circular double stranded DNA loop into which additional DNA segments can be ligated. Another type of vector is a viral vector, wherein additional DNA segments can be ligated into the viral genome. Certain vectors are capable of autonomous replication in a host cell into which they are introduced (e.g., bacterial vectors having a bacterial origin of replication and episomal mammalian vectors). Other vectors (e.g., non-episomal mammalian vectors) are integrated into the genome of a host cell upon introduction into the host cell, and thereby are replicated along with the host genome. Moreover, certain vectors are capable of directing the expression of genes to which they are operatively linked. Such vectors are referred to herein as "expression vectors." In general, expression vectors of utility in recombinant DNA techniques are often in the form of plasmids. In the present specification, "plasmid" and "vector" can be used interchangeably as the plasmid is the most commonly used form of vector. However,

the invention is intended to include such other forms of expression vectors, such as viral vectors (e.g., replication defective retroviruses, adenoviruses, and adeno-associated viruses), which serve equivalent functions.

- 5 The recombinant expression vectors of the invention comprise a nucleic acid of the invention in a form suitable for expression of the nucleic acid in a host cell, which means that the recombinant expression vectors include one or more regulatory sequences, selected on the basis of the host cells to be used for expression, which is operably linked to the nucleic acid sequence to be expressed. Regulatory sequences include those that direct constitutive expression of a nucleotide
10 sequence in many types of host cells and those that direct expression of the nucleotide sequence only in certain host cells or under certain conditions. It will be appreciated by those skilled in the art that the design of the expression vector can depend on such factors as the choice of the host cell to be transformed, the level of expression of polypeptide desired, etc. The expression vectors of the invention can be introduced into host cells to thereby produce polypeptides or peptides,
15 including fusion polypeptides or peptides, encoded by nucleic acids as described herein (e.g., mutated PPO polypeptides, fusion polypeptides, etc.).

In a preferred embodiment of the present invention, the mutated PPO polypeptides are expressed in plants and plants cells such as unicellular plant cells (such as algae) (See Falciatore et al., 1999,
20 Marine Biotechnology 1(3):239-251 and references therein) and plant cells from higher plants (e.g., the spermatophytes, such as crop plants). A mutated PPO polynucleotide may be "introduced" into a plant cell by any means, including transfection, transformation or transduction, electroporation, particle bombardment, agroinfection, biolistics, and the like.

25 Suitable methods for transforming or transfecting host cells including plant cells can be found in Sambrook et al. (Molecular Cloning: A Laboratory Manual. 2nd, ed., Cold Spring Harbor Laboratory, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989) and other laboratory manuals such as Methods in Molecular Biology, 1995, Vol. 44, Agrobacterium protocols, ed: Gartland and Davey, Humana Press, Totowa, New Jersey. As increased tolerance to PPO-
30 inhibiting herbicides is a general trait wished to be inherited into a wide variety of plants like maize, wheat, rye, oat, triticale, rice, barley, soybean, peanut, cotton, rapeseed and canola, manihot, pepper, sunflower and tagetes, solanaceous plants like potato, tobacco, eggplant, and tomato, Vicia species, pea, alfalfa, bushy plants (coffee, cacao, tea), Salix species, trees (oil palm, coconut), perennial grasses, and forage crops, these crop plants are also preferred target plants
35 for a genetic engineering as one further embodiment of the present invention. In a preferred embodiment, the plant is a crop plant. Forage crops include, but are not limited to, Wheatgrass, Canarygrass, Bromegrass, Wildrye Grass, Bluegrass, Orchardgrass, Alfalfa, Salfoin, Birdsfoot Trefoil, Alsike Clover, Red Clover, and Sweet Clover.

40 In one embodiment of the present invention, transfection of a mutated PPO polynucleotide into a plant is achieved by Agrobacterium mediated gene transfer. One transformation method known to those of skill in the art is the dipping of a flowering plant into an Agrobacteria solution, wherein the Agrobacteria contains the mutated PPO nucleic acid, followed by breeding of the transformed gametes. Agrobacterium mediated plant transformation can be performed using for example the

GV3101(pMP90) (Koncz and Schell, 1986, Mol. Gen. Genet. 204:383-396) or LBA4404 (Clontech) Agrobacterium tumefaciens strain. Transformation can be performed by standard transformation and regeneration techniques (Deblaere et al., 1994, Nucl. Acids. Res. 13:4777-4788; Gelvin, Stanton B. and Schilperoort, Robert A, Plant Molecular Biology Manual, 2nd Ed. - Dordrecht : Kluwer Academic Publ., 1995. - in Sect., Ringbuc Zentrale Signatur: BT11-P ISBN 0-7923-2731-4; Glick, Bernard R. and Thompson, John E., Methods in Plant Molecular Biology and Biotechnology, Boca Raton : CRC Press, 1993 360 S., ISBN 0-8493-5164-2). For example, rapeseed can be transformed via cotyledon or hypocotyl transformation (Moloney et al., 1989, Plant Cell Report 8:238-242; De Block et al., 1989, Plant Physiol. 91:694-701). Use of antibiotics for Agrobacterium and plant selection depends on the binary vector and the Agrobacterium strain used for transformation. Rapeseed selection is normally performed using kanamycin as selectable plant marker. Agrobacterium mediated gene transfer to flax can be performed using, for example, a technique described by Mlynarova et al., 1994, Plant Cell Report 13:282-285. Additionally, transformation of soybean can be performed using for example a technique described in European Patent No. 0424 047, U.S. Patent No. 5,322,783, European Patent No. 0397 687, U.S. Patent No. 5,376,543, or U.S. Patent No. 5,169,770. Transformation of maize can be achieved by particle bombardment, polyethylene glycol mediated DNA uptake, or via the silicon carbide fiber technique. (See, for example, Freeling and Walbot "The maize handbook" Springer Verlag: New York (1993) ISBN 3-540-97826-7). A specific example of maize transformation is found in U.S. Patent No. 5,990,387, and a specific example of wheat transformation can be found in PCT Application No. WO 93/07256.

According to the present invention, the introduced mutated PPO polynucleotide may be maintained in the plant cell stably if it is incorporated into a non-chromosomal autonomous replicon or integrated into the plant chromosomes. Alternatively, the introduced mutated PPO polynucleotide may be present on an extra-chromosomal non-replicating vector and be transiently expressed or transiently active. In one embodiment, a homologous recombinant microorganism can be created wherein the mutated PPO polynucleotide is integrated into a chromosome, a vector is prepared which contains at least a portion of an PPO gene into which a deletion, addition, or substitution has been introduced to thereby alter, e.g., functionally disrupt, the endogenous PPO gene and to create a mutated PPO gene. To create a point mutation via homologous recombination, DNA-RNA hybrids can be used in a technique known as chimeraplasty (Cole-Strauss et al., 1999, Nucleic Acids Research 27(5):1323-1330 and Kmiec, 1999, Gene therapy American Scientist 87(3):240-247). Other homologous recombination procedures in Triticum species are also well known in the art and are contemplated for use herein.

In the homologous recombination vector, the mutated PPO gene can be flanked at its 5' and 3' ends by an additional nucleic acid molecule of the PPO gene to allow for homologous recombination to occur between the exogenous mutated PPO gene carried by the vector and an endogenous PPO gene, in a microorganism or plant. The additional flanking PPO nucleic acid molecule is of sufficient length for successful homologous recombination with the endogenous gene. Typically, several hundreds of base pairs up to kilobases of flanking DNA (both at the 5' and 3' ends) are included in the vector (see e.g., Thomas, K. R., and Capecchi, M. R., 1987, Cell 51:503 for a description of homologous recombination vectors or Strepp et al., 1998, PNAS,

95(8):4368-4373 for cDNA based recombination in *Physcomitrella patens*). However, since the mutated PPO gene normally differs from the PPO gene at very few amino acids, a flanking sequence is not always necessary. The homologous recombination vector is introduced into a microorganism or plant cell (e.g., via polyethylene glycol mediated DNA), and cells in which the introduced mutated PPO gene has homologously recombined with the endogenous PPO gene are selected using art-known techniques.

In another embodiment, recombinant microorganisms can be produced that contain selected systems that allow for regulated expression of the introduced gene. For example, inclusion of a mutated PPO gene on a vector placing it under control of the lac operon permits expression of the mutated PPO gene only in the presence of IPTG. Such regulatory systems are well known in the art.

Another aspect of the invention pertains to host cells into which a recombinant expression vector of the invention has been introduced. The terms "host cell" and "recombinant host cell" are used interchangeably herein. It is understood that such terms refer not only to the particular subject cell but they also apply to the progeny or potential progeny of such a cell. Because certain modifications may occur in succeeding generations due to either mutation or environmental influences, such progeny may not, in fact, be identical to the parent cell, but are still included within the scope of the term as used herein. A host cell can be any prokaryotic or eukaryotic cell. For example, a mutated PPO polynucleotide can be expressed in bacterial cells such as *C. glutamicum*, insect cells, fungal cells, or mammalian cells (such as Chinese hamster ovary cells (CHO) or COS cells), algae, ciliates, plant cells, fungi or other microorganisms like *C. glutamicum*. Other suitable host cells are known to those skilled in the art.

A host cell of the invention, such as a prokaryotic or eukaryotic host cell in culture, can be used to produce (i.e., express) a mutated PPO polynucleotide. Accordingly, the invention further provides methods for producing mutated PPO polypeptides using the host cells of the invention. In one embodiment, the method comprises culturing the host cell of invention (into which a recombinant expression vector encoding a mutated PPO polypeptide has been introduced, or into which genome has been introduced a gene encoding a wild-type or mutated PPO polypeptide) in a suitable medium until mutated PPO polypeptide is produced. In another embodiment, the method further comprises isolating mutated PPO polypeptides from the medium or the host cell. Another aspect of the invention pertains to isolated mutated PPO polypeptides, and biologically active portions thereof. An "isolated" or "purified" polypeptide or biologically active portion thereof is free of some of the cellular material when produced by recombinant DNA techniques, or chemical precursors or other chemicals when chemically synthesized. The language "substantially free of cellular material" includes preparations of mutated PPO polypeptide in which the polypeptide is separated from some of the cellular components of the cells in which it is naturally or recombinantly produced. In one embodiment, the language "substantially free of cellular material" includes preparations of a mutated PPO polypeptide having less than about 30% (by dry weight) of non-mutated PPO material (also referred to herein as a "contaminating polypeptide"), more preferably less than about 20% of non-mutated PPO material, still more preferably less than about 10% of non-mutated PPO material, and most preferably less than about 5% non-mutated PPO

material.

When the mutated PPO polypeptide, or biologically active portion thereof, is recombinantly produced, it is also preferably substantially free of culture medium, i.e., culture medium represents
5 less than about 20%, more preferably less than about 10%, and most preferably less than about 5% of the volume of the polypeptide preparation. The language "substantially free of chemical precursors or other chemicals" includes preparations of mutated PPO polypeptide in which the polypeptide is separated from chemical precursors or other chemicals that are involved in the synthesis of the polypeptide. In one embodiment, the language "substantially free of chemical
10 precursors or other chemicals" includes preparations of a mutated PPO polypeptide having less than about 30% (by dry weight) of chemical precursors or non-mutated PPO chemicals, more preferably less than about 20% chemical precursors or non-mutated PPO chemicals, still more preferably less than about 10% chemical precursors or non-mutated PPO chemicals, and most preferably less than about 5% chemical precursors or non-mutated PPO chemicals. In preferred
15 embodiments, isolated polypeptides, or biologically active portions thereof, lack contaminating polypeptides from the same organism from which the mutated PPO polypeptide is derived. Typically, such polypeptides are produced by recombinant expression of, for example, a mutated PPO polypeptide in plants other than, or in microorganisms such as *C. glutamicum*, ciliates, algae, or fungi.

20 In other aspects, a method for treating a plant of the present invention is provided.

In some embodiments, the method comprises contacting the plant with an agronomically acceptable composition.

25 In another aspect, the present invention provides a method for preparing a descendent seed. The method comprises planting a seed of or capable of producing a plant of the present invention. In one embodiment, the method further comprises growing a descendent plant from the seed; and harvesting a descendant seed from the descendent plant. In other
30 embodiments, the method further comprises applying a PPO-inhibiting herbicides herbicidal composition to the descendent plant.

In another embodiment, the invention refers to harvestable parts of the transgenic plant according to the present invention. Preferably, the harvestable parts comprise the PPO nucleic
35 acid or PPO protein of the present invention. The harvestable parts may be seeds, roots, leaves and/or flowers comprising the PPO nucleic acid or PPO protein or parts thereof. Preferred parts of soy plants are soy beans comprising the PPO nucleic acid or PPO protein.

40 In another embodiment, the invention refers to products derived from a plant according to the present invention, parts thereof or harvestable parts thereof. A preferred plant product is fodder, seed meal, oil, or seed-treatment-coated seeds. Preferably, the meal and/or oil comprises the mutated PPO nucleic acids or PPO proteins of the present invention.

In another embodiment, the invention refers to a method for the production of a product, which method comprises

- a) growing the plants of the invention or obtainable by the methods of invention and
- b) producing said product from or by the plants of the invention and/or parts, e.g. seeds, of these plants.

In a further embodiment the method comprises the steps

- a) growing the plants of the invention,
- b) removing the harvestable parts as defined above from the plants and
- c) producing said product from or by the harvestable parts of the invention.

The product may be produced at the site where the plant has been grown, the plants and/or parts thereof may be removed from the site where the plants have been grown to produce the product. Typically, the plant is grown, the desired harvestable parts are removed from the plant, if feasible in repeated cycles, and the product made from the harvestable parts of the plant. The step of growing the plant may be performed only once each time the methods of the invention is performed, while allowing repeated times the steps of product production e.g. by repeated removal of harvestable parts of the plants of the invention and if necessary further processing of these parts to arrive at the product. It is also possible that the step of growing the plants of the invention is repeated and plants or harvestable parts are stored until the production of the product is then performed once for the accumulated plants or plant parts. Also, the steps of growing the plants and producing the product may be performed with an overlap in time, even simultaneously to a large extent or sequentially. Generally the plants are grown for some time before the product is produced.

In one embodiment the products produced by said methods of the invention are plant products such as, but not limited to, a foodstuff, feedstuff, a food supplement, feed supplement, fiber, cosmetic and/or pharmaceutical. Foodstuffs are regarded as compositions used for nutrition and/or for supplementing nutrition. Animal feedstuffs and animal feed supplements, in particular, are regarded as foodstuffs.

In another embodiment the inventive methods for the production are used to make agricultural products such as, but not limited to, plant extracts, proteins, amino acids, carbohydrates, fats, oils, polymers, vitamins, and the like.

It is possible that a plant product consists of one or more agricultural products to a large extent.

As described above, the present invention teaches compositions and methods for increasing the PPO-inhibiting tolerance of a crop plant or seed as compared to a wild-type variety of the plant or seed. In a preferred embodiment, the PPO-inhibiting tolerance of a crop plant or seed is increased such that the plant or seed can withstand a PPO-inhibiting herbicide application of preferably approximately 1-1000 g ai ha⁻¹, more preferably 1-200 g ai ha⁻¹, even more preferably 5-150 g ai

ha⁻¹, and most preferably 10-100 g ai ha⁻¹. As used herein, to “withstand” a PPO-inhibiting herbicide application means that the plant is either not killed or only moderately injured by such application. It will be understood by the person skilled in the art that the application rates may vary, depending on the environmental conditions such as temperature or humidity, and depending on the chosen kind of herbicide (active ingredient ai).

Furthermore, the present invention provides methods that involve the use of at least one PPO-inhibiting herbicide, optionally in combination with one or more herbicidal compounds B, and, optionally, a safener C, as described in detail supra.

In these methods, the PPO-inhibiting herbicide can be applied by any method known in the art including, but not limited to, seed treatment, soil treatment, and foliar treatment. Prior to application, the PPO-inhibiting herbicide can be converted into the customary formulations, for example solutions, emulsions, suspensions, dusts, powders, pastes and granules. The use form depends on the particular intended purpose; in each case, it should ensure a fine and even distribution of the compound according to the invention.

By providing plants having increased tolerance to PPO-inhibiting herbicide, a wide variety of formulations can be employed for protecting plants from weeds, so as to enhance plant growth and reduce competition for nutrients. A PPO-inhibiting herbicide can be used by itself for pre-emergence, post-emergence, pre-planting, and at-planting control of weeds in areas surrounding the crop plants described herein, or a PPO-inhibiting herbicide formulation can be used that contains other additives. The PPO-inhibiting herbicide can also be used as a seed treatment. Additives found in a PPO-inhibiting herbicide formulation include other herbicides, detergents, adjuvants, spreading agents, sticking agents, stabilizing agents, or the like. The PPO-inhibiting herbicide formulation can be a wet or dry preparation and can include, but is not limited to, flowable powders, emulsifiable concentrates, and liquid concentrates. The PPO-inhibiting herbicide and herbicide formulations can be applied in accordance with conventional methods, for example, by spraying, irrigation, dusting, or the like.

Suitable formulations are described in detail in PCT/EP2009/063387 and PCT/EP2009/063386, which are incorporated herein by reference.

It should also be understood that the foregoing relates to preferred embodiments of the present invention and that numerous changes may be made therein without departing from the scope of the invention. The invention is further illustrated by the following examples, which are not to be construed in any way as imposing limitations upon the scope thereof. On the contrary, it is to be clearly understood that resort may be had to various other embodiments, modifications, and equivalents thereof, which, after reading the description herein, may suggest themselves to those skilled in the art without departing from the spirit of the present invention and/or the scope of the appended claims.

EXAMPLES

EXAMPLE 1: Site-directed mutagenesis of *Amaranthus* PPO

All nucleic acid coding sequence and all single and double mutants based on SEQ ID NO: 1, 3, 5, 7, 9, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, 47, were synthesized and cloned by Geneart (Geneart AG, Regensburg, Germany). Rational design mutants were synthesized by Geneart. Random PPO gene libraries were synthesized by Geneart. Plasmids were isolated from *E. coli* TOP10 by performing a plasmid miniprep and confirmed by DNA sequencing.

EXAMPLE 2: Expression and purification of recombinant wildtype and mutant PPO

(Taken from: Franck E. Dayan, Pankaj R. Daga, Stephen O. Duke, Ryan M. Lee, Patrick J. Tranel, Robert J. Doerksen. Biochemical and structural consequences of a glycine deletion in the α -8 helix of protoporphyrinogen oxidase. *Biochimica et Biophysica Acta* 1804 (2010), 1548-56) Clones in pRSET vector were transformed into BL21(DE3)-pLysS strain of *E. coli*. Cells were grown in 250 mL of LB with 100 $\mu\text{g mL}^{-1}$ of carbenicillin, shaking overnight at 37 °C. Cultures were diluted in 1 L of LB with antibiotic and grown at 37 °C shaking for 2 h, induced with 1 mM IPTG and grown at 25 °C shaking for 5 more hours. The cells were harvested by centrifugation at 1600 \times g, washed with 0.09% NaCl, and stored at -80 °C. Cells were lysed using a French press at 140 MPa in 50 mM sodium phosphate pH 7.5, 1 M NaCl, 5 mM imidazole, 5% glycerol, and 1 $\mu\text{g mL}^{-1}$ leupeptin. Following lysis, 0.5 U of benzonase (Novagen, EMD Chemicals, Inc., Gibbstown, NJ) and PMSF (final concentration of 1 mM) were added. Cell debris was removed by centrifugation at 3000 \times g. His-tagged PPO proteins were purified on a nickel activated Hitrap Chelating HP column (GE Healthcare Bio-Sciences Corp., Piscataway, NJ) equilibrated with 20 mM sodium phosphate pH 8.0, 50 mM NaCl, 5 mM imidazole, 5 mM MgCl_2 , 0.1mM EDTA, and 17% glycerol. PPO is eluted with 250 mM imidazole. The active protein was desalted on a PD-10 column (GE Healthcare Bio-Sciences Corp., Piscataway, NJ) equilibrated with a 20 mM sodium phosphate buffer, pH 7.5, 5 mM MgCl_2 , 1 mM EDTA and 17% glycerol. Each litre of culture provided approximately 10 mg of pure PPO, which was stored at -20 °C until being used in assays.

EXAMPLE 3: PPO Enzyme Assay (non-recombinant)

PPO protein (EC 1.3.3.4) was extracted from coleoptiles or shoots (150 g fresh weight) of dark-grown corn, black nightshade, morning glory, and velvetleaf seedlings as described previously (Grossmann et al. 2010). Before harvesting, the seedlings were allowed to green for 2 hours in the light in order to achieve the highest specific enzyme activities in the thylakoid fractions at low chlorophyll concentrations. At high chlorophyll concentrations significant quenching of fluorescence occurs, which limits the amount of green thylakoids that can be used in the test. Plant materials were homogenized in the cold with a Braun blender using a fresh-weight-to-volume ratio of 1:4. Homogenization buffer consisted of tris(hydroxymethyl)aminomethane (Tris)-HCl (50 mM; pH 7.3), sucrose (0.5 M), magnesium chloride (1 mM), ethylenediaminetetraacetic acid (EDTA) (1 mM) and bovine serum albumin (2 g L^{-1}). After filtration through four layers of Miracloth, crude plastid preparations were obtained after centrifugation at 10 000 \times g for 5 min and resuspension in homogenization buffer before centrifugation at 150 \times g for 2 min to remove crude cell debris. The supernatant was centrifuged at 4000 \times g for 15 min and the pellet fraction was resuspended in 1 ml of a buffer containing Tris-HCl (50 mM; pH 7.3), EDTA (2 mM), leupeptin (2 μM), pepstatin (2 μM) and glycerol (200 ml L^{-1}) and stored at -80°C until use. Protein was determined in the enzyme extract with bovine serum albumin as a standard. PPO activity was assayed fluorometrically by

monitoring the rate of Proto formation from chemically reduced protoporphyrinogen IX under initial velocity conditions. The assay mixture consisted of Tris-HCl (100 mM; pH 7.3), EDTA (1 mM), dithiothreitol (5 mM), Tween 80 (0.085%), protoporphyrinogen IX (2 μ M), and 40 μ g extracted protein in a total volume of 200 μ l. The reaction was initiated by addition of substrate

5 protoporphyrinogen IX at 22°C. saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control were prepared in dimethyl sulfoxide (DMSO) solution (0.1 mM

10 concentration of DMSO in the assay) and added to the assay mixture in concentrations of 0.005 pM to 5 μ M before incubation. Fluorescence was monitored directly from the assay mixture using a POLARstar Optima / Galaxy (BMG) with excitation at 405 nm and emission monitored at 630 nm. Non-enzymatic activity in the presence of heat-inactivated extract was negligible. Inhibition of enzyme activity induced by the herbicide was expressed as percentage inhibition relative to untreated controls. Molar concentrations of compound required for 50% enzyme inhibition (IC₅₀

15 values) were calculated by fitting the values to the dose-response equation using non-linear regression analysis.

EXAMPLE 4: PPO Enzyme Assay (recombinant)

Proto was purchased from Sigma-Aldrich (Milwaukee,WI). Protogen was prepared according to

20 Jacobs and Jacobs (N.J. Jacobs, J.M. Jacobs, Assay for enzymatic protoporphyrinogen oxidation, a late step in heme synthesis, Enzyme 28 (1982) 206–219) . Assays were conducted in 100 mM sodium phosphate pH 7.4 with 0.1 mM EDTA, 0.1% Tween 20, 5 μ M FAD, and 500mM imidazole. Dose–response curves with the PPO inhibitors saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS

25 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control, and MC-15608 were obtained in the presence of 150 μ M Protogen. Dose response was measured between the inhibitor concentration range of 1,00E-05 M to 1,00E-12 M. The excitation and emission bandwidths were set at 1.5 and 30 nm, respectively. All assays were made in duplicates or triplicates and measured using a POLARstar

30 Optima / Galaxy (BMG) with excitation at 405 nm and emission monitored at 630 nm. Molar concentrations of compound required for 50% enzyme inhibition (IC₅₀ values) were calculated by fitting the values to the dose-response equation using non-linear regression analysis. The results are shown in Table 4.

Table 4a: IC₅₀ values for various mutated PPO (mutated PPO)

Amino Acid Substitution	SEQ. ID NO.	Relative Ezyme Activity (FU/min)	Saflufenacil	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione
				IC ₅₀ (M)
PPO herbicide sensitive PPO2 WC	2	1000	1,86E-09	5,17E-10
PPO herbicide sensitive PPO2 AC	4	800	1,78E-10	5,96E-11

dG210	6 & 8	80	1,60E-06	2,12E-09
R128L	2	700	2,22E-07	7,73E-10
R128L	2	700	2,22E-07	7,73E-10
R128A	2	730	1,29E-07	1,40E-10
R128C	4	515	5,57E-07	1,16E-10
R128D	4	ND	ND	ND
R128E	4	ND	ND	ND
R128F	4	280	5,25E-07	2,21E-10
R128G	4	440	9,91E-07	4,71E-11
R128H	4	640	1,02E-08	6,15E-11
R128I	4	250	3,65E-07	9,80E-11
R128K	4	180	9,65E-11	ND
R128L	4	280	3,88E-07	1,01E-10
R128M	4	200	6,97E-07	3,56E-11
R128N	4	420	5,79E-07	4,33E-11
R128P	4	ND	ND	ND
R128Q	4	480	1,94E-07	1,09E-11
R128S	4	490	2,46E-07	1,12E-11
R128T	4	510	2,11E-07	3,79E-11
R128V	4	600	2,49E-07	6,70E-11
R128W	4	ND	ND	ND
R128Y	4	230	2,19E-06	5,77E-11
F420A	4	ND	ND	ND
F420V	2	200	1,59E-06	1,61E-09
F420V	2	330		1,61E-09
F420M	2	350	6,77E-07	2,75E-10
F420M	2	700		2,18E-10
F420L	2	200	7,20E-06	9,93E-10
F420I	2	200	9,19E-07	4,95E-10
R128A, F420V	2	510	>0,00001	2,50E-08
R128A+F420M	2	400	>0,00001	6,24E-09
R128A+F420L	2	300	>0,00001	1,62E-08
R128A+F420I	2	330	>0,00001	2,46E-08
R128A_F420A	4	ND	ND	ND
R128L_F420A	4	ND	ND	ND
R128L_F420L	4	300	>0,00001	1,71E-06
R128L_F420I	4	450	>0,00001	1,23E-06
R128L_F420V	4	300	>0,00001	1,51E-06
R128L_F420M	4	400	>0,00001	2,46E-07
R128I_F420A	4	ND	ND	ND
R128I_F420L	4	200	>0,00001	4,66E-07
R128I_F420I	4	100	>0,00001	4,33E-07
R128I_F420V	4	470	>0,00001	4,24E-07

R128I_F420M	4	500	>0,00001	5,82E-08
R128V_F420A	4	ND	ND	ND
R128V_F420L	4	370	>0,00001	4,41E-07
R128V_F420I	4	300	>0,00001	2,23E-07
R128V_F420V	4	300	>0,00001	4,46E-07
R128V_F420M	4	460	>0,00001	4,27E-08
R128M_F420A	4	ND	ND	ND
R128M_F420L	4	300	>0,00001	6,95E-07
R128M_F420I	4	350	>0,00001	4,45E-07
R128M_F420V	4	270	>0,00001	7,04E-07
R128M_F420M	4	480	>0,00001	7,05E-08

Table 4b: IC50 values for various mutated PPO (mutated PPO)

Construct	SEQ. ID NO.	rate (FU/min)	Saflufenacil	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione
			IC50 (M)	
PPO herbicide sensitive PPO2 WC	2	1000	1,86E-09	5,17E-10
PPO herbicide sensitive PPO2 AC	4	800	1,78E-10	5,96E-11
dG210	6 & 8	80	1,60E-06	2,12E-09
R128L	2	700	2,22E-07	7,73E-10
R128K	4	180	9,65E-11	not determined
R128Q	4	481	1,94E-07	1,09E-11
R128S	4	491	2,46E-07	1,13E-11
R128M	4	200	6,97E-07	3,56E-11
R128T	4	721	2,11E-07	3,79E-11
R128N	4	421	5,79E-07	4,33E-11
R128G	4	436	9,91E-07	4,71E-11
R128Y	4	230	2,19E-06	5,77E-11
R128H	4	636	1,02E-08	6,15E-11
R128V	4	923	2,49E-07	7,00E-11
R128I	4	250	3,65E-07	9,80E-11
R128C	4	933	5,57E-07	1,16E-10
R128A	4	731	1,29E-07	1,40E-10
R128F	4	278	5,25E-07	2,21E-10
R128L	4	700	2,22E-07	7,73E-10
R128A, L397D	2	98	≥1,00E-5	5,90E-09
R128A, F420M	2	378	≥1,00E-5	6,24E-09
R128Q, F420M	4	473	≥1,00E-5	1,54E-08

R128A, F420L	2	281	$\geq 1,00E-5$	1,62E-08
R128S, F420M	4	310	$\geq 1,00E-5$	1,77E-08
R128C, F420M	4	329	$\geq 1,00E-5$	2,30E-08
R128A, F420I	2	330	$\geq 1,00E-5$	2,46E-08
R128A, F420V	2	512	$\geq 1,00E-5$	2,50E-08
R128H, F420M	4	252	$\geq 1,00E-5$	2,92E-08
R128G, F420M	4	100	$\geq 1,00E-5$	3,02E-08
R128V, F420M	4	666	$\geq 1,00E-5$	4,27E-08
R128S, F420I	4	150	$\geq 1,00E-5$	4,64E-08
R128Q, F420I	4	202	$\geq 1,00E-5$	5,43E-08
R128T, F420M	4	303	$\geq 1,00E-5$	5,54E-08
R128I, F420M	4	497	$\geq 1,00E-5$	5,82E-08
R128S, F420L	4	110	$\geq 1,00E-5$	6,24E-08
R128Q, F420L	4	150	$\geq 1,00E-5$	6,90E-08
R128M, F420M	4	479	$\geq 1,00E-5$	7,05E-08
R128F, F420M	4	120	$\geq 1,00E-5$	7,84E-08
R128M, F420M	4	306	$\geq 1,00E-5$	8,26E-08
R128N, F420M	4	208	$\geq 1,00E-5$	1,01E-07
R128C, F420I	4	204	$\geq 1,00E-5$	1,20E-07
R128M, F420I	4	250	$\geq 1,00E-5$	1,44E-07
R128H, F420I	4	195	$\geq 1,00E-5$	1,47E-07
R128T, F420V	4	120	$\geq 1,00E-5$	1,50E-07
R128Y, F420M	4	200	$\geq 1,00E-5$	1,61E-07
R128H, F420L	4	185	$\geq 1,00E-5$	1,69E-07
R128N, F420I	4	100	$\geq 1,00E-5$	1,75E-07
R128H, F420V	4	74	$\geq 1,00E-5$	1,82E-07
R128C, F420L	4	217	$\geq 1,00E-5$	1,89E-07
R128Q, F420V	4	113	$\geq 1,00E-5$	2,02E-07
R128N, F420L	4	100	$\geq 1,00E-5$	2,10E-07
R128C, F420V	4	223	$\geq 1,00E-5$	2,16E-07
R128V, F420I	4	300	$\geq 1,00E-5$	2,23E-07
R128T, F420I	4	238	$\geq 1,00E-5$	2,29E-07
R128L, F420M	4	518	$\geq 1,00E-5$	2,46E-07
R128M, F420L	4	211	$\geq 1,00E-5$	2,49E-07
R128T, F420L	4	157	$\geq 1,00E-5$	3,97E-07
R128M, F420V	4	127	$\geq 1,00E-5$	4,00E-07
R128I, F420V	4	464	$\geq 1,00E-5$	4,24E-07
R128I, F420I	4	128	$\geq 1,00E-5$	4,33E-07
R128V, F420L	4	365	$\geq 1,00E-5$	4,41E-07
R128M, F420I	4	343	$\geq 1,00E-5$	4,45E-07
R128V, F420V	4	300	$\geq 1,00E-5$	4,47E-07
R128I, F420L	4	281	$\geq 1,00E-5$	4,66E-07
R128Y, F420I	4	90	$\geq 1,00E-5$	6,11E-07
R128A, Δ G210	4	170	$\geq 1,00E-5$	6,57E-07
R128M, F420L	4	300	$\geq 1,00E-5$	6,95E-07
R128M, F420V	4	261	$\geq 1,00E-5$	7,04E-07

R128F, F420L	4	101	$\geq 1,00E-5$	8,68E-07
R128L, F420I	4	453	$\geq 1,00E-5$	1,23E-06
R128L, F420V	4	289	$\geq 1,00E-5$	1,51E-06
R128L, F420L	4	300	$\geq 1,00E-5$	1,71E-06
R128D	4	Low or no enzyme activity measured		
R128E	4	Low or no enzyme activity measured		
R128P	4	Low or no enzyme activity measured		
R128W	4	Low or no enzyme activity measured		
R128A, F420A	2	Low or no enzyme activity measured		
R128L, F420A	4	Low or no enzyme activity measured		
R128I, F420A	4	Low or no enzyme activity measured		
R128V, F420A	4	Low or no enzyme activity measured		
R128M, F420A	4	Low or no enzyme activity measured		
R128M, F420A	4	Low or no enzyme activity measured		
R128N, F420A	4	Low or no enzyme activity		

		measured		
R128Y, F420A	4	Low or no enzyme activity measured		
R128Y, F420L	4	Low or no enzyme activity measured		
R128Y, F420V	4	Low or no enzyme activity measured		
R128G, F420A	4	Low or no enzyme activity measured		
R128G, F420L	4	Low or no enzyme activity measured		
R128G, F420I	4	Low or no enzyme activity measured		
R128G, F420V	4	Low or no enzyme activity measured		
R128H, F420A	4	Low or no enzyme activity measured		
R128N, F420V	4	Low or no enzyme activity measured		
R128C, F420A	4	Low or no enzyme activity measured		
R128F, F420A	4	Low or no enzyme activity measured		
R128F, F420I	4	Low or no enzyme		

		activity measured		
R128F, F420V	4	Low or no enzyme activity measured		
R128S, F420A	4	Low or no enzyme activity measured		
R128S, F420V	4	Low or no enzyme activity measured		
R128T, F420A	4	Low or no enzyme activity measured		
R128Q, F420A	4	Low or no enzyme activity measured		

IC₅₀ (M): Concentration of inhibitor required for 50% inhibition of enzyme activity; $\geq 1,00E-5$: indicates a very high IC₅₀ over the measurement boundaries, which reflects very high in vitro tolerance.

Table 4c

Common Name	IUPAC Name	SEQ ID	Mutation	rate (FU/min)	IC50 (M)	inhibition (%) at 1x10 ⁻⁵ M
FOMESAFEN		2 or 4	WT	650	1,32E-09	
FOMESAFEN		4	R128A, F420M	362	6,60E-06	
FOMESAFEN		4	R128A, F420L	316	9,91E-06	
FOMESAFEN		4	R128A, F420V	478	1,61E-06	
FOMESAFEN		4	R128I, F420L	202	≥ 1,00E-05	38
FOMESAFEN		4	R128I, F420V	292	2,79E-06	
FOMESAFEN		4	R128V, F420M	413	≥ 1,00E-05	47
FOMESAFEN		4	R128M, F420M	289	≥ 1,00E-05	48
FOMESAFEN		4	R128Y, F420I	99	2,15E-05	
FOMESAFEN		4	R128Y, F420M	174	≥ 1,00E-05	28
FOMESAFEN		4	R128N, F420M	153	1,07E-05	
FOMESAFEN		4	R128C, F420L	192	≥ 1,00E-05	42
FOMESAFEN		4	R128C, F420V	160	2,36E-06	
FOMESAFEN		4	R128C, F420M	277	1,10E-05	
FOMESAFEN		4	R128H, F420M	184	2,91E-06	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	2 or 4	WT	650	2,93E-10	

LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128A, F420M	362	4,57E-08	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128A, F420L	316	6,88E-08	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128A, F420V	478	8,45E-09	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128I, F420L	202	1,30E-07	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128I, F420V	292	1,40E-08	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128V, F420M	413	9,41E-08	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128M, F420M	289	1,31E-07	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128Y, F420I	99	4,80E-08	

LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128Y, F420M	174	1,43E-07	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128N, F420M	153	1,67E-07	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128C, F420L	192	1,42E-07	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128C, F420V	160	1,50E-08	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128C, F420M	277	6,39E-08	
LACTOFEN	(2-ethoxy-1-methyl-2-oxo-ethyl) 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitro-benzoate	4	R128H, F420M	184	6,13E-08	
BUTAFENACIL		2 or 4	WT	650	1,38E-10	
BUTAFENACIL		4	R128A, F420M	362	1,40E-08	
BUTAFENACIL		4	R128A, F420L	316	9,17E-08	
BUTAFENACIL		4	R128A, F420V	478	2,51E-08	
BUTAFENACIL		4	R128I, F420L	202	8,02E-08	
BUTAFENACIL		4	R128I, F420V	292	2,56E-08	

BUTAFENACIL		4	R128V, F420M	413	1,05E-08	
BUTAFENACIL		4	R128M, F420M	289	4,38E-08	
BUTAFENACIL		4	R128Y, F420I	99	5,47E-08	
BUTAFENACIL		4	R128Y, F420M	174	5,04E-08	
BUTAFENACIL		4	R128N, F420M	153	2,84E-08	
BUTAFENACIL		4	R128C, F420L	192	1,10E-07	
BUTAFENACIL		4	R128C, F420V	160	6,69E-08	
BUTAFENACIL		4	R128C, F420M	277	2,31E-08	
BUTAFENACIL		4	R128H, F420M	184	1,28E-08	
CARFENTRAZONE-ETHYL		2 or 4	WT	650	1,03E-09	
CARFENTRAZONE-ETHYL		4	R128A, F420M	362	6,72E-08	
CARFENTRAZONE-ETHYL		4	R128A, F420L	316	4,29E-07	
CARFENTRAZONE-ETHYL		4	R128A, F420V	478	7,97E-07	
CARFENTRAZONE-ETHYL		4	R128I, F420L	202	1,61E-07	
CARFENTRAZONE-ETHYL		4	R128I, F420V	292	2,07E-07	
CARFENTRAZONE-ETHYL		4	R128V, F420M	413	2,29E-08	
CARFENTRAZONE-ETHYL		4	R128M, F420M	289	7,86E-08	

CARFENTRAZONE-ETHYL	4	R128Y, F420I	99	2,82E-07	
CARFENTRAZONE-ETHYL	4	R128Y, F420M	174	8,52E-08	
CARFENTRAZONE-ETHYL	4	R128N, F420M	153	1,88E-07	
CARFENTRAZONE-ETHYL	4	R128C, F420L	192	3,08E-07	
CARFENTRAZONE-ETHYL	4	R128C, F420V	160	3,96E-07	
CARFENTRAZONE-ETHYL	4	R128C, F420M	277	2,99E-08	
CARFENTRAZONE-ETHYL	4	R128H, F420M	184	1,21E-07	
ACIFLUORFEN	2 or 4	WT	650	3,36E-08	
ACIFLUORFEN	4	R128A, F420M	362	$\geq 1,00E-05$	27
ACIFLUORFEN	4	R128A, F420L	316	$\geq 1,00E-05$	20
ACIFLUORFEN	4	R128A, F420V	478	6,67E-06	
ACIFLUORFEN	4	R128I, F420L	202	$\geq 1,00E-05$	16
ACIFLUORFEN	4	R128I, F420V	292	1,21E-05	

ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128V, F420M	413	$\geq 1,00\text{E-}05$	17
ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128M, F420M	289	$\geq 1,00\text{E-}05$	21
ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128Y, F420I	99	$\geq 1,00\text{E-}05$	21
ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128Y, F420M	174	$\geq 1,00\text{E-}05$	15
ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128N, F420M	153	$\geq 1,00\text{E-}05$	39
ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128C, F420L	192	$\geq 1,00\text{E-}05$	17
ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128C, F420V	160	6,72E-06	
ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128C, F420M	277	$\geq 1,00\text{E-}05$	33
ACIFLUORFEN	5-(2-CHLORO-4-TRIFLUOROMETHYL-PHENOXY)-2-NITRO-BENZOIC ACID	4	R128H, F420M	184	$\geq 1,00\text{E-}05$	48
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	2 or 4	WT	650	9,58E-11	
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128A, F420M	362	8,43E-06	

FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128A, F420L	316	$\geq 1,00\text{E-}05$	-8
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128A, F420V	478	6,34E-06	
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128I, F420L	202	$\geq 1,00\text{E-}05$	9
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128I, F420V	292	$\geq 1,00\text{E-}05$	41
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128V, F420M	413	$\geq 1,00\text{E-}05$	34
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128M, F420M	289	$\geq 1,00\text{E-}05$	21
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128Y, F420I	99	$\geq 1,00\text{E-}05$	19
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128Y, F420M	174	$\geq 1,00\text{E-}05$	-2

FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128N, F420M	153	6,15E-06	
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128C, F420L	192	$\geq 1,00E-05$	-11
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128C, F420V	160	7,28E-06	
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128C, F420M	277	$\geq 1,00E-05$	48
FLUMIOXAZIN	2-(7-fluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128H, F420M	184	$\geq 1,00E-05$	30
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisindol-2-yl)phenyl]prop-2-enoate	2 or 4	WT	650	6,69E-10	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisindol-2-yl)phenyl]prop-2-enoate	4	R128A, F420M	362	1,60E-06	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisindol-2-yl)phenyl]prop-2-enoate	4	R128A, F420L	316	$\geq 1,00E-05$	48

CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128A, F420V	478	5,43E-06	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128I, F420L	202	9,51E-06	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128I, F420V	292	4,72E-06	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128V, F420M	413	1,78E-06	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128M, F420M	289	3,84E-06	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128Y, F420I	99	$\geq 1,00E-05$	38
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128Y, F420M	174	1,08E-05	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128N, F420M	153	$\geq 1,00E-05$	48

CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128C, F420L	192	≥ 1,00E-05	42
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128C, F420V	160	9,43E-06	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128C, F420M	277	2,45E-06	
CINIDON-ETHYL	ethyl (Z)-2-chloro-3-[2-chloro-5-(1,3-dioxo-4,5,6,7-tetrahydroisoindol-2-yl)phenyl]prop-2-enoate	4	R128H, F420M	184	≥ 1,00E-05	41
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	2 or 4	WT	650	1,04E-09	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128A, F420M	365	2,17E-07	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128A, F420L	343	5,58E-07	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128A, F420V	550	2,35E-08	

OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128I, F420L	196	4,21E-06	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128I, F420V	326	1,98E-07	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128V, F420M	482	1,05E-06	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128M, F420M	323	7,36E-07	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128Y, F420I	75	1,17E-06	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128Y, F420M	175	1,13E-06	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128N, F420M	174	3,91E-07	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128C, F420L	188	1,49E-06	

OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128C, F420V	225	6,52E-08	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128C, F420M	271	4,16E-07	
OXIFLUORFEN	2-CHLORO-1-(3-ETHOXY-4-NITROPHENOXY)-4-(TRIFLUOROMETHYL)BENZENE	4	R128H, F420M	196	3,68E-07	
OXADIARGYL		2 or 4	WT	650	3,64E-10	
OXADIARGYL		4	R128A, F420M	365	1,97E-08	
OXADIARGYL		4	R128A, F420L	343	1,37E-06	
OXADIARGYL		4	R128A, F420V	550	4,38E-08	
OXADIARGYL		4	R128I, F420L	196	8,64E-07	
OXADIARGYL		4	R128I, F420V	326	2,76E-08	
OXADIARGYL		4	R128V, F420M	482	3,40E-08	
OXADIARGYL		4	R128M, F420M	323	3,33E-08	
OXADIARGYL		4	R128Y, F420I	75	1,73E-07	
OXADIARGYL		4	R128Y, F420M	175	3,60E-08	
OXADIARGYL		4	R128N, F420M	174	1,28E-07	
OXADIARGYL		4	R128C, F420L	188	3,01E-06	
OXADIARGYL		4	R128C, F420V	225	1,46E-07	
OXADIARGYL		4	R128C, F420M	271	6,24E-08	
OXADIARGYL		4	R128H, F420M	196	1,32E-08	

S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	2 or 4	WT	650	1,35E-10	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128A, F420M	365	3,71E-08	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128A, F420L	343	2,77E-07	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128A, F420V	550	4,75E-08	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128I, F420L	196	2,01E-07	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128I, F420V	326	4,38E-08	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128V, F420M	482	3,58E-08	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128M, F420M	323	4,83E-08	

S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128Y, F420I	75	4,64E-07	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128Y, F420M	175	8,92E-08	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128N, F420M	174	1,92E-07	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128C, F420L	188	6,81E-07	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128C, F420V	225	1,24E-07	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128C, F420M	271	6,95E-08	
S-3100	ethyl 2-[[[3-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-pyridyl]oxy]acetate	4	R128H, F420M	196	4,18E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-	2 or 4	WT	650	5,17E-10	

	triazinane-2,4-dione						
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128A, F420M	321	7,02E-09		
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128A, F420M	362	7,95E-09		
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128A, F420M	365	6,10E-09		
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128A, F420L	316	2,96E-08		
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128A, F420L	343	1,56E-08		
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128A, F420V	478	4,14E-08		

	benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione					
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128A, F420V	550	2,13E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128A, F420V	555	3,99E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128I, F420L	202	4,05E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128I, F420L	196	2,45E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128I, F420I	95	1,38E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128I, F420V	292	2,14E-07	

	benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione					
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128I, F420V	326	3,15E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128I, F420M	328	6,10E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128V, F420M	413	6,50E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128V, F420M	482	4,86E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128M, F420M	235	7,69E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128M, F420M	289	7,07E-08	

	benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione					
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128M, F420M	323	4,84E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128Y, F420I	99	4,82E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128Y, F420I	75	2,63E-06	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128Y, F420M	174	2,85E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128Y, F420M	175	1,02E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128G, F420M	153	1,26E-08	

	benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione					
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128Q, F420M	432	1,07E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128H, F420L	193	7,98E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128H, F420I	191	8,22E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128N, F420M	153	7,12E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128N, F420M	174	4,97E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128C, F420L	192	1,00E-07	

	benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione					
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128C, F420L	188	1,83E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128C, F420V	160	1,66E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128C, F420V	225	2,66E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128C, F420M	277	2,53E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128C, F420M	271	2,33E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128F, F420L	129	1,01E-06	

	benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione					
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128F, F420M	136	1,21E-07	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128S, F420M	328	2,40E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128T, F420M	275	4,33E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128H, F420V	95	7,63E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128H, F420M	184	2,64E-08	
BAS 850H	1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione	4	R128H, F420M	196	2,13E-08	

	benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione					
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	2 or 4	WT	650	1,46E-10	
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128A, F420M	365	6,41E-07	
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128A, F420L	343	1,14E-05	
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128A, F420V	550	2,74E-07	
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128I, F420L	196	≥ 1,00E-05	6
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128I, F420V	326	4,32E-06	
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128V, F420M	482	3,11E-06	

850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128M, F420M	323	≥ 1,00E-05	48
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128Y, F420I	75	≥ 1,00E-05	32
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128Y, F420M	175	≥ 1,00E-05	41
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128N, F420M	174	≥ 1,00E-05	43
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128C, F420L	188	≥ 1,00E-05	11
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128C, F420V	225	3,70E-06	
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128C, F420M	271	3,57E-06	
850 analogon	2-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)-4,5,6,7-tetrahydroisindole-1,3-dione	4	R128H, F420M	196	3,07E-06	

850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	2 or 4	WT	650	3,15E-10	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128A, F420M	365	2,56E-09	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128A, F420L	343	1,62E-08	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128A, F420V	550	6,33E-09	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128I, F420L	196	2,69E-07	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128I, F420V	326	9,01E-08	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128V, F420M	482	4,65E-08	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128M, F420M	323	4,94E-08	

850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128Y, F420I	75	4,46E-07	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128Y, F420M	175	1,13E-07	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128N, F420M	174	5,94E-08	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128C, F420L	188	6,72E-08	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128C, F420V	225	2,60E-08	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128C, F420M	271	1,11E-08	
850 analogon	1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione	4	R128H, F420M	196	1,05E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-	2 or 4	WT	650	4,11E-10	

yl]phenoxy]phenoxy]-2-methoxy-acetate						
methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420M	321	8,19E-09		
methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420L	343	4,70E-08		
methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420V	555	2,32E-08		
methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420L	196	7,13E-08		
methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420I	95	2,27E-08		
methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420V	326	1,71E-08		

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420M	328	1,15E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128V, F420M	482	1,49E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128M, F420M	235	1,62E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128Y, F420I	75	2,86E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128G, F420M	153	4,76E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128Q, F420M	432	7,14E-08	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128H, F420L	193	4,47E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128H, F420I	191	7,54E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128N, F420M	174	1,20E-07	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128C, F420V	225	1,16E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128C, F420M	271	1,16E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128F, F420L	129	4,84E-08	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128F, F420M	136	2,81E-09	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128S, F420M	328	3,62E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128T, F420M	275	2,79E-08	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128H, F420V	95	6,93E-09	
	methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128H, F420M	196	1,76E-08	
	2-ethoxyethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-	2 or 4	WT	650	3,80E-10	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	2-ethoxyethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420M	321	1,51E-08	
	2-ethoxyethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420V	555	2,92E-08	
	2-ethoxyethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420M	328	1,39E-08	
	2-ethoxyethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128M, F420M	235	2,24E-08	
	2-ethoxyethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128S, F420M	328	4,68E-08	
	2-ethoxyethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128T, F420M	275	2,93E-08	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	2 or 4	WT	650	5,23E-10	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420M	321	2,27E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420L	343	9,37E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420V	555	4,16E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420L	196	1,07E-07	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420I	95	1,82E-06	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420V	326	3,78E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420M	328	1,06E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128V, F420M	482	1,49E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128M, F420M	235	3,22E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128Y, F420I	75	6,82E-07	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128G, F420M	153	5,14E-08	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128Q, F420M	432	1,72E-07	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128H, F420L	193	6,93E-07	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128H, F420I	191	1,31E-06	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128N, F420M	174	1,48E-07	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128C, F420V	225	1,01E-07	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128C, F420M	271	2,98E-08	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128F, F420L	129	1,18E-06	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128F, F420M	136	6,26E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128S, F420M	328	5,24E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128T, F420M	275	1,17E-07	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128H, F420V	95	9,06E-08	
	cyclohexyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128H, F420M	196	2,97E-07	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	4-pyridylmethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	2 or 4	WT	650	4,27E-10	
	4-pyridylmethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420M	321	1,22E-08	
	4-pyridylmethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420V	555	2,61E-08	
	4-pyridylmethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420M	328	1,56E-08	
	4-pyridylmethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128M, F420M	235	3,34E-08	
	4-pyridylmethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128S, F420M	328	5,65E-08	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	4-pyridylmethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128T, F420M	275	5,88E-08	
	(1-methylcyclopropyl)methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	2 or 4	WT	650	4,16E-10	
	(1-methylcyclopropyl)methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420M	321	1,19E-08	
	(1-methylcyclopropyl)methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420V	555	4,25E-08	
	(1-methylcyclopropyl)methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420M	328	1,37E-08	
	(1-methylcyclopropyl)methyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128M, F420M	235	2,47E-08	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate					
	(1-methylcyclopropyl)methyl 2-[2-(2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128S, F420M	328	6,94E-08	
	(1-methylcyclopropyl)methyl 2-[2-(2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128T, F420M	275	5,77E-08	
	2,2-difluoroethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	2 or 4	WT	650	4,43E-10	
	2,2-difluoroethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420M	321	4,93E-08	
	2,2-difluoroethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128A, F420V	555	6,42E-08	
	2,2-difluoroethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)]pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128I, F420M	328	4,61E-08	

	(trifluoromethyl)pyrimidin-1-yl]phenoxy]-2-methoxy-acetate					
	2,2-difluoroethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128M, F420M	235	1,06E-07	
	2,2-difluoroethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128S, F420M	328	9,94E-08	
	2,2-difluoroethyl 2-[2-[2-chloro-4-fluoro-5-[3-methyl-2,6-dioxo-4-(trifluoromethyl)pyrimidin-1-yl]phenoxy]phenoxy]-2-methoxy-acetate	4	R128T, F420M	275	1,50E-07	

IC50 (M): Concentration of inhibitor required for 50% inhibition of enzyme activity; $\geq 1,00E-5$: indicates a very high IC50 over the measurement boundaries, which reflects very high in vitro tolerance.

EXAMPLE 5: Engineering PPO-derivative herbicide tolerant plants having wildtype or mutated PPO sequences.

PPO-derivative herbicide tolerant soybean (*Glycine max*), corn (*Zea mays*), and Canola (*Brassica napus* or *Brassica Rapa var.* or *Brassica campestris L.*) plants are produced by a method as described by Olhoft *et al.* (US patent 2009/0049567). For transformation of soybean or *Arabidopsis thaliana*, Wildtype or Mutated PPO sequences based on one of the following sequences SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45, 47, are cloned with standard cloning techniques as described in Sambrook *et al.* (Molecular cloning (2001) Cold Spring Harbor Laboratory Press) in a binary vector containing resistance marker gene cassette (AHAS) and mutated PPO sequence (marked as GOI) in between ubiquitin promoter (PcUbi) and nopaline synthase terminator (NOS) sequence. For corn transformation, Wildtype or Mutated PPO sequences are cloned with standard cloning techniques as described in Sambrook *et al.* (Molecular cloning (2001) Cold Spring Harbor Laboratory Press) in a binary vector containing resistance marker gene cassette (AHAS) and mutated PPO sequence (marked as GOI) in between corn ubiquitin promoter (ZmUbi) and nopaline synthase terminator (NOS) sequence. Binary plasmids are introduced to *Agrobacterium tumefaciens* for plant transformation. Plasmid constructs are introduced into soybean's axillary meristem cells at the primary node of seedling explants via *Agrobacterium*-mediated transformation. After inoculation and co-cultivation with *Agrobacteria*, the explants are transferred to shoot introduction media without selection for one week. The explants were subsequently transferred to a shoot induction medium with 1-3 μ M imazapyr (Arsenal) for 3 weeks to select for transformed cells. Explants with healthy callus/shoot pads at the primary node are then transferred to shoot elongation medium containing 1-3 μ M imazapyr until a shoot elongated or the explant died. Transgenic plantlets are rooted, subjected to TaqMan analysis for the presence of the transgene, transferred to soil and grown to maturity in greenhouse.

Transformation of corn plants are done by a method described by McElver and Singh (WO 2008/124495). Plant transformation vector constructs containing mutated PPO sequences are introduced into maize immature embryos via *Agrobacterium*-mediated transformation.

Transformed cells were selected in selection media supplemented with 0.5-1.5 μ M imazethapyr for 3-4 weeks. Transgenic plantlets were regenerated on plant regeneration media and rooted afterwards. Transgenic plantlets are subjected to TaqMan analysis for the presence of the transgene before being transplanted to potting mixture and grown to maturity in greenhouse. *Arabidopsis thaliana* are transformed with wildtype or mutated PPO sequences by floral dip method as described by McElver and Singh (WO 2008/124495). Transgenic *Arabidopsis* plants were subjected to TaqMan analysis for analysis of the number of integration loci. Transformation of *Oryza sativa* (rice) are done by protoplast transformation as described by Peng *et al.* (US 6653529) T0 or T1 transgenic plant of soybean, corn, and rice containing mutated PPO sequences are tested for improved tolerance to PPO-derived herbicides in greenhouse studies and mini-plot studies with the following PPO-inhibiting herbicides: saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control.

Transgenic *Arabidopsis thaliana* plants were assayed for improved tolerance to saflufenacil, 1,5-

dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control in 48-well plates. Therefore, T2 seeds are surface sterilized by stirring for 5 min in ethanol + water (70+30 by volume), rinsing one time with ethanol + water (70+30 by volume) and two times with sterile, deionized water. The seeds are resuspended in 0.1% agar dissolved in water (w/v). Four to five seeds per well are plated on solid nutrient medium consisting of half-strength murashige skoog nutrient solution, pH 5.8 (Murashige and Skoog (1962) *Physiologia Plantarum* 15: 473-497). Compounds are dissolved in dimethylsulfoxid (DMSO) and added to the medium prior solidification (final DMSO concentration 0.1%). Multi well plates are incubated in a growth chamber at 22°C, 75% relative humidity and 110 $\mu\text{mol Phot} \cdot \text{m}^{-2} \cdot \text{s}^{-1}$ with 14 : 10 h light : dark photoperiod. Growth inhibition is evaluated seven to ten days after seeding in comparison to wild type plants.

Additionally, transgenic T1 *Arabidopsis* plants were tested for improved tolerance to PPO-inhibiting herbicides in greenhouse studies with the following PPO-inhibiting herbicides: saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control.

Results are shown in Table 5:

Table 5a: Germination Assay

Tolerance trails with: 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione			
Test Event	SEQ ID NO	Mutation	Tolerance Factor (non-transgenic Arabidopsis = 1)
1	4	R128A, F420V	300
2	4	R128A, F420V	300
3	4	R128A, F420V	3
4	4	R128A, F420V	300
5	4	R128A, F420V	300
6	4	R128A, F420V	200
7	4	R128A, F420V	3
8	4	R128A, F420V	300
9	4	R128A, F420V	300
10	4	R128A, F420V	300
11	4	R128A, F420V	40
12	4	R128A, F420V	3
13	4	R128A, F420V	300
14	4	R128A, F420V	3
15	4	R128A, F420V	200
16	4	R128A, F420V	200
17	4	R128A, F420V	300

18	4	R128A, F420V	3
19	4	R128A, F420V	75
20	4	R128A, F420V	200
21	4	R128A, F420V	300
22	4	R128A, F420V	3
23	4	R128A, F420V	8
24	4	R128A, F420V	75
25	4	R128A, F420V	200
26	4	R128A, F420V	300
1	4	F420V	75
2	4	F420V	75
3	4	F420V	35
4	4	F420V	75
5	4	F420V	300
6	4	F420V	300
7	4	F420V	300
8	4	F420V	300
9	4	F420V	300
10	4	F420V	300
11	4	F420V	3
12	4	F420V	8
13	4	F420V	300
14	4	F420V	20
15	4	F420V	300
16	4	F420V	300
17	4	F420V	300
18	4	F420V	35
19	4	F420V	3
20	4	F420V	300
21	4	F420V	300
22	4	F420V	300
23	4	F420V	300
24	4	F420V	300

Table 5b: Relative tolerance rates of transgenic Arabidopsis plants as compared to a non-transgenic Arabidopsis plant (**non-transgenic** = 1.0), treated with various PPO inhibitors. Growth inhibition is evaluated seven to ten days after seeding in comparison to wild type plants.

Mut PPO	Saflu- fenacil	1,5-dimethyl-6- thioxo-3-(2,2,7- trifluoro-3-oxo-4- (prop-2-ynyl)-3,4- dihydro-2H- benzo[b][1,4]oxazin-	Flumi- oxazin	Fome- safen	Lacto- fen	Sulfen- trazon
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		6-yl)-1,3,5-triazinane-2,4-dione				
AMATU_PPO2_wt	10	13	17	19	8	
AMATU_PPO2_dG210	100	33	107	29	19	203
AMATU_PPO2_R128L	160	23	126	27	22	186
AMATU_PPO2_dG210_R128L	1200	153	271	29	29	244
AMATU_PPO2_F420I	80	367	286	18	17	193
AMATU_PPO2_F420M	168	102	271	29	29	161
AMATU_PPO2_F420L	192	253	286	23	19	111
AMATU_PPO2_R128A_F420I	1200	333	286	29	27	621
AMATU_PPO2_R128A_F420L	1200	333	286	29	29	717
AMATU_PPO2_R128A_F420M	1160	204	286	29	29	

Table 5 c: Phytotox values of transgenic Arabidopsis plants as compared to a non-transgenic Arabidopsis plant (**non-transgenic = 100% damage**), treated with 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione.

Line	Assesment DAT (DAT = Days After Treatment)	SEQ_ID	Substitution	Injury Rating 0 - 100%		
				(0 = no injury, 100 = total control)		
				300	150	75
				1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione g/Ha + 1%MSO		
1	7	2 & 4	R128A_F420V	40	95	95
1	7	2 & 4	R128A_F420V	100	25	0
1	7	2 & 4	R128A_F420V	25	35	35
1	19	2 & 4	R128A_F420V	28	90	90
1	19	2 & 4	R128A_F420V	100	60	25
1	19	2 & 4	R128A_F420V	25	30	30
2	7	2 & 4	F420V	98	95	95
2	7	2 & 4	F420V	25	90	15
2	7	2 & 4	F420V	25	15	15

2	19	2 & 4	F420V	95	90	98
2	19	2 & 4	F420V	55	85	40
2	19	2 & 4	F420V	45	45	30

Table 5 d: Relative tolerance rates of transgenic Arabidopsis plants as compared to a non-transgenic Arabidopsis plant **on a scale from 0 – 100, where 100 is 100% damage**, treated with single and mixtures of PPO inhibitors (e.g. Saflufenacil plus 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione). Plant growth injury is evaluated seven to ten days after application in comparison to wild type plants.

			non-transgenic Arabidopsis	non-transgenic Arabidopsis	non-transgenic Arabidopsis	Ø non-transgenic Arabidopsis	Ø R128A, F420V 1	Ø R128A, F420V 2	Ø R128A, F420V 3	Ø F420V 1	Ø F420V 2	Ø F420V 3	Ø R128A, F420V 1 to 3	Ø F420V 1 to 3
			POST	POST	POST	POST	POST	POST	POST	POST	POST	POST	Ø R128A, F420V 1 to 3	Ø F420V 1 to 3
PPO Herbicide (+ 1% MSO)	g ai/ha	DAT	7	7	7	7	7	7	7	7	7	7		
Saflufenacil + 1,5-dimethyl-6- thioxo-3-(2,2,7- trifluoro-3-oxo-4- (prop-2-ynyl)-3,4- dihydro-2H- benzo[b][1,4]oxa- zin-6-yl)-1,3,5- triazinane-2,4- dione	50 + 25		98	98	98	98	23	23	21	33	33	27	22	31
	25 + 50		98	98	98	98	16	19	16	27	22	16	17	22
	100 + 50		98	98	98	98	15	26	23	55	47	43	21	48
	50 + 100		98	98	98	98	10	20	28	35	33	31	19	33
	200 + 100		98	98	98	98	25	23	28	63	60	66	25	63
	100 + 200		98	98	98	98	30	29	26	58	45	56	28	53
Saflufenacil	75		98	98	98	98	16	22	18	39	36	51	18	42
	150		98	98	98	98	18	24	18	60	55	66	20	60
	300		98	98	98	98	22	22	19	77	72	78	21	76
1,5-dimethyl-6- thioxo-3-(2,2,7- trifluoro-3-oxo-4- (prop-2-ynyl)-3,4-	75		98	98	98	98	18	24	11	17	9	8	18	11
	150		98	98	98	98	23	20	30	28	11	12	24	17
	300		98	98	98	98	26	33	36	36	22	22	32	26

dihydro-2H- benzo[b][1,4]oxa- zin-6-yl)-1,3,5- triazine-2,4- dione													
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Table 5 e shows phytotox values on a scale from 0 – 100, were 100 is 100% damage.

		ARBTH WT	AMATU_PPO2_R128A_F420V	AMATU_PPO2_R128A_F420V	AMATU_PPO2_R128A_F420V	AMATU_PPO2_L397D_F420V	AMATU_PPO2_L397D_F420V
	event	1	A	B	D	O	P
compound	g ai/ha						
KIXOR +	75 + 400 + 3750	100	8	0	20	0	7
VALOR (Flumioxazin) +	50 + 200 + 3750	100	0	0	12	0	7
DESTINY HC	25 + 100 + 3750	100	0	17	12	0	3
KIXOR +	75 + 120 + 3750	100	5	3	13	15	22
SPOTLIGHT (Carfentrazone) +	50 + 60 + 3750	100	0	3	3	5	7
DESTINY HC	25 + 30 + 3750	100	0	7	3	3	3
KIXOR +	75 + 200 + 3750	100	3	8	22	13	15
BAS 850 00 H +	50 + 100 + 3750	100	0	7	13	10	10
DESTINY HC	25 + 50 + 3750	100	0	15	15	7	7
BAS 850 00 H +	200 + 400 + 3750	100	10	12	20	17	17
VALOR (Flumioxazin) +	100 + 200 + 3750	100	2	7	13	10	10
DESTINY HC	50 + 100 + 3750	100	0	0	3	3	0
BAS 850 00 H +	200 + 120 + 3750	100	8	20	23	17	20

SPOTLIGHT (Carfentrazone) +	100 + 60 + 3750						
		100	3	12	7	8	7
DESTINY HC	50 + 30 + 3750	100	0	7	7	0	3

Table 5f shows phytotox values on a scale from 0 – 100, were 100 is 100% damage

		ARBTH WT	AMATU_PPO2 F420V				AMATU_PPO2 R128A_F420V				AMATU_PPO2 L397D				AMATU_PPO2 L397D_F420V			
	repetition	1	1	2	1	2	1	2	1	2	1	2	1	2	1	2	1	2
compound	g ai/ha																	
Kixor	200	100	85	90	95	80	10	10	40	10	95	95	85	90	30	0	10	10
	100	100	65	70	70	65	10	0	10	10	85	85	80	80	10	0	20	10
	50	100	50	30	50	50	0	0	10	10	65	65	50	70	10	20	10	40
BAS 850H	300	100	70	50	40	50	20	30	20	30	90	100	70	85	10	20	50	10
	150	100	60	40	40	65	10	10	40	50	75	70	70	70	20	10	30	0
	75	100	30	40	30	40	0	0	10	20	70	80	60	65	10	10	40	10
Carfentrazone	200	100	40	10	50	20	30	40	10	10	65	60	50	65	20	20	20	10
	100	100	10	10	40	20	10	10	10	10	60	50	30	30	20	20	50	10
	50	100	10	10	40	10	10	10	30	0	30	60	20	30	30	10	50	20
Kixor + Carfentrazone	75 +120	100	40	70	75	65	10	10	10	10	90	80	55	65	40	30	10	10
	37,5 + 60	100	30	65	70	50	10	30	0	0	70	80	55	50	10	10	10	10
	18,75 + 30	100	30	30	30	30	10	30	30	0	60	70	10	20	10	10	75	20

Table 5g shows phytotox values on a scale from 0 – 100, were 100 is 100% damage

compound	repetition g ai/ha	wild type	AMATU_PPO2_F420M		AMATU_PPO2_R128A_F420M		AMATU_PPO2_R128A_F420V		AMATU_PPO2_L397D_F420V		AMATU_PPO2_L397D	
	Event		A	B	A	B	A	D	O	A	E	O
Oxyfluorfen Kixor MSO 1%	800 + 75 + 3750	100	70	73	15	5	75	55	7,5	75	78	73
	800 + 50 + 3750	100	65	63	18	10	50	53	23	83	78	68
	800 + 25 + 3750	100	65	58	13	13	63	43	5	83	68	53
Oxyfluorfen Flumioxazin MSO 1%	800 + 400 + 3750	100	60	60	13	20	63	60	20	83	63	43
	800 + 200 + 3750	100	65	55	25	23	73	43	35	80	60	38
	800 + 100 + 3750	100	63	53	40	35	70	40	5	85	50	38
Oxyfluorfen BAS 850H MSO 1%	800 + 200 + 3750	100	75	70	60	58	70	60	20	90	95	83
	800 + 100 + 3750	100	73	65	63	50	75	55	13	93	100	78
	800 + 50 + 3750	100	73	50	43	50	73	60	25	88	88	70
Fomesafen BAS 850H MSO 1%	300 + 200 + 3750	100	85	85	63	55	80	78	60	97	90	73
	300 + 100 + 3750	100	85	85	58	55	85	78	70	95	93	83
	300 + 50 + 3750	100	93	83	48	55	85	80	63	94	90	75
Oxyfluorfen	800 + 600 +	100	85	95	60	50	90	83	58	93	68	40

Fomesafen	3750											
MSO 1%	800 + 450 + 3750	100	88	85	58	48	80	80	50	94	58	35
	800 + 300 + 3750	100	80	80	60	43	80	80	65	97	58	45
Flumioxazin	100 + 120 + 3750	100	68	70	58	55	45	28	0	78	80	60
	100 + 60 + 3750	100	60	60	50	43	40	45	0	83	73	60
Carfentrazone	100 + 30 + 3750	100	65	60	45	43	53	43	5	97	70	60
MSO 1%												
Oxyfluorfen	800 + 120 + 3750	100	45	43	43	35	65	68	25	88	68	53
	800 + 60 + 3750	100	38	25	10	33	58	60	35	88	58	53
Carfentrazone	800 + 30 + 3750	100	38	18	10	25	65	58	30	95	55	30
MSO 1%												

EXAMPLE 6: Tissue Culture Conditions.

An *in vitro* tissue culture mutagenesis assay has been developed to isolate and characterize plant tissue (e.g., maize, rice tissue) that is tolerant to protoporphyrinogen oxidase inhibiting herbicides, (saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenoxy, sulfentrazone, and photosynthesis inhibitor diuron as negative control). The assay utilizes the somaclonal variation that is found in *in vitro* tissue culture. Spontaneous mutations derived from somaclonal variation can be enhanced by chemical mutagenesis and subsequent selection in a stepwise manner, on increasing concentrations of herbicide.

The present invention provides tissue culture conditions for encouraging growth of friable, embryogenic maize or rice callus that is regenerable. Calli were initiated from 4 different maize or rice cultivars encompassing *Zea mays* and Japonica (Taipei 309, Nipponbare, Koshihikari) and Indica (Indica 1) varieties, respectively. Seeds were surface sterilized in 70% ethanol for approximately 1 min followed by 20% commercial Clorox bleach for 20 minutes. Seeds were rinsed with sterile water and plated on callus induction media. Various callus induction media were tested. The ingredient lists for the media tested are presented in Table 6.

Table 6

Ingredient	Supplier	R001M	R025M	R026M	R327M	R008M	MS711R
B5 Vitamins	Sigma					1.0 X	
MS salts	Sigma			1.0 X	1.0 X	1.0 X	1.0 X
MS Vitamins	Sigma			1.0 X	1.0 X		
N6 salts	Phytotech	4.0 g/L	4.0g/L				
N6 vitamins	Phytotech	1.0 X	1.0 X				
L-Proline	Sigma	2.9 g/L	0.5 g/L				1.2 g/L
Casamino Acids	BD	0.3 g/L	0.3 g/L	2 g/L			
Casein Hydrolysate	Sigma						1.0 g/L
L-Asp Monohydrate	Phytotech						150 mg/L
Nicotinic Acid	Sigma						0.5 mg/L
Pyridoxine HCl	Sigma						0.5 mg/L
Thiamine HCl	Sigma						1.0 mg/L
Myo-inositol	Sigma						100 mg/L
MES	Sigma	500 mg/L	500 mg/L	500 mg/L	500 mg/L	500 mg/L	500 mg/L
Maltose	VWR	30 g/L	30 g/L	30 g/L	30 g/L		
Sorbitol	Duchefa			30 g/L			
Sucrose	VWR					10 g/L	30 g/L
NAA	Duchefa					50 µg/L	
2,4-D	Sigma	2.0 mg/L					1.0 mg/L
MgCl ₂ ·6H ₂ O	VWR					750 mg/L	
→pH		5.8	5.8	5.8	5.8	5.8	5.7
Gelrite	Duchefa	4.0 g/L				2.5 g/L	
Agarose Type1	Sigma		7.0 g/L	10 g/L	10 g/L		
→Autoclave		15 min	15 min	15 min	15 min	15 min	20 min
Kinetin	Sigma		2.0 mg/L	2.0 mg/L			
NAA	Duchefa		1.0 mg/L	1.0 mg/L			
ABA	Sigma		5.0 mg/L				
Cefotaxime	Duchefa		0.1 g/L	0.1 g/L	0.1 g/L		
Vancomycin	Duchefa		0.1 g/L	0.1 g/L	0.1 g/L		
G418 Disulfate	Sigma		20 mg/L	20 mg/L	20 mg/L		

R001M callus induction media was selected after testing numerous variations. Cultures were kept in the dark at 30°C. Embryogenic callus was subcultured to fresh media after 10-14 days.

5

EXAMPLE 7: Selection of Herbicide-tolerant Calli.

Once tissue culture conditions were determined, further establishment of selection conditions were established through the analysis of tissue survival in kill curves with saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control. Careful consideration of

10

accumulation of the herbicide in the tissue, as well as its persistence and stability in the cells and the culture media was performed. Through these experiments, a sub-lethal dose has been established for the initial selection of mutated material. After the establishment of the starting dose of saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control in selection media, the tissues were selected in a step-wise fashion by increasing the concentration of the PPO inhibitor with each transfer until cells are recovered that grew vigorously in the presence of toxic doses. The resulting calli were further subcultured every 3-4 weeks to R001M with selective agent. Over 26,000 calli were subjected to selection for 4-5 subcultures until the selective pressure was above toxic levels as determined by kill curves and observations of continued culture. Alternatively, liquid cultures initiated from calli in MS711R with slow shaking and weekly subcultures. Once liquid cultures were established, selection agent was added directly to the flask at each subculture. Following 2-4 rounds of liquid selection, cultures were transferred to filters on solid R001M media for further growth.

EXAMPLE 8: Regeneration of Plants.

Tolerant tissue was regenerated and characterized molecularly for PPO gene sequence mutations and/or biochemically for altered PPO activity in the presence of the selective agent. In addition, genes involved directly and/or indirectly in tetrapyrrole biosynthesis and/or metabolism pathways were also sequenced to characterize mutations. Finally, enzymes that change the fate (e.g. metabolism, translocation, transportaion) were also sequence to characterized mutations. Following herbicide selection, calli were regenerated using a media regime of R025M for 10 – 14 days, R026M for ca. 2 weeks, R327M until well formed shoots were developed, and R008S until shoots were well rooted for transfer to the greenhouse. Regeneration was carried out in the light. No selection agent was included during regeneration. Once strong roots were established, M0 regenerants were transplant to the greenhouse in square or round pots. Transplants were maintained under a clear plastic cup until they were adapted to greenhouse conditions. The greenhouse was set to a day/night cycle of 27°C/21°C (80°F/70°F) with 600W high pressure sodium lights supplementing light to maintain a 14 hour day length. Plants were watered according to need, depending in the weather and fertilized daily.

EXAMPLE 9: Sequence Analysis.

Leaf tissue was collected from clonal plants separated for transplanting and analyzed as individuals. Genomic DNA was extracted using a Wizard® 96 Magnetic DNA Plant System kit (Promega, US Patent Nos. 6,027,945 & 6,368,800) as directed by the manufacturer. Isolated DNA was PCR amplified using the appropriate forward and reverse primer.

PCR amplification was performed using Hotstar Taq DNA Polymerase (Qiagen) using touchdown thermocycling program as follows: 96°C for 15 min, followed by 35 cycles (96°C, 30 sec; 58°C – 0.2 °C per cycle, 30 sec; 72°C, 3 min and 30 sec), 10 min at 72°C. PCR products were verified for concentration and fragment size via agarose gel electrophoresis. Dephosphorylated PCR products were analyzed by direct sequence using the PCR primers (DNA Landmarks, or Entelechon). Chromatogram trace files (.scf) were analyzed for mutation relative to the wild-type gene using

Vector NTI Advance 10™ (Invitrogen). Based on sequence information, mutations were identified in several individuals. Sequence analysis was performed on the representative chromatograms and corresponding AlignX alignment with default settings and edited to call secondary peaks.

5 **EXAMPLE 10: Demonstration of Herbicide-tolerance.**

T0 or T1 transgenic plant of soybean, corn, Canola varieties and rice containing PPO1 and or PPO2 sequences are tested for improved tolerance to herbicides in greenhouse studies and mini-plot studies with the following herbicides: saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS
10 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control. For the pre-emergence treatment, the herbicides are applied directly after sowing by means of finely distributing nozzles. The containers are irrigated gently to promote germination and growth and subsequently covered with transparent plastic hoods until the plants have rooted. This cover causes uniform germination of the test plants,
15 unless this has been impaired by the herbicides. For post emergence treatment, the test plants are first grown to a height of 3 to 15 cm, depending on the plant habit, and only then treated with the herbicides. For this purpose, the test plants are either sown directly, and grown in the same containers or they are first grown separately and transplanted into the test containers a few days prior to treatment.

20

For testing of T0 plants, cuttings can be used. In the case of soybean plants, an optimal shoot for cutting is about 7.5 to 10 cm tall, with at least two nodes present. Each cutting is taken from the original transformant (mother plant) and dipped into rooting hormone powder (indole-3-butyric acid, IBA). The cutting is then placed in oasis wedges inside a bio-dome. Wild type cuttings are also
25 taken simultaneously to serve as controls. The cuttings are kept in the bio-dome for 5-7 days and then transplanted to pots and then acclimated in the growth chamber for two more days. Subsequently, the cuttings are transferred to the greenhouse, acclimated for approximately 4 days, and then subjected to spray tests as indicated. Depending on the species, the plants are kept at 10-25°C or 20-35°C. The test period extends over 3 weeks. During this time, the plants are tended
30 and their response to the individual treatments is evaluated. Herbicide injury evaluations are taken at 2 and 3 weeks after treatment. Plant injury is rated on a scale of 0% to 100%, 0% being no injury and 100% being complete death.

Transgenic *Arabidopsis thaliana* plants were assayed for improved tolerance to saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control, in 48-well plates.
35 Therefore, T2 seeds are surface sterilized by stirring for 5 min in ethanol + water (70+30 by volume), rinsing one time with ethanol + water (70+30 by volume) and two times with sterile, deionized water. The seeds are resuspended in 0.1% agar dissolved in water (w/v) Four to five seeds per well are plated on solid nutrient medium consisting of half-strength murashige skoog nutrient solution, pH 5.8 (Murashige and Skoog (1962) *Physiologia Plantarum* 15: 473-497). Compounds are dissolved in dimethylsulfoxid (DMSO) and added to the medium prior solidification (final DMSO concentration 0.1%). Multi well plates are incubated in a growth chamber at 22°C,
40

75% relative humidity and $110 \mu\text{mol Phot} \cdot \text{m}^{-2} \cdot \text{s}^{-1}$ with 14 : 10 h light : dark photoperiod. Growth inhibition is evaluated seven to ten days after seeding in comparison to wild type plants.

Additionally, transgenic T1 *Arabidopsis* plants were tested for improved tolerance to herbicides in greenhouse studies with the following herbicides: saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control. Results are shown in Table 5 and Figure 2.

EXAMPLE 11: Herbicide Selection Using Tissue Culture.

Media was selected for use and kill curves developed as specified above. For selection, different techniques were utilized. Either a step wise selection was applied, or an immediate lethal level of herbicide was applied. In either case, all of the calli were transferred for each new round of selection. Selection was 4-5 cycles of culture with 3-5 weeks for each cycle. Calli were placed onto nylon membranes to facilitate transfer (200 micron pore sheets, Biodesign, Saco, Maine).

Membranes were cut to fit 100x20 mm Petri dishes and were autoclaved prior to use. 25-35 calli (average weight/calli being 22mg) were utilized in every plate. In addition, one set of calli were subjected to selection in liquid culture media with weekly subcultures followed by further selection on semi-solid media. Mutant lines were selected using saflufenacil, 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4), flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control. Efficiencies of obtaining mutants was high either based on a percentage of calli that gave rise to a regenerable, mutant line or the number of lines as determined by the gram of tissue utilized.

EXAMPLE 12: Maize whole plant transformation and PPO inhibitor tolerance testing.

Immature embryos were transformed according to the procedure outlined in Peng et al. (WO2006/136596). Plants were tested for the presence of the T-DNA by Taqman analysis with the target being the nos terminator which is present in all constructs. Healthy looking plants were sent to the greenhouse for hardening and subsequent spray testing. The plants were individually transplanted into MetroMix 360 soil in 4" pots. Once in the greenhouse (day/night cycle of 27°C /21°C with 14 hour day length supported by 600W high pressure sodium lights), they were allowed to grow for 14 days. They were then sprayed with a treatment of 25 to 200 g ai/ha saflufenacil + 1.0% v/v methylated seed oil (MSO) and / or 25 - 200 g ai/ha 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (CAS 1258836-72-4) plus 1% MSO. Other PPO inhibiting herbicides were also tested in a similar fashion for confirming cross resistance: flumioxazin, butafenacil, acifluorfen, lactofen, bifenox, sulfentrazone, and photosynthesis inhibitor diuron as negative control. Herbicide injury evaluations were taken at 7, 14 and 21 days after treatment. Herbicide injury evaluations were taken 2, 7, 14 and 21 days post-spray to look for injury to new growth points and overall plant health. The top survivors were transplanted into gallon pots filled with MetroMix 360 for seed production. Results are shown in Table 7 and Figures 3, and 4.

Table 7a Transgenic T0 corn events were sprayed in the greenhouse with the indicated amount of compound + 1% (v/v) MSO at V2 stage. Herbicide injury was evaluated 7 days after treatment with

a 0 to 9 rating scale where 0 is no injury relative to an unsprayed wild type check and 9 is completely dead.

Table 7a

SEQ ID	Event	0	BAS800H (g ai/ha)		BAS850H (g ai/ha)		
			50	75	50	75	100
AmtuPPX2L_R128A_F420V	1	0					
	2	0					
	3	0					
	4	0					
	5	0					
	6	0					
	7	0					
	8	0					
	9				4		
	10				4		
	11				4		
	12				4		
	13				3		
	14				4		
	15				4		
	16				3		
	17						4
	18						4
AmtuPPX2L_R128A_F420I	1	1					
	2	1					
	3	1					
	4	0					
	5	0					
	6	2					
	7	0					
	8	1					
	9	1					
	10	1					
	11		8				
	12		1				
	13		4				
	14		1				
	15		0				
	16				6		
	17				0		
	18				2		
	19				2		

	20			1		
	21				5	
	22				1	
	23	0				
	24	0				
	25	0				
	26	0				
	27	0				
	28		0			
	29		0			
	30			0		
	31			1		
	32			0		
	33			0		
	34			3		
	35				1	
	36	0				
	37	0				
	38	0				
	39				4	
	40			0		
	41			2		
	42			1		
	43					4
AmtuPPX2L_R128A _F420L	1	0				
	2			3		
	3					2
	4	0				
	5			2		
	6					2
	7	0				
	8			2		
	9	0				
	10			2		
	11	0				
	12			3		
	13	0				
	14			3		
	15	0				
	16			2		
	17	0				
	18					2
	19	0				
	20					2
	21	0				

	22	0					
	23	0					
	24	0					
	25	2					
AmtuPPX2L_R128A _F420V	1	0					
	2				1		
	3						1
	4	0					
	5				4		
	6						5
	7	0					
	8				3		
	9						1
	10	0					
	11				6		
	12	0					
	13				3		
	14	0					
	15				1		
	16	0					
	17				3		
	18	0					
	19						1
	20	0					
	21						5
	22	0					
	23						1
	24	0					
	25	3					
	26	1					
	27	1					
Tp-Fdx::c- AmtuPPX2L_R128A _F420V	1	0					
	2	0					
	3	0					
	4				0		
	5				1		
	6				0		
	7	0					
	8				0		
	9						0
	10	0					
	11				0		
	12	0					

	13				0		
	14						1
	15			2			
AmtuPPX2L_R128L_F420M	1	0			1		
	2				1		
	3				0		
	4				5		
	5				1		
	6				5		
	7				3		
	8				2		
	9				8		
	10				2		
	11				2		
	12				0		
	13	0			0		
	14	0			2		
	15				0		
	16				0		
	17				3		
	18				3		
	19				6		
	20				1		
	21				4		
	22				3		
	23				2		
	24				2		
	25				0		
	26				0		
	27				0		
	28				2		
	29				2		
	30				1		
	31				0		
	32				2		
	33				2		
	34				1		
	35				4		
	36				1		
	37				2		
AmtuPPX2L_R128M_F420I	1	0			7		
	2	0			0		
	3	0		0	1		0
	4				1		

	5			1		
	6			0		
	7			2		
	8	0		1	0	
	9	0		0	1	
	10	0		0		
	11	0		1		
	12	0		1		
	13	0		4		
	14	0		0		
	15	0		1		
	16	0		1		
	17			2		
	18			4		
	19			2		
	20			0		
	21			0		
	22			0		
	23			0		
	24			1		
	25			4		
	26			0		
	27			0		
	28			0		
	29			2		
	30			3		
	31	0		3		
	32	0		1	2	
	33			4		
	34	0		3		
	35	0		1		2
	36			4		
	37			1		
AmtuPPX2L_R128 M_F420L	1	1		1		
	2			0		
	3			4		
	4			0		
	5	0		1	2	
	6	0		0		
	7			0		
	8			1		
	9			6		
	10			0		
	11			0		
	12			0		

	13	0			1		
	14	0			3		
	15				2		
	16	0			1		
	17	0			3		
	18				0		
AmtuPPX2L_R128 M_F420V	1	0			0		
	2	0			3		
	3	0			0		
	4	0			0		
	5	0		1	0		0
	6	0			5		
	7				6		
	8				1		
	9				5		
	10				1		
	11				0		
	12				0		
	13				0		
	14	2			0		
	15	0			0		
	16	1			1		
	17	0			0		1
	18				1		
	19				0		
	20				1		
	21				0		
	22				1		
	23				0		
	24				0		
	25				0		
	26	2			0		
	27	0			0		
	28	1			1		
	29	0			0		1
	30				1		
	31				0		
	32				1		
	33				0		
	34				1		
	35				0		
	36				0		
	37				0		
	38				2		
	39				0		

	40				1		
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Table 7 b Transgenic T1 corn events were sprayed in the field with 100 g ai BAS800H and 50 g ai BAS850H + 1% (v/v) MSO at V2-V3 developmental stage. Herbicide injury was evaluated at 3, 7, 14, and 21 days after treatment (DAT) with a 0 to 100 rating scale where 0 is no injury relative to an unsprayed wild type check and 100 is completely dead

5

Construct	SEQ ID	Event	3 DAT	7 DAT	14 DAT	21 DAT
RTP11136-1	AmtuPPX2L_R128A_F420V	1	20	30	0	0
RTP11141-1	AmtuPPX2L_R128A_F420I	2	70	80	70	80
RTP11141-1		3	20	10	10	10
RTP11141-1		4	10	0	30	20
RTP11141-1		5	10	0	20	10
RTP11141-1		6	10	0	10	0
RTP11141-1		7	10	0	30	20
RTP11141-1		8	80	80	70	70
RTP11141-1		9	10	0	10	0
RTP11141-1		10	10	10	40	30
RTP11141-1		11	10	10	30	20
RTP11142-2	AmtuPPX2L_R128A_F420L	12	10	30	10	10
RTP11142-2		13	10	10	30	20
RTP11142-2		14	10	10	20	20
RTP11142-2		15	10	10	30	20
RTP11142-2		16	20	30	40	20
RTP11142-2		17	10	0	20	0
RTP11142-2		18	10	10	10	0
RTP11142-2		19	20	10	10	0
RTP11142-2		20	10	10	10	0
RTP11142-2		21	10	10	10	0
RTP11142-2		22	10	0	10	0
RTP11142-2		23	20	40	50	50
RTP11142-2		24	50	80		
RTP11142-2		25	10	10	0	0
RTP11142-2		26	0	10	10	0
RTP11142-2		27	10	20	20	0
RTP11142-2		28	10	20	20	10
RTP11142-2		29	10	20	30	10
RTP11142-2		30	10	40	40	20
RTP11142-2		31	0	30	40	20
RTP11143-2	AmtuPPX2L_R128A_F420V	32	10	40	40	20
RTP11143-2		33	10	30	30	10
RTP11143-2		34	10	20	20	10
RTP11143-2		35	10	40	40	20
RTP11143-2		36	10	20	10	0

RTP11144-2	Tp-Fdx::c- AmtuPPX2L_R128A_F420V	37	20	10	10	0
RTP11144-2		38	20	10	10	0
RTP11144-2		39	0	0	10	0
RTP11144-2		40	30	20	20	0
RTP11144-2		41	40	10	10	0
RTP11144-2		42	20	10	0	0
RTP11144-2		43	0	10	0	0
RTP11144-2		44	30	10	10	0
RTP11144-2		45	20	20	0	0

EXAMPLE 13: Soybean transformation and PPO Inhibitor tolerance testing.

Soybean cv Jake was transformed as previously described by Siminszky et al., Phytochem Rev.

5:445-458 (2006). After regeneration, transformants were transplanted to soil in small pots, placed
 5 in growth chambers (16 hr day/ 8 hr night; 25°C day/ 23°C night; 65% relative humidity; 130-150
 microE m⁻² s⁻¹) and subsequently tested for the presence of the T-DNA via Taqman analysis.

After a few weeks, healthy, transgenic positive, single copy events were transplanted to larger pots
 and allowed to grow in the growth chamber. An optimal shoot for cutting was about 3-4 inches tall,

10 with at least two nodes present. Each cutting was taken from the original transformant (mother
 plant) and dipped into rooting hormone powder (indole-3-butyric acid, IBA). The cutting was then
 placed in oasis wedges inside a bio-dome. The mother plant was taken to maturity in the

greenhouse and harvested for seed. Wild type cuttings were also taken simultaneously to serve as
 negative controls. The cuttings were kept in the bio-dome for 5-7 days and then transplanted to 3
 inch pots and then acclimated in the growth chamber for two more days. Subsequently, the

15 cuttings were transferred to the greenhouse, acclimated for approximately 4 days, and then
 sprayed with a treatment of 0 - 200 g ai/ha saflufenacil plus 1% MSO and / or 25 - 200 g ai/ha 1,5-
 dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-
 1,3,5-triazinane-2,4-dione (CAS 1258836-72-4) plus 1% MSO. Other PPO inhibiting herbicides

were also tested in a similar fashion for confirming cross resistance: flumioxazin, butafenacil,

20 acifluorfen, lactofen, bifenoxy, sulfentrazone, and photosynthesis inhibitor diuron as negative
 control. Herbicide injury evaluations were taken at 2, 7, 14 and 21 days after treatment. Results
 are shown in Table 8, and Figures 5, 6, and 7.

Table 8 b. Greenhouse data - segregating T1 individuals. Rated for injury (0-9 point scale) 1 week after treatment

GOI	wild type	AmtuPPX2L R128A_F420L	AmtuPPX2L R128A_F420L	AmtuPPX2L R128A_F420L	AmtuPPX2L R128A_F420V	AmtuPPX2L L397D_F420V	AmtuPPX2L R128A_F420M	AmtuPPX2L R128A_F420I	AmtuPPX2L R128A_F420I
Event		SDS-10642	SDS-10787	SDS-11034	SDS-10652	SDS-10990	SDS-10985	SDS-10791	SDS-10648
unsprayed	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	1	1	1
	1	0	*	*	0	0	0	0	0
Saflufenacil 150 g ai/ha	9	0	6	0	3	9	1	1	5
	9	0	5	0	5	3	0	0	5
	9	0	4	5	3	0	*	9	4
	9	0	0	1	4	1	3	9	4
	9	0	4	0	6	3	2	9	6
	9	1	4	0	4	3	0	9	5
1,5-dimethyl-6- thioxo-3-(2,2,7- trifluoro-3-oxo-4- (prop-2-ynyl)-3,4- dihydro-2H- benzo[b][1,4]oxazin- 6-yl)-1,3,5- triazine-2,4-dione 100 g ai/ha	9	1	4	0	3	3	0	1	3
	9	0	9	2	5	4	1	2	4
9	6	4	4	4	9	9	9	3	4
	5	5	5	4	4	7	9	2	9
	5	9	9	4	4	6	6	4	4

	9	5	9	3	4	6	9	9	5
	9	5	5	2	4	9	6	3	5
	8	9	5	3	9	7	9	5	5
	9	5	6	3	4	6	9	4	4
	9	4	6	2	4	9	6	4	5
Fomesafen 600 g ai/ha	5	0	1	2	1	1	3	6	5
	5	1	1	0	2	3	0	3	3
	4	0	0	0	0	0	0	3	3
	4	1	0	2	0	4	1	1	3
	4	0	2	0	1	5	3	4	3
	5	1	5	1	0	5	3	2	3
	4	1	2	1	2	0	1	1	3
	5	0	3	1	4	1	5	1	4
	9	3	9	5	9	6	9	3	9
	9	3	5	4	6	5	6	3	9
Flumioxazin 150 g ai/ha	9	2	4	6	6	6	4	3	5
	9	1	5	5	5	5	5	1	9
	9	3	5	9	5	6	9	1	5
	9	9	9	3	4	6	6	3	9
	9	1	4	6	4	9	4	1	5
	9	2	5	5	6	6	9	3	9
	9	1	5	1	9	3	3	9	3
	9	0	5	3	*	3	3	9	4
	7	3	4	3	6	4	8	9	3
	7	1	6	9	3	9	3	2	4
Sulfentrazone 350 g ai/ha	8	2	9	0	5	4	*	1	5
	9	0	9	1	3	3	4	2	5
	9	0	5	1	5	9	3	9	3
	9	3	5	1	6	3	1	9	8

Sulfentrazone 700 g ai/ha	9	3	3	3	2	3	1	3	3	3
	9	1	4	3	3	3	9	3	3	2
	9	3	6	3	7	9	2	3	3	9
	9	2	4	2	7	4	3	4	3	3
	9	2	5	1	4	4	4	9	4	4
	9	2	6	3	4	4	3	2	4	4
	9	0	5	4	6	9	2	9	4	4
	9	2	6	2	4	9	9	9	4	4
	8	2	6	4	4	4	1	4	4	5
	7	4	*	9	4	8	3	8	7	7
Oxyfluorfen 600 g ai/ha	8	3	5	5	5	4	4	4	6	6
	9	2	8	4	6	3	3	5	8	8
	7	8	5	4	6	4	4	5	5	6
	8	3	6	5	9	9	3	5	8	8
	9	2	6	5	4	4	4	4	9	9
	7	3	5	6	4	5	9	3	3	3
	9	3	6	5	5	9	5	9	5	5
	9	4	6	6	5	5	4	4	9	9
	9	3	5	6	4	4	4	5	9	9
	9	3	8	6	4	5	8	5	4	4
Oxyfluorfen 1200 g ai/ha	8	2	5	5	3	5	8	5	5	5
	9	4	5	6	4	5	5	9	9	9
	9	3	9	6	4	4	4	5	5	9
	9	3	9	6	4	4	4	5	5	9
	8	2	5	5	3	5	8	5	5	5
	9	4	5	6	4	5	5	9	9	9
	9	3	9	6	4	4	4	5	5	9
	8	3	5	5	5	5	5	4	4	5

Table 8 c: Field data - T1 generation. Rated for injury (1-5 point scale) 3 days after treatment.

GOI	wild type	AmtuPPX2L R128A_F420M	AmtuPPX2L R128A_F420I	AmtuPPX2L R128A_F420I	AmtuPPX2L R128A_F420I	AmtuPPX2L R128A_F420V	AmtuPPX2L L397D_F420V
Event		SDS-11052	SDS-10648	SDS-10791	SDS-11014	SDS-11035	SDS-11034
unsprayed	1	1	1	1	1	1	1
1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzof[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (=„benzoxazin“; BAS 850H) 100 g ai/ha	5	3	3	2	2	2	3
benzoxazin 50 g ai/ha	5	3	3	2	2	2	2
Saflufenacil 150 g ai/ha	5	2	2	2	2	2	2
Saflufenacil 75 g ai/ha	5	2	2	2	2	2	2

Rating	Phenotype (phytotoxicity) of surviving plants
1	no obvious damage (no phytotoxicity)
2	minor amount of leaf damage, plant will survive
3	moderate amount of leaf damage, plant will survive
4	severe amount of leaf damage, plant will survive
5	no surviving plants - all plants dead/dying

Table 8d: Field data - T1 generation soybeans rated for injury with 1-5 point scale.										
Injury rating taken 3 days after treatment										
Genotype	GOI	Event	benzoxazin +Saflufenacil (100gai/ha + 100gai/ha)	benzoxazin + Saflufenacil (50gai/ha + 50 gai/ha)	benzoxazi n (100 gai/ha)	benzoxazin (50 gai/ha)	Saflufenacil (150 gai/ha)	Saflufenacil (75 gai/ha)		
			Rating							
Wildtype		Jake	5	5	5	5	5	5	5	5
LTM377-1	AmtuPPX2L_dG210	SDS-10656	4	4	4	4	4	3,5	3,5	3,5
LTM377-1	AmtuPPX2L_dG210	SDS-10562	*	*	3	3	4	4	4	4
LTM377-1	AmtuPPX2L_dG210	SDS-10566	*	*	3	3	4	4	4	4
LTM387-1	AmtuPPX2L_R128A_ F420V	SDS-11034	*	*	2	2	2	2	3	3
LTM387-1	AmtuPPX2L_R128A_ F420V	SDS-11035	*	*	2	2	2	2	2	2
LTM387-1	AmtuPPX2L_R128A_ F420V	SDS-10998	2,5	2,5	2,5	2,5	2	2	2	2
LTM387-1	AmtuPPX2L_R128A_ F420V	SDS-11105	3,5	3	3	3	2,5	2,5	2,5	2,5
LTM387-1	AmtuPPX2L_R128A_ F420V	SDS-11110	3,5	3	3	3	2,5	2,5	2,5	2,5

Table 8 e: Field data - T1 generation soybeans rated for injury with 1-5 point scale.				
Injury rating taken 3 days after treatment				
Genotype	GOI	Event	Saflufenacil (150 gai/ha)	Saflufenacil (75 gai/ha)
			Rating	
Wildtype		Jake	5	5
LTM382-2	AmtuPPX2L_F420L	SDS-10533	2,5	2,5
LTM382-2	AmtuPPX2L_F420L	SDS-10544	2,5	2,5
LTM382-2	AmtuPPX2L_F420L	SDS-10558	2	2,5
LTM383-1	AmtuPPX2L_F420M	SDS-10645	3	4
LTM383-1	AmtuPPX2L_F420M	SDS-10761	3	3
LTM383-1	AmtuPPX2L_F420M	SDS-10633	3	3
LTM383-1	AmtuPPX2L_F420M	SDS-10635	3,5	3,5
LTM383-1	AmtuPPX2L_F420M	SDS-10646	2,5	2,5
LTM384-1	AmtuPPX2L_R128A_F420L	SDS-10642	2	2
LTM384-1	AmtuPPX2L_R128A_F420L	SDS-10787	2,5	3
LTM385-1	AmtuPPX2L_R128A_F420M	SDS-11052	3	3
LTM385-1	AmtuPPX2L_R128A_F420M	SDS-10985	2	2
LTM385-1	AmtuPPX2L_R128A_F420M	SDS-10990	2,5	2,5
LTM385-1	AmtuPPX2L_R128A_F420M	SDS-11011	2	2
LTM386-1	AmtuPPX2L_R128A_F420I	SDS-10648	3	3
LTM386-1	AmtuPPX2L_R128A_F420I	SDS-10791	2	2
LTM386-1	AmtuPPX2L_R128A_F420I	SDS-11014	2	2
LTM386-1	AmtuPPX2L_R128A_F420I	SDS-10658	3,5	3,5
LTM386-1	AmtuPPX2L_R128A_F420I	SDS-10776	2,5	2
LTM386-1	AmtuPPX2L_R128A_F420I	SDS-11036	2,5	2,5
LTM386-1	AmtuPPX2L_R128A_F420I	SDS-11111	2,5	2,5
LTM386-1	AmtuPPX2L_R128A_F420I	SDS-11118	2	2

Table 8f Soy T0 plants greenhouse data

SEQ ID	event number	Herbicide treatment g ai/ha & injury scores 1 WAT					
			Saflufenacil		1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione		
			0	100	200	25	50
AmtuPPX2L_R128L_F420V	1	0	4	6	3	4	5
	2	0	1	2	0	1	3

Table 8g Field data - T1 generation. Rated for injury (1-5 point scale) 7 or 14 days after treatment (DAT) 1. Herbicide treatment 1 occurred at the V3-V4 stage and herbicide treatment 2 occurred 10 days later at ~V6 stage.

SEQ ID 2 or 4	Event #	Injury rating							
		saflufenacil + BAS 850H	saflufenacil + BAS 850H	BAS 850H	BAS 850H	BAS 850H	saflufenacil	saflufenacil	
	wild type	5	5	5	5	5	5	5	
AmtuPPX2L_R128A_F420L	1	2,5	3	2,5	3	1	1	1	
AmtuPPX2L_R128A_F420L	2	3	3,5	3,5	3,5	3	2	2	
AmtuPPX2L_R128A_F420M	3	2	3	3	3,5	1,5	1,5	1,5	
AmtuPPX2L_R128A_F420M	4	2	3	3	3,5	1,5	1	1	
AmtuPPX2L_R128A_F420I	5	2,5	3	3	3,5	1,5	1	1	
AmtuPPX2L_R128A_F420I	6	3	3,5	3	3,5	3	3	3	
AmtuPPX2L_R128A_F420I	7	2	3	3	3,5	1,5	1,5	1,5	
AmtuPPX2L_R128A_F420I	8	1	2	2,5	2,5	1	2	2	
AmtuPPX2L_R128A_F420I	9	1	1	2,5	1,5	1	1	1	
AmtuPPX2L_R128A_F420V	10	3	3	3,5	3	3	3	3	

Table 8h Greenhouse data – T2 generation; Data are the average injury score (0-9 scale) of up to 4 individuals per homozygous T2 event. Injury was evaluated 1 week after treatment in the greenhouse. BAS800H refers to Saflufenacil/Kixor; BAS 850H refers to 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl)-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazinane-2,4-dione (or “Benzoxazin”), BAS850-Analog refers to 1-methyl-6-(trifluoromethyl)-3-(2,2,7-trifluoro-3-oxo-4-prop-2-ynyl-1,4-benzoxazin-6-yl)pyrimidine-2,4-dione (described in detail in WO2011/57935)

Herbicide	g ai/ha	WT	AmtuPPX2L_R1 28A_F420L	AmtuPPX2L_ R128A_F420M	AmtuPPX2L_ R128A_F420I	AmtuPPX2L_R1 28A_F420V
unsprayed check	0	0,5	1,3	1,0	1,0	1,3
saflufenacil	100	9,0	4,3	4,0	2,0	2,7
BAS 850H	50					
1% (v/v) MSO						
saflufenacil	200	9,0	4,5	5,0	1,8	2,8
BAS 850H	100					
1% (v/v) MSO						
saflufenacil	100	9,0	4,8	5,0	0,5	2,0
flumioxazin	140					
1% (v/v) MSO						
saflufenacil	100	9,0	0,7	1,0	0,3	1,0
sulfentrazone	560					
1% (v/v) MSO						
saflufenacil	100	9,0	5,0	6,0	5,0	4,7
BAS 850-Analog	50					
1% (v/v) MSO						

Table 8i Greenhouse data – T2 generation; Various mixture ratios of saflufenacil and 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl))-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazine-2,4-dione. Data are the average injury score (0-9 scale) of up to 4 individuals per homozygous T2 event. Injury was evaluated 1 week after treatment in the greenhouse. BAS800H refers to Saflufenacil/Kixor; BAS 850H refers to 1,5-dimethyl-6-thioxo-3-(2,2,7-trifluoro-3-oxo-4-(prop-2-ynyl))-3,4-dihydro-2H-benzo[b][1,4]oxazin-6-yl)-1,3,5-triazine-2,4-dione ("Benzoxazin"), all mutants based on AmtuPPX2L (SEQ ID NO:2 or 4)

5

Herbicide	g ai/ha	wild type	R128A_F420 L (event a)	R128A_F420L (event b)	R128A_F420V (event a)	R128A_F420V (event b)	R128A_F420V (event c)
saflufenacil + BAS 850H	unsprayed	0,3	0,3	1,3	2,0	0,8	1,0
	6.25 + 3.125	8,3	4,0	6,0	0,5	0,3	0,0
	12.5 + 6.25	9,0	0,7	6,0	1,0	0,3	1,0
	25 + 12.5	9,0	1,5	7,5	1,7	1,0	3,5
	50 + 25	9,0	2,8	7,5	1,0	2,5	2,0
	100 + 50	9,0	5,0	6,0	2,3	2,3	4,0
	200 + 100	9,0	5,0	6,7	3,5	3,5	4,5
	400 + 200	9,0	4,7	8,5	3,3	2,8	4,3
	800 + 400	9,0	5,3	8,5	3,0	3,8	4,3

Rating	Phenotype (phytotoxicity) of surviving plants
1	no obvious damage (no phytotoxicity)
2	minor amount of leaf damage, plant will survive
3	moderate amount of leaf damage, plant will survive
4	severe amount of leaf damage, plant will survive
5	no surviving plants - all plants dead/dying

The following gives a definition of the injury scores measured above:

Score	Description of injury
5	
0	No Injury
1	Minimal injury, only a few patches of leaf injury or chlorosis.
10	
2	Minimal injury with slightly stronger chlorosis. Overall growth points remain undamaged.
3	Slightly stronger injury on secondary leaf tissue, but primary leaf and growth points are still undamaged.
15	
4	Overall plant morphology is slightly different, some chlorosis and necrosis in secondary growth points and leaf tissue. Stems are intact. Regrowth is highly probable within 1 week.
5	Overall plant morphology is clearly different, some chlorosis and necrosis on a few leaves and growth points, but primary growth point is intact. Stem tissue is still green. Regrowth is highly probably within 1 week.
20	
6	Strong injury can be seen on the new leaflet growth. Plant has a high probability to survive only through regrowth at different growth points. Most of the leaves are chlorotic/necrotic but stem tissue is still green. May have regrowth but with noticeable injured appearance.
25	
7	Most of the active growth points are necrotic. There may be a single growth point that could survive and may be partially chlorotic or green and partially necrotic. Two leaves may still be chlorotic with some green; the rest of the plant including stem is necrotic.
30	
8	Plant will likely die, and all growth points are necrotic. One leaf may still be chlorotic with some green. The remainder of the plant is necrotic.
9	Plant is dead.
35	
*	Not tested

Claims:

1. A method for controlling undesired vegetation at a plant cultivation site, the method comprising the steps of:
 - 5 a) providing, at said site, a plant that comprises at least one nucleic acid comprising a nucleotide sequence encoding a mutated protoporphyrinogen oxidase (PPO) which is resistant or tolerant to a "PPO inhibiting herbicide" and/or
 - b) applying to said site an effective amount of said herbicide,wherein the mutated PPO comprises a sequence of SEQ ID NO: 2, a variant, derivative,
10 orthologue, paralogue or homologue thereof, in which the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, or His, and/or the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala, Leu, Val, Ile, or Met.
- 15 2. The method according to claim 1, wherein the nucleotide sequence of a) comprises the sequence of SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45 or 47, or a variant or derivative thereof.
3. The method according to claims any of claims 1 to 2, wherein the plant comprises at least
20 one additional heterologous nucleic acid comprising a nucleotide sequence encoding a herbicide tolerance enzyme.
4. The method according to any of claims 1 to 3 wherein the PPO inhibiting herbicide is applied in conjunction with one or more additional herbicides.
- 25 5. An isolated and/or recombinant and/or synthetic nucleic acid encoding a mutated PPO polypeptide, wherein the nucleic acid comprises the nucleotide sequence of SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45 or 47, or a variant or derivative thereof, wherein the mutated PPO comprises a sequence of SEQ
30 ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which the amino acid at or corresponding to position 128 of SEQ ID NO:2 is other than Arginine; and/or the amino acid at or corresponding to position 420 of SEQ ID NO: 2 is other than Phenylalanine.
- 35 6. The nucleic acid of claim 6, wherein the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, or His, and/or the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala, Leu, Val, Ile, or Met.
- 40 7. A mutated PPO polypeptide comprising a sequence of SEQ ID NO: 2, a variant, derivative, orthologue, paralogue or homologue thereof, in which the amino acid at or corresponding to position 128 of SEQ ID NO:2 is Leu, Ala, Val, Ile, Met, Tyr, Gly, Asn, Cys, Phe, Ser, Thr, Gln, or His, and the amino acid at or corresponding to position 420 of SEQ ID NO:2 is Ala, Leu, Val, Ile, or Met, wherein said mutated PPO polypeptide confers in-

creased resistance or tolerance to a PPO inhibiting herbicide in a plant as compared to a wild type plant.

- 5 8. A transgenic plant cell transformed by and expressing a nucleic acid encoding a mutated PPO polypeptide as defined in claim 7, wherein expression of the nucleic acid in the plant cell results in increased resistance or tolerance to a PPO inhibiting herbicide as compared to a wild type variety of the plant cell.
- 10 9. The transgenic plant cell of claim 8, wherein the mutated PPO polypeptide encoding nucleic acid comprises a polynucleotide sequence selected from the group consisting of: a) a polynucleotide as shown in SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45 or 47, or a variant or derivative thereof; b) a polynucleotide encoding a polypeptide as shown in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, or a variant or derivative thereof; c) a
15 a polynucleotide comprising at least 60 consecutive nucleotides of any of a) or b); and d) a polynucleotide complementary to the polynucleotide of any of a) through c).
- 20 10. A transgenic plant comprising a plant cell as defined in claim 8 or 9, wherein expression of the mutated PPO polypeptide encoding nucleic acid in the plant results in the plant's increased resistance to PPO inhibiting herbicide as compared to a wild type plant.
11. A plant cell mutagenized to obtain a plant cell which expresses a nucleic acid encoding a mutated PPO polypeptide as defined in claim 7.
- 25 12. A plant that expresses a mutagenized or recombinant mutated PPO polypeptide as defined in claim 7, and wherein said mutated PPO confers upon the plant increased herbicide tolerance as compared to the corresponding wild-type variety of the plant when expressed therein.
- 30 13. A method for growing a plant as defined in claim 10 or 12 while controlling weeds in the vicinity of said plant, said method comprising the steps of:
 - a) growing said plant ; and
 - b) applying a herbicide composition comprising a PPO-inhibiting herbicide to the plant and weeds, wherein the herbicide normally inhibits protoporphyrinogen oxidase, at a level of the herbicide that would inhibit the growth of a corresponding
35 wild-type plant.
- 40 14. A seed produced by a plant comprising a plant cell as defined in claim 8, 9, or 11, or by a plant as defined in claim 10 or 12, wherein the seed is true breeding for an increased resistance to a PPO inhibiting herbicide as compared to a wild type variety of the seed.
15. A method of producing a transgenic plant cell with an increased resistance to a PPO inhibiting herbicide as compared to a wild type variety of the plant cell comprising, trans-

forming the plant cell with an expression cassette comprising a nucleic acid encoding a mutated PPO polypeptide as defined in claim 7.

- 5 16. A method of producing a transgenic plant comprising, (a) transforming a plant cell with an expression cassette comprising a nucleic acid encoding a mutated PPO polypeptide as defined in claim 7, and (b) generating a plant with an increased resistance to PPO inhibiting herbicide from the plant cell.
- 10 17. The method of claim 15 or 16, wherein the nucleic acid encoding the mutated PPO polypeptide comprises a polynucleotide sequence selected from the group consisting of : a) a polynucleotide as shown in SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45 or 47, or a variant or derivative thereof; b) a polynucleotide encoding a polypeptide as shown in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, 36, 38, 40, 42, 44, 46, or 48, or a variant or derivative thereof; c) a polynucleotide comprising at least 60 consecutive nucleotides of any of a) or b); and d) a polynucleotide complementary to the polynucleotide of any of a) through c).
- 15 18. The method of any of claims 15 to 17, wherein the expression cassette further comprises a transcription initiation regulatory region and a translation initiation regulatory region that are functional in the plant.
- 20 19. An expression cassette comprising a nucleic acid encoding a mutated PPO polypeptide as defined in claim 7, a transcription initiation regulatory region and a translation initiation regulatory region that are functional in the plant, and a chloroplast-targeting sequence comprising a nucleotide sequence that encodes a chloroplast transit peptide.
- 25 20. The expression cassette of claim 19, wherein the targeting sequence comprises a nucleotide sequence that encodes a transit peptide comprising the amino acid sequence of SEQ ID NO: 49, 50, 51, 52, or 53.
- 30 21. A method of identifying or selecting a transformed plant cell, plant tissue, plant or part thereof comprising: i) providing a transformed plant cell, plant tissue, plant or part thereof, wherein said transformed plant cell, plant tissue, plant or part thereof comprises a polynucleotide as shown in SEQ ID NO: 1, 3, 5, 7, 9, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33, 35, 37, 39, 41, 43, 45 or 47, or a variant or derivative thereof, wherein the polynucleotide encodes a mutated PPO polypeptide as defined in claim 7 that is used as a selection marker, and wherein said transformed plant cell, plant tissue, plant or part thereof may comprise a further isolated polynucleotide; ii) contacting the transformed plant cell, plant tissue, plant or part thereof with at least one PPO inhibiting compound; iii) determining whether the plant cell, plant tissue, plant or part thereof is affected by the PPO inhibiting compound; and iv) identifying or selecting the transformed plant cell, plant tissue, plant or part thereof.
- 35 40

22. A combination useful for weed control, comprising (a) a polynucleotide encoding a mutated PPO polypeptide as defined in claim 7, which polynucleotide is capable of being expressed in a plant to thereby provide to that plant tolerance to a PPO inhibiting herbicide; and (b) a PPO inhibiting herbicide.
- 5
23. A process for preparing a combination useful for weed control comprising (a) providing a polynucleotide encoding a mutated PPO polypeptide as defined in claim 7, which polynucleotide is capable of being expressed in a plant to thereby provide to that plant tolerance to a PPO inhibiting herbicide; and (b) providing a PPO inhibiting herbicide.
- 10
24. The process according to claim 23, wherein said step of providing a polynucleotide comprises providing a plant containing the polynucleotide.
25. The process according to claim 23, wherein said step of providing a polynucleotide comprises providing a seed containing the polynucleotide.
- 15
26. The process according to claim 25, further comprising a step of applying the PPO inhibiting herbicide to the seed.
- 20
27. Use of a combination as defined in claim 22 to control weeds at a plant cultivation site.

[illegible]

Figure 1 - continued

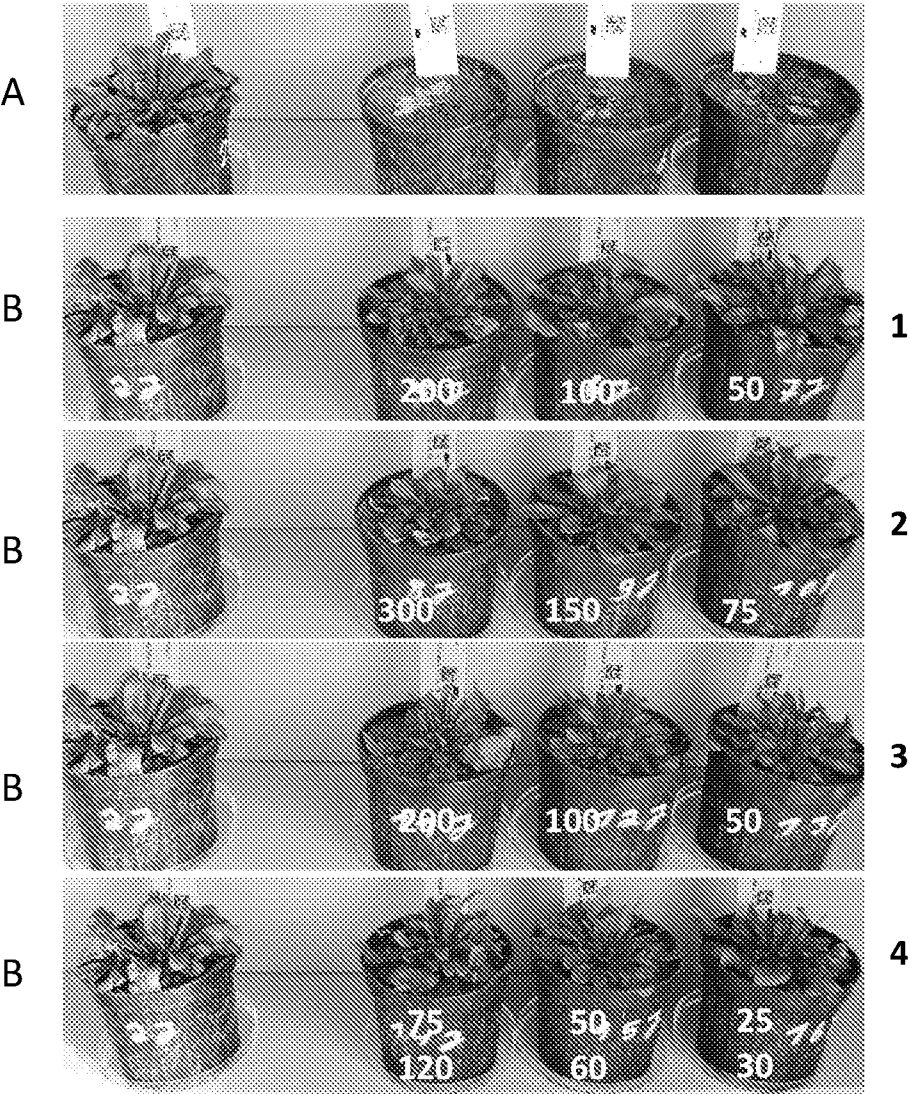
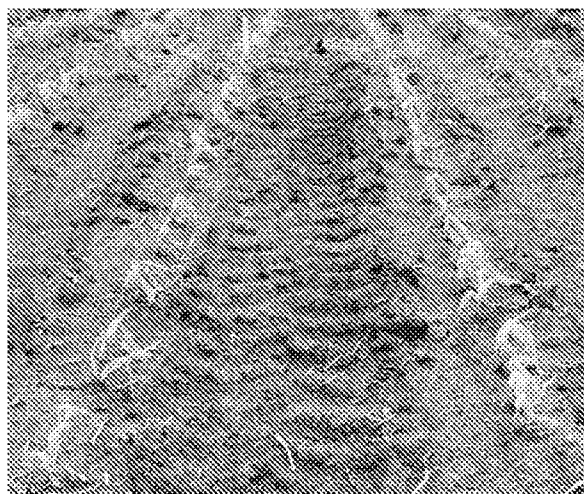
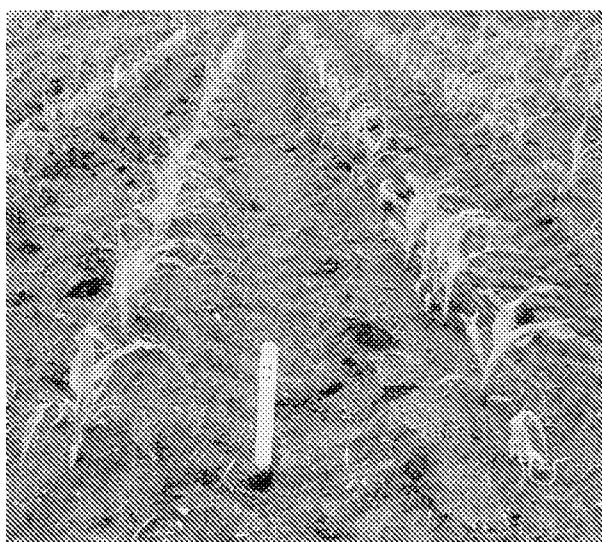


Figure 2



1



2



3

Figure 3

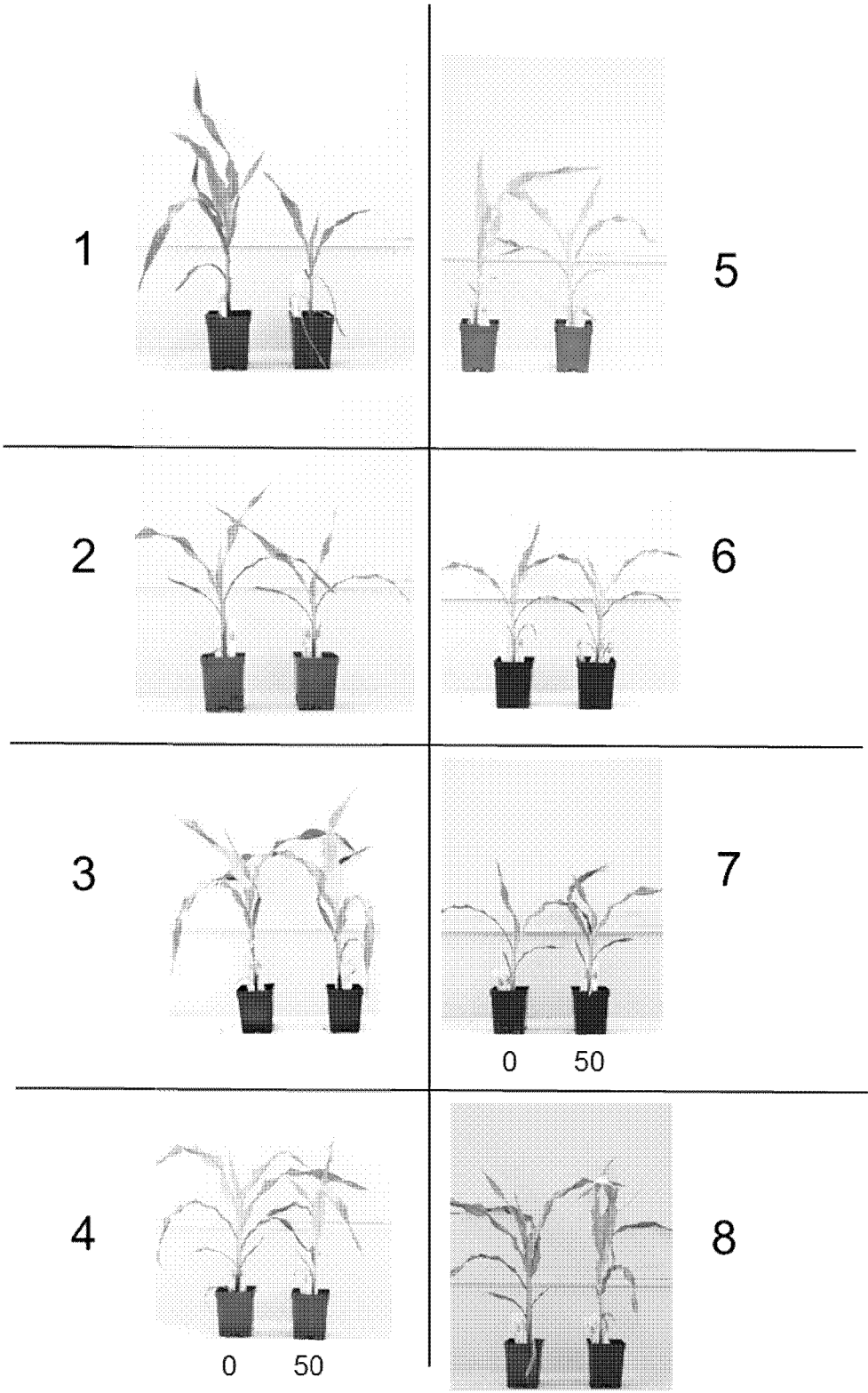


Figure 4

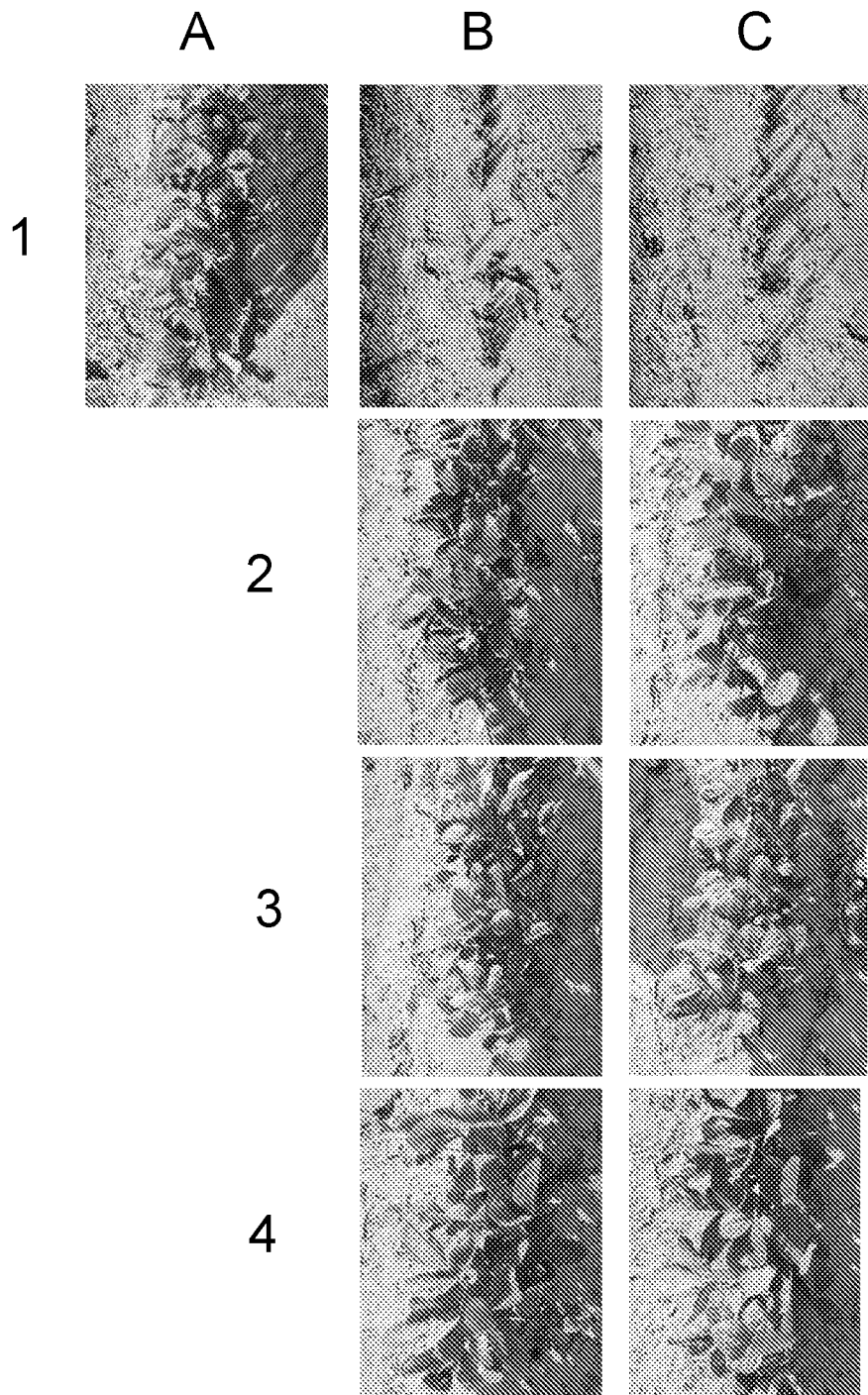


Figure 5

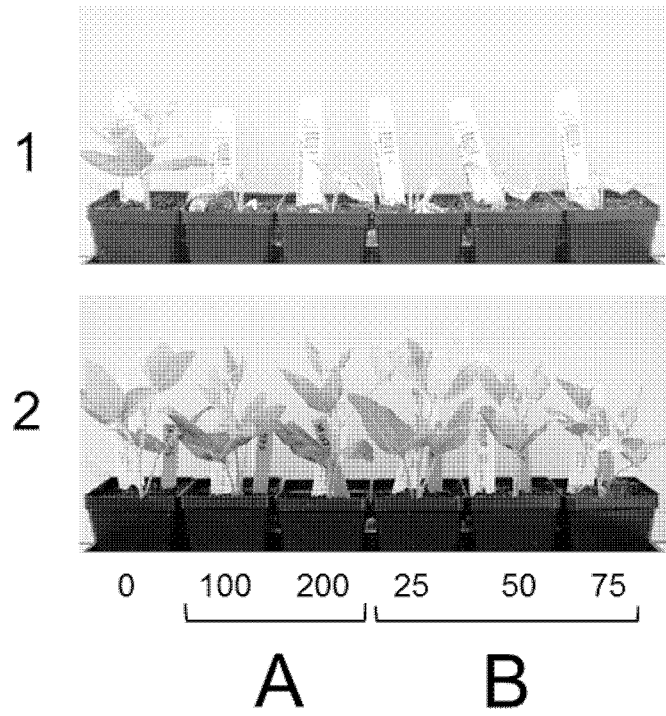


Figure 6

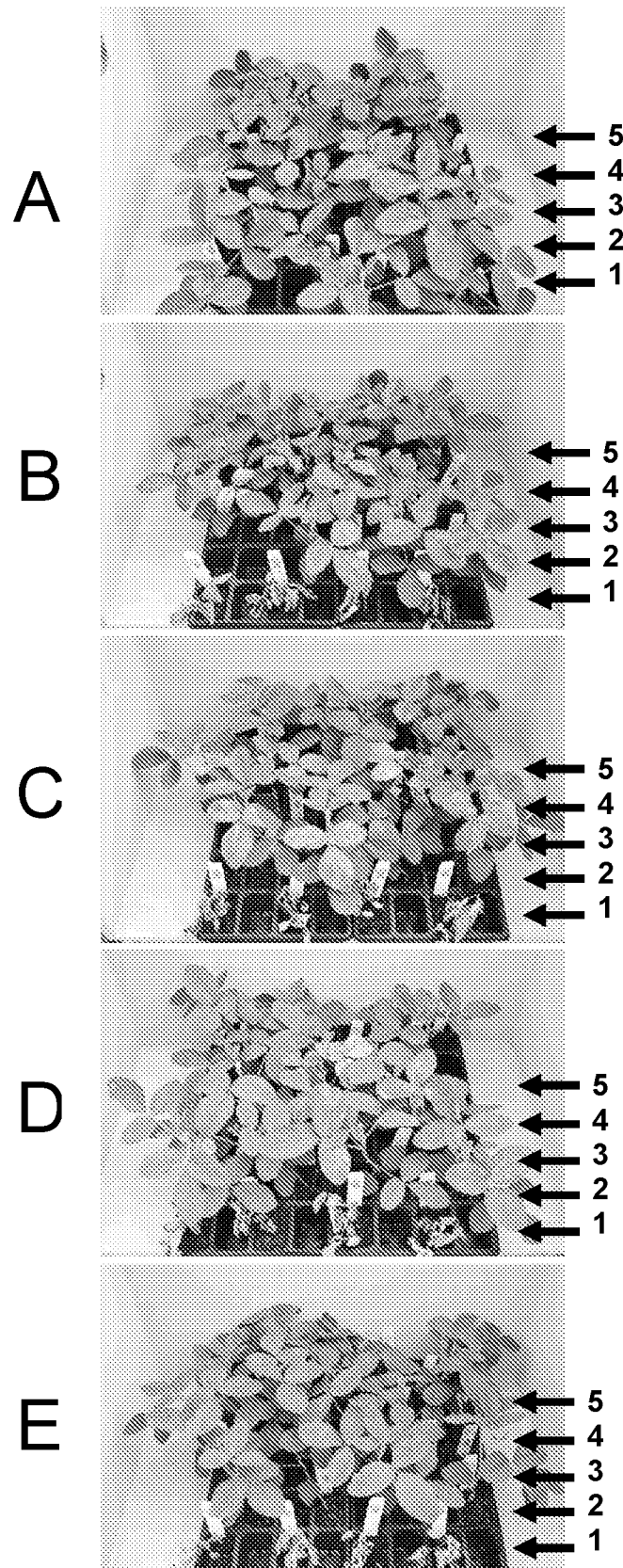


Figure 7

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cagcttgggt aagatgaact caaactccag tgtgaggtgc tgtccttgtc atataaccag	900
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eof-seql

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 gacaagatgg aaaaggatct tcctggattt ttttatgcag gtaaccataa ggggtggactt 1500
 tcagtgggaa aagcgatggc ctccggatgc aaggctgcgg aacttgtaat atcctatctg 1560
 gactctcata tatacgtgaa gatggatgag aagaccgcgt aa 1602

<210> 6
 <211> 533
 <212> PRT
 <213> *Amaranthus tuberculatum*

<400> 6

Met Val Ile Gln Ser Ile Thr His Leu Ser Pro Asn Leu Ala Leu Pro
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Ser Pro Leu Ser Val Ser Thr Lys Asn Tyr Pro Val Ala Val Met Gly
 20 25 30

Asn Ile Ser Glu Arg Glu Glu Pro Thr Ser Ala Lys Arg Val Ala Val
 35 40 45

Val Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Lys Leu Lys Ser
 50 55 60

His Gly Leu Ser Val Thr Leu Phe Glu Ala Asp Ser Arg Ala Gly Gly
 65 70 75 80

Lys Leu Lys Thr Val Lys Lys Asp Gly Phe Ile Trp Asp Glu Gly Ala
 85 90 95

Asn Thr Met Thr Glu Ser Glu Ala Glu Val Ser Ser Leu Ile Asp Asp
 100 105 110

Leu Gly Leu Arg Glu Lys Gln Gln Leu Pro Ile Ser Gln Asn Lys Arg
 115 120 125

Tyr Ile Ala Arg Asp Gly Leu Pro Val Leu Leu Pro Ser Asn Pro Ala
 130 135 140

Ala Leu Leu Thr Ser Asn Ile Leu Ser Ala Lys Ser Lys Leu Gln Ile
 145 150 155 160

Met Leu Glu Pro Phe Leu Trp Arg Lys His Asn Ala Thr Glu Leu Ser
 165 170 175

Asp Glu His Val Gln Glu Ser Val Gly Glu Phe Phe Glu Arg His Phe
 180 185 190

Gly Lys Glu Phe Val Asp Tyr Val Ile Asp Pro Phe Val Ala Gly Thr
 195 200 205

eof-seql

Cys Gly Asp Pro Gln Ser Leu Ser Met His His Thr Phe Pro Glu Val
 210 215 220
 Trp Asn Ile Glu Lys Arg Phe Gly Ser Val Phe Ala Gly Leu Ile Gln
 225 230 235 240
 Ser Thr Leu Leu Ser Lys Lys Glu Lys Gly Gly Glu Asn Ala Ser Ile
 245 250 255
 Lys Lys Pro Arg Val Arg Gly Ser Phe Ser Phe Gln Gly Gly Met Gln
 260 265 270
 Thr Leu Val Asp Thr Met Cys Lys Gln Leu Gly Glu Asp Glu Leu Lys
 275 280 285
 Leu Gln Cys Glu Val Leu Ser Leu Ser Tyr Asn Gln Lys Gly Ile Pro
 290 295 300
 Ser Leu Gly Asn Trp Ser Val Ser Ser Met Ser Asn Asn Thr Ser Glu
 305 310 315 320
 Asp Gln Ser Tyr Asp Ala Val Val Val Thr Ala Pro Ile Arg Asn Val
 325 330 335
 Lys Glu Met Lys Ile Met Lys Phe Gly Asn Pro Phe Ser Leu Asp Phe
 340 345 350
 Ile Pro Glu Val Thr Tyr Val Pro Leu Ser Val Met Ile Thr Ala Phe
 355 360 365
 Lys Lys Asp Lys Val Lys Arg Pro Leu Glu Gly Phe Gly Val Leu Ile
 370 375 380
 Pro Ser Lys Glu Gln His Asn Gly Leu Lys Thr Leu Gly Thr Leu Phe
 385 390 395 400
 Ser Ser Met Met Phe Pro Asp Arg Ala Pro Ser Asp Met Cys Leu Phe
 405 410 415
 Thr Thr Phe Val Gly Gly Ser Arg Asn Arg Lys Leu Ala Asn Ala Ser
 420 425 430
 Thr Asp Glu Leu Lys Gln Ile Val Ser Ser Asp Leu Gln Gln Leu Leu
 435 440 445
 Gly Thr Glu Asp Glu Pro Ser Phe Val Asn His Leu Phe Trp Ser Asn
 450 455 460
 Ala Phe Pro Leu Tyr Gly His Asn Tyr Asp Cys Val Leu Arg Ala Ile
 465 470 475 480

col f-seq1

Asp Lys Met Glu Lys Asp Leu Pro Gly Phe Phe Tyr Ala Gly Asn His
485 490 495

Lys Gly Gly Leu Ser Val Gly Lys Ala Met Ala Ser Gly Cys Lys Ala
500 505 510

Ala Glu Leu Val Ile Ser Tyr Leu Asp Ser His Ile Tyr Val Lys Met
515 520 525

Asp Glu Lys Thr Ala
530

<210> 7
<211> 1602
<212> DNA
<213> Amaranthus tuberculatus

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aagctaaaat cccatggttt gagtgtgaca ttgtttgaag ctaattctag agctggaggc 240
aaacttaaaa ctgttaaaaa agatgggtttt atttgggatg agggggcaaa tactatgaca 300
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tcaacattgt tatctaagaa ggaaaagggt ggagaaaatg cttctattaa gaagcctcgt 780
gtacgtgggt cattttcatt tcaaggtgga atgcagacac ttgttgacac aatgtgcaaa 840
cagcttgggtg aagatgaact caaactccag tgtgaggtgc tgtccttgtc atataaccag 900
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ggaggaagca gaaatagaaa acttgcaaac gcttcaacgg atgaattgaa gcaaatagtt 1320

eof-seql

tcttctgacc ttcagcagct gttgggcact gaggacgaac cttcatttgt caatcatctc 1380
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 gacaagatgg aaaaggatct tcctggattt ttttatgcag gtaaccataa ggggtggactt 1500
 tcagtgggaa aagcgatggc ctccggatgc aaggctgcgg aacttgtaat atcctatctg 1560
 gactctcata tatacgtgaa gatggatgag aagaccgcgt aa 1602

<210> 8
 <211> 533
 <212> PRT
 <213> *Amaranthus tuberculatum*

<400> 8

Met Val Ile Gln Ser Ile Thr His Leu Ser Pro Asn Leu Ala Leu Pro
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Ser Pro Leu Ser Val Ser Thr Lys Asn Tyr Pro Val Ala Val Met Gly
 20 25 30

Asn Ile Ser Glu Arg Glu Glu Pro Thr Ser Ala Lys Arg Val Ala Val
 35 40 45

Val Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Lys Leu Lys Ser
 50 55 60

His Gly Leu Ser Val Thr Leu Phe Glu Ala Asn Ser Arg Ala Gly Gly
 65 70 75 80

Lys Leu Lys Thr Val Lys Lys Asp Gly Phe Ile Trp Asp Glu Gly Ala
 85 90 95

Asn Thr Met Thr Glu Ser Glu Ala Glu Val Ser Ser Leu Ile Asp Asp
 100 105 110

Leu Gly Leu Arg Glu Lys Gln Gln Leu Pro Ile Ser Gln Asn Lys Arg
 115 120 125

Tyr Ile Ala Arg Asp Gly Leu Pro Val Leu Leu Pro Ser Asn Pro Ala
 130 135 140

Ala Leu Leu Thr Ser Asn Ile Leu Ser Ala Lys Ser Lys Leu Gln Ile
 145 150 155 160

Met Leu Glu Pro Phe Leu Trp Arg Lys His Asn Ala Thr Glu Leu Ser
 165 170 175

Asp Glu His Val Gln Glu Ser Val Gly Glu Phe Phe Glu Arg His Phe
 180 185 190

Gly Lys Glu Phe Val Asp Tyr Val Ile Asp Pro Phe Val Ala Gly Thr
 195 200 205

eof-seq1

Cys Gly Asp Pro Gln Ser Leu Ser Met Tyr His Thr Phe Pro Glu Val
 210 215 220
 Trp Asn Ile Glu Lys Arg Phe Gly Ser Val Phe Ala Gly Leu Ile Gln
 225 230 235 240
 Ser Thr Leu Leu Ser Lys Lys Glu Lys Gly Gly Glu Asn Ala Ser Ile
 245 250 255
 Lys Lys Pro Arg Val Arg Gly Ser Phe Ser Phe Gln Gly Gly Met Gln
 260 265 270
 Thr Leu Val Asp Thr Met Cys Lys Gln Leu Gly Glu Asp Glu Leu Lys
 275 280 285
 Leu Gln Cys Glu Val Leu Ser Leu Ser Tyr Asn Gln Lys Gly Ile Pro
 290 295 300
 Ser Leu Gly Asn Trp Ser Val Ser Ser Met Ser Asn Asn Thr Ser Glu
 305 310 315 320
 Asp Gln Ser Tyr Asp Ala Val Val Val Thr Ala Pro Ile Arg Asn Val
 325 330 335
 Lys Glu Met Lys Ile Met Lys Phe Gly Asn Pro Phe Ser Leu Asp Phe
 340 345 350
 Ile Pro Glu Val Thr Tyr Val Pro Leu Ser Val Met Ile Thr Ala Phe
 355 360 365
 Lys Lys Asp Lys Val Lys Arg Pro Leu Glu Gly Phe Gly Val Leu Ile
 370 375 380
 Pro Ser Lys Glu Gln His Asn Gly Leu Lys Thr Leu Gly Thr Leu Phe
 385 390 395 400
 Ser Ser Met Met Phe Pro Asp Arg Ala Pro Ser Asp Met Cys Leu Phe
 405 410 415
 Thr Thr Phe Val Gly Gly Ser Arg Asn Arg Lys Leu Ala Asn Ala Ser
 420 425 430
 Thr Asp Glu Leu Lys Gln Ile Val Ser Ser Asp Leu Gln Gln Leu Leu
 435 440 445
 Gly Thr Glu Asp Glu Pro Ser Phe Val Asn His Leu Phe Trp Ser Asn
 450 455 460
 Ala Phe Pro Leu Tyr Gly His Asn Tyr Asp Ser Val Leu Arg Ala Ile
 465 470 475 480

eol f-seql

Asp Lys Met Glu Lys Asp Leu Pro Gly Phe Phe Tyr Al a Gly Asn Hi s
485 490 495

Lys Gly Gly Leu Ser Val Gly Lys Al a Met Al a Ser Gly Cys Lys Al a
500 505 510

Al a Glu Leu Val Ile Ser Tyr Leu Asp Ser Hi s Ile Tyr Val Lys Met
515 520 525

Asp Glu Lys Thr Al a
530

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<211> 1644
<212> DNA
<213> Arabidopsi s thal i ana

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gaattgttgc agatagcaat ggcgtctgga gcagtagcag atcatcaa at tgaagcggtt 180
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ccaatttcac agaaaaagcg gtatatgttg cggaatgggtg tacctgtgat gctacctacc 480
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ccttttgttg gtggaacaag tgctgcggac cctgattccc tttcaatgaa gcattctttc 720
ccagatctct ggaatagttt tggctctatt atagtcgggtg caatcagaac aaagtttgct 780
gctaaagggtg gtaaaagtag agacacaaag agttctcctg gcacaaaaaa gggttcgcgt 840
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tcacatgatg agatcaattt agactccaag gtactctctt tgtcttaca tttctggatca 960
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cccttcagc taaactttct ccccagatt aattacatgc ccctctcgggt tttaatcacc 1140
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aaggagcaaa agcatggttt caaaactcta ggtacacttt tttcatcaat gatgtttcca 1260
gatcgttccc ctagtgacgt tcatctatat acaactttta ttggtgggag taggaaccag 1320

eof-seql

gaactagcca aagcttcac tgacgaatta aaacaagttg tgacttctga ccttcagcga 1380
ctgttggggg ttgaaggtag acccgtgtct gtcaaccatt actattggag gaaagcattc 1440
ccgttgtatg acagcagcta tgactcagtc atggaagcaa ttgacaagat ggagaatgat 1500
ctacctgggt tcttctatgc aggtaatcat cgaggggggc tctctgttgg gaaatcaata 1560
gcatcaggtt gcaaagcagc tgaccttgtg atctcatacc tggagtcttg ctcaaagac 1620
aagaaaccaa atgacagctt ataa 1644

<210> 10
<211> 547
<212> PRT
<213> Arabidopsis thaliana
<400> 10

Met Gly Leu Ile Lys Asn Gly Thr Leu Tyr Cys Arg Phe Gly Ile Ser
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Trp Asn Phe Ala Ala Val Phe Phe Ser Thr Tyr Phe Arg His Cys Phe
20 25 30

Arg Leu Val Arg Asp Phe Asp Ser Glu Leu Leu Gln Ile Ala Met Ala
35 40 45

Ser Gly Ala Val Ala Asp His Gln Ile Glu Ala Val Ser Gly Lys Arg
50 55 60

Val Ala Val Val Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Lys
65 70 75 80

Leu Lys Ser Arg Gly Leu Asn Val Thr Val Phe Glu Ala Asp Gly Arg
85 90 95

Val Gly Gly Lys Leu Arg Ser Val Met Gln Asn Gly Leu Ile Trp Asp
100 105 110

Glu Gly Ala Asn Thr Met Thr Glu Ala Glu Pro Glu Val Gly Ser Leu
115 120 125

Leu Asp Asp Leu Gly Leu Arg Glu Lys Gln Gln Phe Pro Ile Ser Gln
130 135 140

Lys Lys Arg Tyr Ile Val Arg Asn Gly Val Pro Val Met Leu Pro Thr
145 150 155 160

Asn Pro Ile Glu Leu Val Thr Ser Ser Val Leu Ser Thr Gln Ser Lys
165 170 175

Phe Gln Ile Leu Leu Glu Pro Phe Leu Trp Lys Lys Lys Ser Ser Lys
180 185 190

Val Ser Asp Ala Ser Ala Glu Glu Ser Val Ser Glu Phe Phe Gl n Arg
 195 200 eol f-seql 205
 Hi s Phe Gly Gl n Gl u Val Val Asp Tyr Leu Ile Asp Pro Phe Val Gly
 210 215 220
 Gly Thr Ser Ala Ala Asp Pro Asp Ser Leu Ser Met Lys Hi s Ser Phe
 225 230 235 240
 Pro Asp Leu Trp Asn Ser Phe Gly Ser Ile Ile Val Gly Ala Ile Arg
 245 250 255
 Thr Lys Phe Ala Ala Lys Gly Gly Lys Ser Arg Asp Thr Lys Ser Ser
 260 265 270
 Pro Gly Thr Lys Lys Gly Ser Arg Gly Ser Phe Ser Phe Lys Gly Gly
 275 280 285
 Met Gl n Ile Leu Pro Asp Thr Leu Cys Lys Ser Leu Ser Hi s Asp Gl u
 290 295 300
 Ile Asn Leu Asp Ser Lys Val Leu Ser Leu Ser Tyr Asn Ser Gly Ser
 305 310 315 320
 Arg Gl n Gl u Asn Trp Ser Leu Ser Cys Val Ser Hi s Asn Gl u Thr Gl n
 325 330 335
 Arg Gl n Asn Pro Hi s Tyr Asp Ala Ala Pro Leu Cys Asn Val Lys Gl u
 340 345 350
 Met Lys Val Met Lys Gly Gly Gl n Pro Phe Gl n Leu Asn Phe Leu Pro
 355 360 365
 Gl u Ile Asn Tyr Met Pro Leu Ser Val Leu Ile Thr Thr Phe Thr Lys
 370 375 380
 Gl u Lys Val Lys Arg Pro Leu Gl u Gly Phe Gly Val Leu Ile Pro Ser
 385 390 395 400
 Lys Gl u Gl n Lys Hi s Gly Phe Lys Thr Leu Gly Thr Leu Phe Ser Ser
 405 410 415
 Met Met Phe Pro Asp Arg Ser Pro Ser Asp Val Hi s Leu Tyr Thr Thr
 420 425 430
 Phe Ile Gly Gly Ser Arg Asn Gl n Gl u Leu Ala Lys Ala Ser Thr Asp
 435 440 445
 Gl u Leu Lys Gl n Val Val Thr Ser Asp Leu Gl n Arg Leu Leu Gly Val
 450 455 460

eol f-seql

Glu Gly Glu Pro Val Ser Val Asn His Tyr Tyr Trp Arg Lys Ala Phe
 465 470 475 480

Pro Leu Tyr Asp Ser Ser Tyr Asp Ser Val Met Glu Ala Ile Asp Lys
 485 490 495

Met Glu Asn Asp Leu Pro Gly Phe Phe Tyr Ala Gly Asn His Arg Gly
 500 505 510

Gly Leu Ser Val Gly Lys Ser Ile Ala Ser Gly Cys Lys Ala Ala Asp
 515 520 525

Leu Val Ile Ser Tyr Leu Glu Ser Cys Ser Asn Asp Lys Lys Pro Asn
 530 535 540

Asp Ser Leu
 545

<210> 11
 <211> 1647
 <212> DNA
 <213> Nicotiana tabacum

<400> 11
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 agtgtcaatt gcaatggctg gagaacacga tgctccgttg ccaaagatta cacagttcct 180
 tcctcagcgg tcgacggcgg acccgccgcg gagctggact gtgttatagt tggagcagga 240
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eof-seq1

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 <211> 548
 <212> PRT
 <213> Nicotiana tabacum

<400> 12

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Ser Ser Ser Ser Pro Leu Ala Phe Leu Asn Arg Thr Ser Phe Ile Pro
 20 25 30

Phe Ser Ser Ile Ser Lys Arg Asn Ser Val Asn Cys Asn Gly Trp Arg
 35 40 45

Thr Arg Cys Ser Val Ala Lys Asp Tyr Thr Val Pro Ser Ser Ala Val
 50 55 60

Asp Gly Gly Pro Ala Ala Glu Leu Asp Cys Val Ile Val Gly Ala Gly
 65 70 75 80

Ile Ser Gly Leu Cys Ile Ala Gln Val Met Ser Ala Asn Tyr Pro Asn
 85 90 95

Leu Met Val Thr Glu Ala Arg Asp Arg Ala Gly Gly Asn Ile Thr Thr
 100 105 110

Val Glu Arg Asp Gly Tyr Leu Trp Glu Glu Gly Pro Asn Ser Phe Gln
 115 120 125

Pro Ser Asp Pro Met Leu Thr Met Ala Val Asp Cys Gly Leu Lys Asp
 130 135 140

Asp Leu Val Leu Gly Asp Pro Asn Ala Pro Arg Phe Val Leu Trp Lys
 145 150 155 160

Gly Lys Leu Arg Pro Val Pro Ser Lys Leu Thr Asp Leu Pro Phe Phe
 165 170 175

eof-seql

Asp Leu Met Ser Ile Pro Gly Lys Leu Arg Ala Gly Phe Gly Ala Ile
 180 185 190
 Gly Leu Arg Pro Ser Pro Pro Gly His Glu Glu Ser Val Glu Gln Phe
 195 200 205
 Val Arg Arg Asn Leu Gly Gly Glu Val Phe Glu Arg Leu Ile Glu Pro
 210 215 220
 Phe Cys Ser Gly Val Tyr Ala Gly Asp Pro Ser Lys Leu Ser Met Lys
 225 230 235 240
 Ala Ala Phe Gly Lys Val Trp Lys Leu Glu Glu Thr Gly Gly Ser Ile
 245 250 255
 Ile Gly Gly Thr Phe Lys Ala Ile Lys Glu Arg Ser Ser Thr Pro Lys
 260 265
 Ala Pro Arg Asp Pro Arg Leu Pro Lys Pro Lys Gly Gln Thr Val Gly
 275 280 285
 Ser Phe Arg Lys Gly Leu Arg Met Leu Pro Asp Ala Ile Ser Ala Arg
 290 295 300
 Leu Gly Ser Lys Leu Lys Leu Ser Trp Lys Leu Ser Ser Ile Thr Lys
 305 310 315 320
 Ser Glu Lys Gly Gly Tyr His Leu Thr Tyr Glu Thr Pro Glu Gly Val
 325 330 335
 Val Ser Leu Gln Ser Arg Ser Ile Val Met Thr Val Pro Ser Tyr Val
 340 345 350
 Ala Ser Asn Ile Leu Arg Pro Leu Ser Val Ala Ala Ala Asp Ala Leu
 355 360 365
 Ser Asn Phe Tyr Tyr Pro Pro Val Gly Ala Val Thr Ile Ser Tyr Pro
 370 375 380
 Gln Glu Ala Ile Arg Asp Glu Arg Leu Val Asp Gly Glu Leu Lys Gly
 385 390 395 400
 Phe Gly Gln Leu His Pro Arg Thr Gln Gly Val Glu Thr Leu Gly Thr
 405 410 415
 Ile Tyr Ser Ser Ser Leu Phe Pro Asn Arg Ala Pro Lys Gly Arg Val
 420 425 430
 Leu Leu Leu Asn Tyr Ile Gly Gly Ala Lys Asn Pro Glu Ile Leu Ser
 435 440 445

eof-seq1

Lys Thr Glu Ser Gln Leu Val Glu Val Val Asp Arg Asp Leu Arg Lys
450 455 460

Met Leu Ile Lys Pro Lys Ala Gln Asp Pro Leu Val Val Gly Val Arg
465 470 475 480

Val Trp Pro Gln Ala Ile Pro Gln Phe Leu Val Gly His Leu Asp Thr
485 490 495

Leu Ser Thr Ala Lys Ala Ala Met Asn Asp Asn Gly Leu Glu Gly Leu
500 505 510

Phe Leu Gly Gly Asn Tyr Val Ser Gly Val Ala Leu Gly Arg Cys Val
515 520 525

Glu Gly Ala Tyr Glu Val Ala Ser Glu Val Thr Gly Phe Leu Ser Arg
530 535 540

Tyr Ala Tyr Lys
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<210> 13
<211> 1668
<212> DNA
<213> Cichorium intybus

<400> 13
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eof-seql

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<210> 14
 <211> 555
 <212> PRT
 <213> Cichorium

<400> 14

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 Pro Arg Tyr Leu Ile Thr Tyr Ser Pro Ala His Arg Lys Cys Asn Arg
 35 40 45
 Trp Arg Phe Arg Cys Ser Ile Ala Lys Asp Ser Pro Ile Thr Pro Pro
 50 55 60
 Ile Ser Asn Glu Phe Asn Ser Gln Pro Leu Leu Asp Cys Val Ile Val
 65 70 75 80
 Gly Ala Gly Ile Ser Gly Leu Cys Ile Ala Gln Ala Leu Ala Thr Lys
 85 90 95
 His Ala Ser Val Ser Pro Asp Val Ile Val Thr Glu Ala Arg Asp Arg
 100 105 110
 Val Gly Gly Asn Ile Ser Thr Val Glu Arg Asp Gly Tyr Leu Trp Glu
 115 120 125
 Glu Gly Pro Asn Ser Phe Gln Pro Ser Asp Ala Met Leu Thr Met Val
 130 135 140

eol f-seql

Val 145	Asp	Ser	Gly	Leu	Lys 150	Asp	Asp	Leu	Val	Leu 155	Gly	Asp	Pro	Thr	Ala 160
Pro	Arg	Phe	Val	Leu 165	Trp	Gly	Gly	Asp	Leu 170	Lys	Pro	Val	Pro	Ser 175	Lys
Pro	Ala	Asp	Leu 180	Pro	Phe	Phe	Asp	Leu 185	Met	Ser	Phe	Pro	Gly 190	Lys	Leu
Arg	Ala	Gly 195	Phe	Gly	Ala	Leu	Gly 200	Phe	Arg	Pro	Ser	Pro 205	Pro	Asp	Arg
Glu	Glu 210	Ser	Val	Glu	Glu	Phe 215	Val	Arg	Arg	Asn	Leu 220	Gly	Asp	Glu	Val
Phe 225	Glu	Arg	Leu	Ile	Glu 230	Pro	Phe	Cys	Ser	Gly 235	Val	Tyr	Ala	Gly	Asp 240
Pro	Ser	Lys	Leu	Ser 245	Met	Lys	Ala	Ala	Phe 250	Gly	Lys	Val	Trp	Asn 255	Leu
Glu	Gln	Asn	Gly 260	Gly	Ser	Ile	Val	Gly 265	Gly	Ala	Phe	Lys	Ala 270	Ile	Gln
Asp	Arg	Lys 275	Asn	Ser	Gln	Lys	Pro 280	Pro	Arg	Asp	Pro	Arg 285	Leu	Pro	Lys
Pro	Lys 290	Gly	Gln	Thr	Val	Gly 295	Ser	Phe	Arg	Lys	Gly 300	Gln	Ala	Met	Leu
Pro 305	Asn	Ala	Ile	Ser	Thr 310	Arg	Leu	Gly	Ser	Arg 315	Val	Lys	Leu	Cys	Trp 320
Lys	Leu	Thr	Ser	Ile 325	Ser	Lys	Leu	Glu	Asn 330	Arg	Gly	Tyr	Asn	Leu 335	Thr
Tyr	Glu	Thr	Pro 340	Gln	Gly	Phe	Glu	Ser 345	Leu	Gln	Thr	Lys	Thr 350	Ile	Val
Met	Thr	Val 355	Pro	Ser	Tyr	Val	Ala 360	Ser	Asp	Leu	Leu	Arg 365	Pro	Leu	Ser
Leu	Gly 370	Ala	Ala	Asp	Ala	Leu 375	Ser	Lys	Phe	Tyr	Tyr 380	Pro	Pro	Val	Ala
Ala 385	Val	Ser	Ile	Ser	Tyr 390	Pro	Lys	Asp	Ala	Ile 395	Arg	Ala	Asp	Arg	Leu 400
Ile	Asp	Gly	Gln	Leu 405	Lys	Gly	Phe	Gly	Gln 410	Leu	His	Pro	Arg	Ser 415	Gln

eol f-seql

Gly Val Glu Thr Leu Gly Thr Ile Tyr Ser Ser Ser Leu Phe Pro Asn
420 425 430

Arg Ala Pro Pro Gly Arg Val Leu Leu Leu Asn Tyr Ile Gly Gly Ala
435 440 445

Thr Asn Pro Glu Ile Leu Ser Lys Thr Glu Gly Glu Ile Val Asp Ala
450 455 460

Val Asp Arg Asp Leu Arg Thr Met Leu Ile Arg Arg Asp Ala Glu Asp
465 470 475 480

Pro Leu Thr Leu Gly Val Arg Val Trp Pro Arg Ala Ile Pro Gln Phe
485 490 495

Leu Ile Gly His Tyr Asp Ile Leu Asp Ser Ala Lys Ala Ala Leu Ser
500 505 510

Ser Gly Gly Phe Gln Gly Met Phe Leu Gly Gly Asn Tyr Val Ser Gly
515 520 525

Val Ala Leu Gly Lys Cys Val Glu Ala Ala Tyr Asp Val Ala Ala Glu
530 535 540

Val Met Asn Phe Leu Ser Gln Gly Val Tyr Lys
545 550 555

<210> 15
<211> 1689
<212> DNA
<213> Spinacia oleracea

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cgaggaggaa gctctatccg ctgctctaca atctcaacct ctaattccgc ggctgcagcc 180
aattaccaga acaaaaacat aggcacaaac ggagttgacg gcggcgagg cgaggagggt 240
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caatttgtcc gtcgtaatct tgggtgatgag gtctttgaac gcttgatcga acctttttgt 720
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eol f-seql

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 gataaatag 1689

<210> 16
 <211> 562
 <212> PRT
 <213> Spi naci a
 <400> 16

Met Ser Ala Met Ala Leu Ser Ser Thr Met Ala Leu Ser Leu Pro Gln
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 Ser Ser Met Ser Leu Ser His Cys Arg His Asn Arg Ile Thr Ile Leu
 20 25 30
 Ile Pro Ser Ser Ser Leu Arg Arg Arg Gly Gly Ser Ser Ile Arg Cys
 35 40 45
 Ser Thr Ile Ser Thr Ser Asn Ser Ala Ala Ala Ala Asn Tyr Gln Asn
 50 55 60
 Lys Asn Ile Gly Thr Asn Gly Val Asp Gly Gly Gly Gly Gly Gly Gly
 65 70 75 80
 Val Leu Asp Cys Val Ile Val Gly Gly Gly Ile Ser Gly Leu Cys Ile
 85 90 95
 Ala Gln Ala Leu Ser Thr Lys Tyr Ser Asn Leu Ser Thr Asn Phe Ile
 100 105 110

eol f-seq1

Val Thr Glu Ala Lys Asp Arg Val Gly Gly Asn Ile Thr Thr Met Glu
115 120 125

Ala Asp Gly Tyr Leu Trp Glu Glu Gly Pro Asn Ser Phe Gl n Pro Ser
130 135 140

Asp Ala Val Leu Thr Met Ala Val Asp Ser Gly Leu Lys Gl u Gl u Leu
145 150 155 160

Val Leu Gly Asp Pro Asn Ser Pro Arg Phe Val Leu Trp Asn Gly Lys
165 170 175

Leu Arg Pro Val Pro Ser Lys Leu Thr Asp Leu Pro Phe Phe Asp Leu
180 185 190

Met Ser Phe Pro Gly Lys Ile Arg Ala Gly Leu Gly Ala Leu Gly Leu
195 200 205

Arg Pro Ser Pro Pro Ala His Glu Glu Ser Val Glu Gl n Phe Val Arg
210 215 220

Arg Asn Leu Gly Asp Glu Val Phe Glu Arg Leu Ile Glu Pro Phe Cys
225 230 235 240

Ser Gly Val Tyr Ala Gly Asp Pro Ser Lys Leu Ser Met Lys Ala Ala
245 250 255

Phe Gly Arg Val Trp Val Leu Glu Gl n Lys Gly Gly Ser Ile Ile Gly
260 265 270

Gly Thr Leu Lys Thr Ile Gl n Glu Arg Lys Asp Asn Pro Lys Pro Pro
275 280 285

Arg Asp Pro Arg Leu Pro Lys Pro Lys Gly Gl n Thr Val Gly Ser Phe
290 295 300

Arg Lys Gly Leu Ser Met Leu Pro Thr Ala Ile Ser Glu Arg Leu Gly
305 310 315 320

Asn Lys Val Lys Val Ser Trp Thr Leu Ser Gly Ile Ala Lys Ser Ser
325 330 335

Asn Gly Glu Tyr Asn Leu Thr Tyr Glu Thr Pro Asp Gly Leu Val Ser
340 345 350

Val Arg Thr Lys Ser Val Val Met Thr Val Pro Ser Tyr Val Ala Ser
355 360 365

Ser Leu Leu Arg Pro Leu Ser Asp Val Ala Ala Glu Ser Leu Ser Lys
370 375 380

eol f-seql

Phe His Tyr Pro Pro Val Ala Ala Val Ser Leu Ser Tyr Pro Lys Glu
385 390 395 400

Ala Ile Arg Ser Glu Cys Leu Ile Asp Gly Glu Leu Lys Gly Phe Gly
405 410 415

Gln Leu His Ser Arg Ser Gln Gly Val Glu Thr Leu Gly Thr Ile Tyr
420 425 430

Ser Ser Ser Leu Phe Pro Gly Arg Ala Pro Pro Gly Arg Thr Leu Ile
435 440 445

Leu Asn Tyr Ile Gly Gly Asp Thr Asn Pro Gly Ile Leu Asp Lys Thr
450 455 460

Lys Asp Glu Leu Ala Glu Ala Val Asp Arg Asp Leu Arg Arg Ile Leu
465 470 475 480

Ile Asn Pro Asn Ala Lys Ala Pro Arg Val Leu Gly Val Arg Val Trp
485 490 495

Pro Gln Ala Ile Pro Gln Phe Leu Ile Gly His Phe Asp Leu Leu Asp
500 505 510

Ala Ala Lys Ala Ala Leu Thr Asp Gly Gly His Lys Gly Leu Phe Leu
515 520 525

Gly Gly Asn Tyr Val Ser Gly Val Ala Leu Gly Arg Cys Ile Glu Gly
530 535 540

Ala Tyr Glu Ser Ala Ala Glu Val Val Asp Phe Leu Ser Gln Tyr Ser
545 550 555 560

Asp Lys

<210> 17
<211> 1596
<212> DNA
<213> Spinacia oleracea

<400> 17
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ctcaaaactg ttgtaaagga tggtttgatt tgggatgaag gggcaaatac catgacagag 300
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ccaatttcac aaaacaaaag atacattgcc agagatggtc ttcctgtgct gttaccttca 420

eol f-seql

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cataatggaa gccttacatc agagaattgg tcagtgtctt ctatgtcaaa cagcaccatc      960
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<210> 18
 <211> 531
 <212> PRT
 <213> Spinacia

<400> 18

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Leu Val Ser Pro Thr Lys Asn Asn Pro Val Met Gly Asn Val Ser Glu
20 25 30

Arg Asn Gln Val Asn Gln Pro Ile Ser Ala Lys Arg Val Ala Val Val
35 40 45

Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Lys Leu Lys Ser Asn
50 55 60

Gly Leu Asn Val Thr Leu Phe Glu Ala Asp Ser Arg Ala Gly Gly Lys
65 70 75 80

eol f-seql

Leu Lys Thr Val Val₈₅ Lys Asp Gly Leu Ile₉₀ Trp Asp Glu Gly Ala₉₅ Asn

Thr Met Thr Glu₁₀₀ Ser Asp Glu Glu Val₁₀₅ Thr Ser Leu Phe Asp₁₁₀ Asp Leu

Gly Ile Arg₁₁₅ Glu Lys Leu Glu₁₂₀ Leu Pro Ile Ser Glu Asn₁₂₅ Lys Arg Tyr

Ile Ala₁₃₀ Arg Asp Gly Leu Pro₁₃₅ Val Leu Leu Pro Ser₁₄₀ Asn Pro Val Ala

Leu₁₄₅ Leu Lys Ser Asn Ile₁₅₀ Leu Ser Ala Lys Ser₁₅₅ Lys Leu Glu Ile Met₁₆₀

Leu Glu Pro Phe Leu₁₆₅ Trp Lys Lys His Asn₁₇₀ Gly Ala Lys Val Ser₁₇₅ Asp

Glu Asn Ala Glu₁₈₀ Glu Ser Val Ala Glu₁₈₅ Phe Phe Glu Arg His₁₉₀ Phe Gly

Lys Glu Phe₁₉₅ Val Asp Tyr Leu Ile₂₀₀ Asp Pro Phe Val Ala₂₀₅ Gly Thr Ser

Gly Gly₂₁₀ Asp Pro Glu Ser Leu₂₁₅ Ser Met Arg His Ala₂₂₀ Phe Pro Glu Leu

Trp₂₂₅ Asn Ile Glu Asn Arg₂₃₀ Phe Gly Ser Val Ile₂₃₅ Ser Gly Phe Ile Glu₂₄₀

Ser Lys Leu Ser Ser₂₄₅ Lys Lys Glu Lys Gly₂₅₀ Gly Glu Lys Glu Ser₂₅₅ Ser

Asn Lys Lys Pro₂₆₀ Arg Val Arg Gly Ser₂₆₅ Phe Ser Phe Glu Gly₂₇₀ Gly Met

Glu Thr Leu₂₇₅ Val Asp Thr Ile Cys₂₈₀ Lys Glu Phe Gly Glu₂₈₅ Asp Glu Leu

Lys Leu₂₉₀ Glu Ser Glu Val Leu₂₉₅ Ser Leu Ser Tyr Ser₃₀₀ His Asn Gly Ser

Leu Thr Ser Glu Asn Trp₃₁₀ Ser Val Ser Ser Met₃₁₅ Ser Asn Ser Thr Ile₃₂₀

Glu Asp Glu Pro Tyr₃₂₅ Asp Ala Val Val Val₃₃₀ Thr Ala Pro Ile Asn₃₃₅ Asn

Val Lys Glu Leu₃₄₀ Lys Ile Met Lys Val₃₄₅ Glu Asn Pro Phe Ser₃₅₀ Leu Asp

eol f-seql

Phe Ile Pro Glu Val Ser Cys Leu Pro Leu Ser Val Ile Ile Thr Thr
355 360 365

Phe Lys Lys Thr Asn Val Lys Arg Pro Leu Glu Gly Phe Gly Val Leu
370 375 380

Val Pro Ser Asn Glu Gln His Asn Gly Leu Lys Thr Leu Gly Thr Leu
385 390 395 400

Phe Ser Ser Met Met Phe Pro Asp Arg Ala Pro Ser Asp Val Tyr Leu
405 410 415

Tyr Thr Thr Phe Val Gly Gly Ser Arg Asn Arg Glu Leu Ala Lys Ala
420 425 430

Ser Thr Asp Glu Leu Lys Gln Ile Val Ser Ser Asp Leu Gln Gln Leu
435 440 445

Leu Gly Thr Glu Gly Glu Pro Thr Phe Val Asn His Phe Tyr Trp Ser
450 455 460

Lys Ala Phe Pro Leu Tyr Gly Arg Asn Tyr Asp Ser Val Leu Arg Ala
465 470 475 480

Ile Glu Lys Met Glu Arg Asp Leu Pro Gly Leu Phe Tyr Ala Gly Asn
485 490 495

His Lys Gly Gly Leu Ser Val Gly Lys Ser Ile Ala Ser Gly Tyr Lys
500 505 510

Ala Ala Glu Leu Ala Ile Ser Tyr Leu Glu Ser Asn Lys Met Thr Glu
515 520 525

Glu Thr Ile
530

<210> 19
<211> 1674
<212> DNA
<213> Solanum tuberosum

<400> 19
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acggtggaaa gagatggata cttatgggaa gaaggtccta acagtttcca gccttcggat 420

eol f-seql

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ctattccttg ggggtaatta tgtgtctggt gtagcattgg gaaggtgtgt tgaaggtgct 1620
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<210> 20
<211> 557
<212> PRT
<213> Solanum tuberosum

<400> 20

Met Thr Thr Thr Ala Val Ala Asn His Pro Ser Ile Phe Thr His Arg
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Ser Pro Leu Pro Ser Pro Ser Ser Ser Ser Ser Pro Ser Phe Leu
20 25 30

Phe Leu Asn Arg Thr Asn Phe Ile Pro Tyr Phe Ser Thr Ser Lys Arg
35 40 45

Asn Ser Val Asn Cys Asn Gly Trp Arg Thr Arg Cys Ser Val Ala Lys
50 55 60

Asp Tyr Thr Val Pro Pro Ser Glu Val Asp Gly Asn Glu Phe Pro Glu
65 70 75 80

eof-seql

Leu Asp Cys Val Val Val Gly Ala Gly Ile Ser Gly Leu Cys Ile Ala
 85 90 95
 Lys Val Ile Ser Ala Asn Tyr Pro Asn Leu Met Val Thr Glu Ala Arg
 100 105 110
 Asp Arg Ala Gly Gly Asn Ile Thr Thr Val Glu Arg Asp Gly Tyr Leu
 115 120 125
 Trp Glu Glu Gly Pro Asn Ser Phe Gln Pro Ser Asp Pro Met Leu Thr
 130 135 140
 Met Ala Val Asp Cys Gly Leu Lys Asp Asp Leu Val Leu Gly Asp Pro
 145 150 155 160
 Asp Ala Pro Arg Phe Val Leu Trp Lys Asp Lys Leu Arg Pro Val Pro
 165 170 175
 Gly Lys Leu Thr Asp Leu Pro Phe Phe Asp Leu Met Ser Ile Pro Gly
 180 185 190
 Lys Leu Arg Ala Gly Phe Gly Ala Ile Gly Leu Arg Pro Ser Pro Pro
 195 200 205
 Gly Tyr Glu Glu Ser Val Glu Gln Phe Val Arg Arg Asn Leu Gly Ala
 210 215 220
 Glu Val Phe Glu Arg Leu Ile Glu Pro Phe Cys Ser Gly Val Tyr Ala
 225 230 235 240
 Gly Asp Pro Ser Lys Leu Ile Met Lys Ala Ala Phe Gly Lys Val Trp
 245 250 255
 Lys Leu Glu Gln Thr Gly Gly Ser Ile Ile Gly Gly Thr Phe Lys Ala
 260 265 270
 Ile Lys Glu Arg Ser Ser Asn Pro Lys Pro Pro Arg Asp Pro Arg Leu
 275 280 285
 Pro Thr Pro Lys Gly Gln Thr Val Gly Ser Phe Arg Lys Gly Leu Arg
 290 295 300
 Met Leu Pro Asp Ala Ile Cys Glu Arg Leu Gly Ser Lys Val Lys Leu
 305 310 315 320
 Ser Trp Lys Leu Ser Ser Ile Thr Lys Ser Glu Lys Gly Gly Tyr Leu
 325 330 335
 Leu Thr Tyr Glu Thr Pro Glu Gly Val Val Ser Leu Arg Ser Arg Ser
 340 345 350

eof-seql

Ile Val Met Thr Val Pro Ser Tyr Val Ala Ser Asn Ile Leu Arg Pro
355 360 365

Leu Ser Val Ala Ala Ala Asp Ala Leu Ser Ser Phe Tyr Tyr Pro Pro
370 375 380

Val Ala Ala Val Thr Ile Ser Tyr Pro Gln Glu Ala Ile Arg Asp Glu
385 390 395 400

Arg Leu Val Asp Gly Glu Leu Lys Gly Phe Gly Gln Leu His Pro Arg
405 410 415

Ser Gln Gly Val Glu Thr Leu Gly Thr Ile Tyr Ser Ser Ser Leu Phe
420 425 430

Pro Asn Arg Ala Pro Asn Gly Arg Val Leu Leu Leu Asn Tyr Ile Gly
435 440 445

Gly Ala Thr Asn Thr Glu Ile Val Ser Lys Thr Glu Ser Gln Leu Val
450 455 460

Glu Ala Val Asp Arg Asp Leu Arg Lys Met Leu Ile Lys Pro Lys Ala
465 470 475 480

Gln Asp Pro Phe Val Thr Gly Val Arg Val Trp Pro Gln Ala Ile Pro
485 490 495

Gln Phe Leu Val Gly His Leu Asp Thr Leu Gly Thr Ala Lys Thr Ala
500 505 510

Leu Ser Asp Asn Gly Leu Asp Gly Leu Phe Leu Gly Gly Asn Tyr Val
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Ser Gly Val Ala Leu Gly Arg Cys Val Glu Gly Ala Tyr Glu Ile Ala
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Ser Glu Val Thr Gly Phe Leu Ser Gln Tyr Ala Tyr Lys
545 550 555

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<212> DNA
<213> Zea mays

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ggcggaggca tcagtggcct ctgcaccgcg caggcgctgg ccacgcggca cggcgtcggg 240

eol f-seql

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 <211> 535
 <212> PRT
 <213> Zea mays

<400> 22

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Val Arg Cys Ala Ala Val Ala Gly Gly Ala Ala Glu Ala Pro Ala Ser
 35 40 45

Thr Gly Ala Arg Leu Ser Ala Asp Cys Val Val Val Gly Gly Gly Ile
 50 55 60

eof-seql

Ser Gly Leu Cys Thr Ala Gln Ala Leu Ala Thr Arg His Gly Val Gly
65 70 75 80

Asp Val Leu Val Thr Glu Ala Arg Ala Arg Pro Gly Gly Asn Ile Thr
85 90 95

Thr Val Glu Arg Pro Glu Glu Gly Tyr Leu Trp Glu Glu Gly Pro Asn
100 105 110

Ser Phe Gln Pro Ser Asp Pro Val Leu Thr Met Ala Val Asp Ser Gly
115 120 125

Leu Lys Asp Asp Leu Val Phe Gly Asp Pro Asn Ala Pro Arg Phe Val
130 135 140

Leu Trp Glu Gly Lys Leu Arg Pro Val Pro Ser Lys Pro Ala Asp Leu
145 150 155 160

Pro Phe Phe Asp Leu Met Ser Ile Pro Gly Lys Leu Arg Ala Gly Leu
165 170 175

Gly Ala Leu Gly Ile Arg Pro Pro Pro Gly Arg Glu Glu Ser Val
180 185 190

Glu Glu Phe Val Arg Arg Asn Leu Gly Ala Glu Val Phe Glu Arg Leu
195 200 205

Ile Glu Pro Phe Cys Ser Gly Val Tyr Ala Gly Asp Pro Ser Lys Leu
210 215 220

Ser Met Lys Ala Ala Phe Gly Lys Val Trp Arg Leu Glu Glu Thr Gly
225 230 235 240

Gly Ser Ile Ile Gly Gly Thr Ile Lys Thr Ile Gln Glu Arg Ser Lys
245 250 255

Asn Pro Lys Pro Pro Arg Asp Ala Arg Leu Pro Lys Pro Lys Gly Gln
260 265 270

Thr Val Ala Ser Phe Arg Lys Gly Leu Ala Met Leu Pro Asn Ala Ile
275 280 285

Thr Ser Ser Leu Gly Ser Lys Val Lys Leu Ser Trp Lys Leu Thr Ser
290 295 300

Ile Thr Lys Ser Asp Asp Lys Gly Tyr Val Leu Glu Tyr Glu Thr Pro
305 310 315 320

Glu Gly Val Val Ser Val Gln Ala Lys Ser Val Ile Met Thr Ile Pro
325 330 335

eof-seq1

Ser Tyr Val Ala Ser Asn Ile Leu Arg Pro Leu Ser Ser Asp Ala Ala
340 345 350

Asp Ala Leu Ser Arg Phe Tyr Tyr Pro Pro Val Ala Ala Val Thr Val
355 360 365

Ser Tyr Pro Lys Glu Ala Ile Arg Lys Glu Cys Leu Ile Asp Gly Glu
370 375 380

Leu Gln Gly Phe Gly Gln Leu His Pro Arg Ser Gln Gly Val Glu Thr
385 390 400

Leu Gly Thr Ile Tyr Ser Ser Ser Leu Phe Pro Asn Arg Ala Pro Asp
405 410 415

Gly Arg Val Leu Leu Leu Asn Tyr Ile Gly Gly Ala Thr Asn Thr Gly
420 425 430

Ile Val Ser Lys Thr Glu Ser Glu Leu Val Glu Ala Val Asp Arg Asp
435 440 445

Leu Arg Lys Met Leu Ile Asn Ser Thr Ala Val Asp Pro Leu Val Leu
450 455 460

Gly Val Arg Val Trp Pro Gln Ala Ile Pro Gln Phe Leu Val Gly His
465 470 475 480

Leu Asp Leu Leu Glu Ala Ala Lys Ala Ala Leu Asp Arg Gly Gly Tyr
485 490 495

Asp Gly Leu Phe Leu Gly Gly Asn Tyr Val Ala Gly Val Ala Leu Gly
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Arg Cys Val Glu Gly Ala Tyr Glu Ser Ala Ser Gln Ile Ser Asp Phe
515 520 525

Leu Thr Lys Tyr Ala Tyr Lys
530 535

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<212> DNA
<213> Zea mays

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eol f-seql

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 <212> PRT
 <213> Zea mays

<400> 24

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Ala Met Ala Gly Ser Asp Asp Pro Arg Ala Ala Pro Ala Arg Ser Val
 35 40 45

eol f-seql

Ala Val Val Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Arg Leu
50 55 60

Arg Gln Ser Gly Val Asn Val Thr Val Phe Glu Ala Ala Asp Arg Ala
65 70 75 80

Gly Gly Lys Ile Arg Thr Asn Ser Glu Gly Gly Phe Val Trp Asp Glu
85 90 95

Gly Ala Asn Thr Met Thr Glu Gly Glu Trp Glu Ala Ser Arg Leu Ile
100 105 110

Asp Asp Leu Gly Leu Gln Asp Lys Gln Gln Tyr Pro Asn Ser Gln His
115 120 125

Lys Arg Tyr Ile Val Lys Asp Gly Ala Pro Ala Leu Ile Pro Ser Asp
130 135 140

Pro Ile Ser Leu Met Lys Ser Ser Val Leu Ser Thr Lys Ser Lys Ile
145 150 155 160

Ala Leu Phe Phe Glu Pro Phe Leu Tyr Lys Lys Ala Asn Thr Arg Asn
165 170 175

Ser Gly Lys Val Ser Glu Glu His Leu Ser Glu Ser Val Gly Ser Phe
180 185 190

Cys Glu Arg His Phe Gly Arg Glu Val Val Asp Tyr Phe Val Asp Pro
195 200 205

Phe Val Ala Gly Thr Ser Ala Gly Asp Pro Glu Ser Leu Ser Ile Arg
210 215 220

His Ala Phe Pro Ala Leu Trp Asn Leu Glu Arg Lys Tyr Gly Ser Val
225 230 235 240

Ile Val Gly Ala Ile Leu Ser Lys Leu Ala Ala Lys Gly Asp Pro Val
245 250 255

Lys Thr Arg His Asp Ser Ser Gly Lys Arg Arg Asn Arg Arg Val Ser
260 265 270

Phe Ser Phe His Gly Gly Met Gln Ser Leu Ile Asn Ala Leu His Asn
275 280 285

Glu Val Gly Asp Asp Asn Val Lys Leu Gly Thr Glu Val Leu Ser Leu
290 295 300

Ala Cys Thr Phe Asp Gly Val Pro Ala Leu Gly Arg Trp Ser Ile Ser
305 310 315 320

eol f-seql

Val Asp Ser Lys Asp₃₂₅ Ser Gly Asp Lys Asp₃₃₀ Leu Ala Ser Asn Gln Thr₃₃₅

Phe Asp Ala Val₃₄₀ Ile Met Thr Ala Pro₃₄₅ Leu Ser Asn Val Arg₃₅₀ Arg Met

Lys Phe Thr₃₅₅ Lys Gly Gly Ala Pro₃₆₀ Val Val Leu Asp Phe₃₆₅ Leu Pro Lys

Met Asp₃₇₀ Tyr Leu Pro Leu Ser₃₇₅ Leu Met Val Thr Ala₃₈₀ Phe Lys Lys Asp

Asp₃₈₅ Val Lys Lys Pro Leu₃₉₀ Glu Gly Phe Gly Val₃₉₅ Leu Ile Pro Tyr Lys₄₀₀

Glu Gln Gln Lys His₄₀₅ Gly Leu Lys Thr Leu₄₁₀ Gly Thr Leu Phe Ser₄₁₅ Ser

Met Met Phe Pro₄₂₀ Asp Arg Ala Pro Asp₄₂₅ Asp Gln Tyr Leu Tyr₄₃₀ Thr Thr

Phe Val Gly₄₃₅ Gly Ser His Asn Arg₄₄₀ Asp Leu Ala Gly Ala₄₄₅ Pro Thr Ser

Ile Leu₄₅₀ Lys Gln Leu Val Thr₄₅₅ Ser Asp Leu Lys Lys₄₆₀ Leu Leu Gly Val

Glu Gly₄₆₅ Gln Pro Thr Phe Val₄₇₀ Lys His Val Tyr₄₇₅ Trp Gly Asn Ala Phe₄₈₀

Pro Leu Tyr Gly₄₈₅ His Asp Tyr Ser Ser Val₄₉₀ Leu Glu Ala Ile Glu₄₉₅ Lys

Met Glu Lys Asn₅₀₀ Leu Pro Gly Phe Phe₅₀₅ Tyr Ala Gly Asn Ser₅₁₀ Lys Asp

Gly Leu Ala₅₁₅ Val Gly Ser Val Ile₅₂₀ Ala Ser Gly Ser Lys₅₂₅ Ala Ala Asp

Leu Ala₅₃₀ Ile Ser Tyr Leu Glu₅₃₅ Ser His Thr Lys His₅₄₀ Asn Asn Ser His

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 <211> 1692
 <212> DNA
 <213> Chlamydomonas reinhardtii

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eol f-seql

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 <211> 563
 <212> PRT
 <213> Chl amydomonas

<400> 26

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eol f-seq1

Pro Thr Pro Phe Ser Val Ala Ser Pro Ala Thr Ala Ala Ser Pro Ala
35 40 45

Thr Ala Ala Ala Arg Arg Thr Leu His Arg Thr Ala Ala Ala Ala Thr
50 55 60

Gly Ala Pro Thr Ala Ser Gly Ala Gly Val Ala Lys Thr Leu Asp Asn
65 70 75 80

Val Tyr Asp Val Ile Val Val Gly Gly Gly Leu Ser Gly Leu Val Thr
85 90 95

Gly Gln Ala Leu Ala Ala Gln His Lys Ile Gln Asn Phe Leu Val Thr
100 105 110

Glu Ala Arg Glu Arg Val Gly Gly Asn Ile Thr Ser Met Ser Gly Asp
115 120 125

Gly Tyr Val Trp Glu Glu Gly Pro Asn Ser Phe Gln Pro Asn Asp Ser
130 135 140

Met Leu Gln Ile Ala Val Asp Ser Gly Cys Glu Lys Asp Leu Val Phe
145 150 155 160

Gly Asp Pro Thr Ala Pro Arg Phe Val Trp Trp Glu Gly Lys Leu Arg
165 170 175

Pro Val Pro Ser Gly Leu Asp Ala Phe Thr Phe Asp Leu Met Ser Ile
180 185 190

Pro Gly Lys Ile Arg Ala Gly Leu Gly Ala Ile Gly Leu Ile Asn Gly
195 200 205

Ala Met Pro Ser Phe Glu Glu Ser Val Glu Gln Phe Ile Arg Arg Asn
210 215 220

Leu Gly Asp Glu Val Phe Phe Arg Leu Ile Glu Pro Phe Cys Ser Gly
225 230 235 240

Val Tyr Ala Gly Asp Pro Ser Lys Leu Ser Met Lys Ala Ala Phe Asn
245 250 255

Arg Ile Trp Ile Leu Glu Lys Asn Gly Gly Ser Leu Val Gly Gly Ala
260 265 270

Ile Lys Leu Phe Gln Glu Arg Gln Ser Asn Pro Ala Pro Pro Arg Asp
275 280 285

Pro Arg Leu Pro Pro Lys Pro Lys Gly Gln Thr Val Gly Ser Phe Arg
290 295 300

Lys Gly Leu Lys Met Leu Pro Asp Ala Ile Glu Arg Asn Ile Pro Asp
 305 310 315 320

Lys Ile Arg Val Asn Trp Lys Leu Val Ser Leu Gly Arg Glu Ala Asp
 325 330 335

Gly Arg Tyr Gly Leu Val Tyr Asp Thr Pro Glu Gly Arg Val Lys Val
 340 345 350

Phe Ala Arg Ala Val Ala Leu Thr Ala Pro Ser Tyr Val Val Ala Asp
 355 360 365

Leu Val Lys Glu Gln Ala Pro Ala Ala Ala Glu Ala Leu Gly Ser Phe
 370 375 380

Asp Tyr Pro Pro Val Gly Ala Val Thr Leu Ser Tyr Pro Leu Ser Ala
 385 390 395 400

Val Arg Glu Glu Arg Lys Ala Ser Asp Gly Ser Val Pro Gly Phe Gly
 405 410 415

Gln Leu His Pro Arg Thr Gln Gly Ile Thr Thr Leu Gly Thr Ile Tyr
 420 425 430

Ser Ser Ser Leu Phe Pro Gly Arg Ala Pro Glu Gly His Met Leu Leu
 435 440 445

Leu Asn Tyr Ile Gly Gly Thr Thr Asn Arg Gly Ile Val Asn Gln Thr
 450 455 460

Thr Glu Gln Leu Val Glu Gln Val Asp Lys Asp Leu Arg Asn Met Val
 465 470 475 480

Ile Lys Pro Asp Ala Pro Lys Pro Arg Val Val Gly Val Arg Val Trp
 485 490 495

Pro Arg Ala Ile Pro Gln Phe Asn Leu Gly His Leu Glu Gln Leu Asp
 500 505 510

Lys Ala Arg Lys Ala Leu Asp Ala Ala Gly Leu Gln Gly Val His Leu
 515 520 525

Gly Gly Asn Tyr Val Ser Gly Val Ala Leu Gly Lys Val Val Glu His
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Gly Tyr Glu Ser Ala Ala Asn Leu Ala Lys Ser Val Ser Lys Ala Ala
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Val Lys Ala

eol f-seql

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 <211> 1734
 <212> DNA
 <213> Polytomella sp

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 <211> 577
 <212> PRT

eof-seql

<213> Polytomella

<400> 28

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20      25      30

Leu  Asn  Phe  Ser  Thr  His  Ser  Pro  Phe  Asp  Ser  Thr  Tyr  Asp  Val  Val
35      40      45

Val  Val  Gly  Ala  Gly  Ile  Ser  Gly  Leu  Ser  Thr  Ala  Gln  Ala  Leu  Ser
50      55      60

Ile  Gln  His  Lys  Ile  Asp  Asn  Val  Leu  Val  Thr  Glu  Ala  Asp  His  Arg
65      70      75      80

Val  Gly  Gly  Lys  Ile  Thr  Thr  Lys  Arg  Asn  Lys  Asp  Phe  Leu  Trp  Glu
85      90      95

Glu  Gly  Pro  Asn  Ser  Cys  Leu  Met  Asn  Asp  Ala  Leu  Tyr  Arg  Ala  Ala
100     105     110

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Pro  Arg  Trp  Ile  Leu  Trp  Gly  Arg  Arg  Leu  Arg  Val  Ala  Pro  Ile  Gly
130     135     140

Ser  Tyr  Ala  Leu  Lys  Ser  Asp  Leu  Leu  Ser  Thr  Gln  Gly  Leu  Leu  Arg
145     150     155     160

Ala  Ile  Arg  Gly  Val  Thr  Gly  Phe  Gly  Val  Ser  Pro  Ala  Pro  Pro  Lys
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Gly  Gln  Glu  Glu  Ser  Val  Glu  Gly  Phe  Val  Arg  Arg  Thr  Leu  Gly  Asp
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Glu  Ile  Phe  Glu  Arg  Leu  Val  Glu  Pro  Phe  Cys  Ser  Gly  Val  Tyr  Ala
195     200     205

Gly  Asp  Pro  Ser  Lys  Leu  Ser  Met  Arg  Ala  Ala  Phe  Gly  Lys  Leu  Val
210     215     220

Glu  Phe  Glu  Glu  Thr  Gly  Asp  Gly  Ser  Leu  Leu  Arg  Gly  Val  Phe  Arg
225     230     235     240

Tyr  Val  Met  Asn  Lys  Arg  Arg  Glu  Arg  Arg  Thr  Gly  Gly  Ala  Lys  Asp
245     250     255

```

eol f-seql

Gly Asp Thr Val₂₆₀ Pro Leu Asn Glu Thr₂₆₅ Ala Lys Ala Pro Lys₂₇₀ Ser Ser

Ser Gly Pro₂₇₅ Thr Val Ser Ser Phe₂₈₀ Glu Gly Gly Ile Glu₂₈₅ Ile Leu Pro

Lys Ala₂₉₀ Ile Ala Gln Lys Leu₂₉₅ Gly Asp Arg Val Arg₃₀₀ Leu Gly Leu Arg

Leu₃₀₅ Val Arg Ile Asp Pro₃₁₀ Thr Gln Leu Ala Asp₃₁₅ Gly Thr Thr Ala Tyr₃₂₀

Arg Leu Ser Tyr Arg₃₂₅ Arg Met Ser His Gln₃₃₀ Gly Asp Asp Asp Ser₃₃₅ Ser

Arg Thr Ala Gly₃₄₀ Ala Val Pro Arg Thr₃₄₅ Ala Glu Gly Asp Val₃₅₀ Ala Ala

Gly Asp Glu₃₅₅ Asp Ala Val Val Glu₃₆₀ Val Val Ala Lys Lys₃₆₅ Val Val Leu

Thr Thr₃₇₀ Pro Ala Phe Asp Ala₃₇₅ Ala Asp Ile Leu Ser₃₈₀ Arg Ser Gly Leu

Val₃₈₅ Ala Ala Ala Asn Pro₃₉₀ Leu Lys Glu Val Asp₃₉₅ Tyr Pro Pro Val Ala₄₀₀

Leu Val Val Leu Ser₄₀₅ Tyr Asp Val Asp Ser₄₁₀ Ile Ser Ala Ile His Arg₄₁₅

Val Ser His Val₄₂₀ Ala His Gly Leu Ser₄₂₅ Gly Phe Gly Gln Leu₄₃₀ His Pro

Arg Pro Glu₄₃₅ Gly Leu Arg Thr Leu₄₄₀ Gly Thr Ile Tyr Gly₄₄₅ Ser Thr Leu

Phe Pro Asn Arg Ser Pro Val₄₅₅ Ala Arg Thr Thr Leu₄₆₀ Leu Asn Phe Val

Gly₄₆₅ Gly Ser Thr Asp Arg₄₇₀ Ala Val Gly Ser Ala₄₇₅ Asp Pro Met Ala Leu₄₈₀

Ala Met Glu Val Asp₄₈₅ Leu Asp Leu Lys Lys₄₉₀ Ser Gly Leu Ile Arg₄₉₅ Glu

Gly Ala Ala Lys₅₀₀ Pro Glu Val Leu Gly₅₀₅ Val Lys Val Tyr Pro Lys Ala

Ile Pro Gln₅₁₅ Phe Asp Ile Gly His₅₂₀ Leu Asp Arg Val Glu₅₂₅ Lys Ala Lys

eol f-seql

Met Met Leu Lys Asn Glu Arg Gly Gly Ala Asp Trp Ser Gly Val Lys
530 535 540

Leu Ala Gly Asn Tyr Val Cys Gly Val Ala Val Gly Arg Cys Ile Glu
545 550 555 560

Phe Gly Phe Glu Ile Ala Glu Asn Leu Ala Gln Glu Leu Ala Arg Lys
565 570 575

Lys

<210> 29
<211> 1635
<212> DNA
<213> Sorghum bicolor

<400> 29
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cgcgagctc ccgccaggctc ggtcgccgctc gtcggcgccg gggtcagcgg gctcgtggcg 180
gcgtacaggc tcaggaagag cggcgtgaat gtgacggtgt tcgaggcggc cgacagggcg 240
ggaggaaaga tacggaccaa ttccgagggc gggtttctct gggatgaagg agcgaacacc 300
atgacagaag gtgaattgga ggccagtaga ctgatagatg atctcgggtct acaagacaaa 360
cagcagtatc ctaactccca acacaagcgt tacattgtca aagatggagc accagcactg 420
attccttcgg atcccatttc gctgatgaaa agcagtgttc tttctacaaa atcaaagatt 480
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tctgatgagc atttgagtga gagtggtggg agcttctttg aacgccactt cggaagagaa 600
gttggtgact atcttattga tccatttgta gctggaacaa gtgcaggaga tccagagtca 660
ctatctatatt gtcatgcatt cccagcactg tggaatttgg aaagaaaata tggttcagtt 720
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gattcatcag cgaaaagaag gaatagacgc gtgtcgtttt catttcatgg tggaatgcag 840
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gtgttgatcat tggcgtgtac attagatgga gccctgcac caggcgggtg gtcaatttct 960
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gatcgagctc ctgacgacca atatttatat acaacatttg ttgggggtag ccacaataga 1320
gatcttgctg gagctccaac gtctattctg aaacaacttg tgacctctga ccttaaaaaa 1380

eol f-seql

ctcttaggcg tacaggggca accaactttt gtcaagcata tatactgggg aaatgctttt	1440
cctttgtatg gtcattgatta caattctgta ttggaagcta tagaaaagat ggagaaaaat	1500
cttccagggt tcttctacgc aggaataaac aaggatgggc ttgctgttgg gagggtttata	1560
gcttcaggaa gcaaggctgc tgaccttgca atctcgtatc ttgaatctca caccaagcat	1620
aataatttac attga	1635

<210> 30
 <211> 544
 <212> PRT
 <213> Sorghum

<400> 30

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Pro Tyr Arg Pro Thr Ser Ala Arg Ser Leu Arg Leu Arg Pro Val Leu
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Ala Met Ala Gly Ser Asp Asp Ser Arg Ala Ala Pro Ala Arg Ser Val
 35 40 45

Ala Val Val Gly Ala Gly Val Ser Gly Leu Val Ala Ala Tyr Arg Leu
 50 55 60

Arg Lys Ser Gly Val Asn Val Thr Val Phe Glu Ala Ala Asp Arg Ala
 65 70 75 80

Gly Gly Lys Ile Arg Thr Asn Ser Glu Gly Gly Phe Leu Trp Asp Glu
 85 90 95

Gly Ala Asn Thr Met Thr Glu Gly Glu Leu Glu Ala Ser Arg Leu Ile
 100 105 110

Asp Asp Leu Gly Leu Gln Asp Lys Gln Gln Tyr Pro Asn Ser Gln His
 115 120 125

Lys Arg Tyr Ile Val Lys Asp Gly Ala Pro Ala Leu Ile Pro Ser Asp
 130 135 140

Pro Ile Ser Leu Met Lys Ser Ser Val Leu Ser Thr Lys Ser Lys Ile
 145 150 155 160

Ala Leu Phe Phe Glu Pro Phe Leu Tyr Lys Lys Ala Asn Thr Arg Asn
 165 170 175

Pro Gly Lys Val Ser Asp Glu His Leu Ser Glu Ser Val Gly Ser Phe
 180 185 190

Phe Glu Arg His Phe Gly Arg Glu Val Val Asp Tyr Leu Ile Asp Pro
 195 200 205

eof-seql

Phe Val Ala Gly Thr Ser Ala Gly Asp Pro Glu Ser Leu Ser Ile Cys
 210 215 220
 His Ala Phe Pro Ala Leu Trp Asn Leu Glu Arg Lys Tyr Gly Ser Val
 225 230 235 240
 Val Val Gly Ala Ile Leu Ser Lys Leu Thr Ala Lys Gly Asp Pro Val
 245 250 255
 Lys Thr Arg Arg Asp Ser Ser Ala Lys Arg Arg Asn Arg Arg Val Ser
 260 265 270
 Phe Ser Phe His Gly Gly Met Gln Ser Leu Ile Asn Ala Leu His Asn
 275 280 285
 Glu Val Gly Asp Asp Asn Val Lys Leu Gly Thr Glu Val Leu Ser Leu
 290 295 300
 Ala Cys Thr Leu Asp Gly Ala Pro Ala Pro Gly Gly Trp Ser Ile Ser
 305 310 315 320
 Asp Asp Ser Lys Asp Ala Ser Gly Lys Asp Leu Ala Lys Asn Gln Thr
 325 330 335
 Phe Asp Ala Val Ile Met Thr Ala Pro Leu Ser Asn Val Gln Arg Met
 340 345 350
 Lys Phe Thr Lys Gly Gly Ala Pro Phe Val Leu Asp Phe Leu Pro Lys
 355 360 365
 Val Asp Tyr Leu Pro Leu Ser Leu Met Val Thr Ala Phe Lys Lys Glu
 370 375 380
 Asp Val Lys Lys Pro Leu Glu Gly Phe Gly Val Leu Ile Pro Tyr Lys
 385 390 395 400
 Glu Gln Gln Lys His Gly Leu Lys Thr Leu Gly Thr Leu Phe Ser Ser
 405 410 415
 Met Met Phe Pro Asp Arg Ala Pro Asp Asp Gln Tyr Leu Tyr Thr Thr
 420 425 430
 Phe Val Gly Gly Ser His Asn Arg Asp Leu Ala Gly Ala Pro Thr Ser
 435 440 445
 Ile Leu Lys Gln Leu Val Thr Ser Asp Leu Lys Lys Leu Leu Gly Val
 450 455 460
 Gln Gly Gln Pro Thr Phe Val Lys His Ile Tyr Trp Gly Asn Ala Phe
 465 470 475 480

eol f-seql

Pro Leu Tyr Gly His Asp Tyr Asn Ser Val Leu Glu Ala Ile Glu Lys
485 490 495

Met Glu Lys Asn Leu Pro Gly Phe Phe Tyr Ala Gly Asn Asn Lys Asp
500 505 510

Gly Leu Ala Val Gly Ser Val Ile Ala Ser Gly Ser Lys Ala Ala Asp
515 520 525

Leu Ala Ile Ser Tyr Leu Glu Ser His Thr Lys His Asn Asn Leu His
530 535 540

<210> 31
<211> 1611
<212> DNA
<213> Oryza sativa

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agcggcgcgg ccgaggcgcc cgcggcgccc ggggcgcggg tgcggcgga ctgcgtcgtg 180
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ggcgacgtgc tcgtcacgga ggcccgcc cgccccggcg gcaacatcac caccgccgag 300
cgcgccggcg agggctacct ctgggaggag gggccaaca gcttcagcc ttccgacccc 360
gtcctacca tggccgtgga cagcgggctc aaggacgatc tcgtgttcgg ggaccccaac 420
gcgccgcggt tcgtgctgtg ggaggggaag ctaaggccgg tgccgtccaa gcccggcgac 480
ctgccgttct tcgacctcat gagcatcccc ggcaagctca gggccggcct tggcgcgctc 540
ggcgttcgag cgccacctcc agggcgtgag gagtcggtgg aggacttcgt gcggcgcaac 600
ctcggcgcgg aggtctttga gcgcctcatt gagcctttct gctcaggtgt gtatgctggt 660
gaccttcaa agctcagtat gaaggctgca tttgggaagg tgtggaggct ggaggatact 720
ggaggtagca ttattggtgg aaccatcaaa acaatccagg agagggggaa aaaccccaaa 780
ccgccgaggg atccccgcct tccaacgcca aaggggcaga cagttgcatc tttcaggaag 840
ggcttgacta tgctcccgga tgctattaca tctaggttgg gtagcaaagt caaactttca 900
tggaagtga caagcattac aaagtcagac aacaaaggat atgcattagt gtatgaaaca 960
ccagaagggg tggctcgggt gcaagctaaa actgttgtca tgaccatccc atcatatggt 1020
gctagtata tcttgccggc actttcaagt gatgcagcag atgctctgtc aatattctat 1080
tatccaccag ttgctgctgt aactgtttca tatccaaaag aagcaattag aaaagaatgc 1140
ttaattgacg gagagctcca gggtttcggc cagctgcatc cgcgtagtca gggagttgag 1200
actttaggaa caatatatag ctcatcactc tttccaaatc gtgctccagc tggaaggggtg 1260
ttacttctga actacatagg aggttctaca aatacagggg ttgtttccaa gactgaaagt 1320

eof-seq1

gagctggtag aagcagttga ccgtgacctc aggaagatgc tgataaatcc taaagcagtg 1380
gaccctttgg tccttggcgt ccgggtatgg ccacaagcca taccacagtt cctcattggc 1440
catcttgatc atcttgaggc tgcaaatct gccctgggca aaggtgggta tgatggattg 1500
ttcctcggag ggaactatgt tgcaggagtt gccctgggcc gatgcgttga aggtgcatat 1560
gagagtcct cacaatatc tgactacttg accaagtacg cctacaagtg a 1611

<210> 32
<211> 536
<212> PRT
<213> Oryza sativa

<400> 32

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Ala Pro Pro Leu Arg Ile Arg Asp Ala Ala Arg Arg Thr Arg Arg Arg
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Gly His Val Arg Cys Ala Val Ala Ser Gly Ala Ala Glu Ala Pro Ala
35 40 45

Ala Pro Gly Ala Arg Val Ser Ala Asp Cys Val Val Val Gly Gly Gly
50 55 60

Ile Ser Gly Leu Cys Thr Ala Gln Ala Leu Ala Thr Lys His Gly Val
65 70 75 80

Gly Asp Val Leu Val Thr Glu Ala Arg Ala Arg Pro Gly Gly Asn Ile
85 90 95

Thr Thr Ala Glu Arg Ala Gly Glu Gly Tyr Leu Trp Glu Glu Gly Pro
100 105 110

Asn Ser Phe Gln Pro Ser Asp Pro Val Leu Thr Met Ala Val Asp Ser
115 120 125

Gly Leu Lys Asp Asp Leu Val Phe Gly Asp Pro Asn Ala Pro Arg Phe
130 135 140

Val Leu Trp Glu Gly Lys Leu Arg Pro Val Pro Ser Lys Pro Gly Asp
145 150 155 160

Leu Pro Phe Phe Asp Leu Met Ser Ile Pro Gly Lys Leu Arg Ala Gly
165 170 175

Leu Gly Ala Leu Gly Val Arg Ala Pro Pro Pro Gly Arg Glu Glu Ser
180 185 190

Val Glu Asp Phe Val Arg Arg Asn Leu Gly Ala Glu Val Phe Glu Arg
195 200 205

eof-seq1

Leu Ile Glu Pro Phe Cys Ser Gly Val Tyr Ala Gly Asp Pro Ser Lys
 210 215 220
 Leu Ser Met Lys Ala Ala Phe Gly Lys Val Trp Arg Leu Glu Asp Thr
 225 230 235 240
 Gly Gly Ser Ile Ile Gly Gly Thr Ile Lys Thr Ile Gln Glu Arg Gly
 245 250 255
 Lys Asn Pro Lys Pro Pro Arg Asp Pro Arg Leu Pro Thr Pro Lys Gly
 260 265 270
 Gln Thr Val Ala Ser Phe Arg Lys Gly Leu Thr Met Leu Pro Asp Ala
 275 280 285
 Ile Thr Ser Arg Leu Gly Ser Lys Val Lys Leu Ser Trp Lys Leu Thr
 290 295 300
 Ser Ile Thr Lys Ser Asp Asn Lys Gly Tyr Ala Leu Val Tyr Glu Thr
 305 310 315 320
 Pro Glu Gly Val Val Ser Val Gln Ala Lys Thr Val Val Met Thr Ile
 325 330 335
 Pro Ser Tyr Val Ala Ser Asp Ile Leu Arg Pro Leu Ser Ser Asp Ala
 340 345 350
 Ala Asp Ala Leu Ser Ile Phe Tyr Tyr Pro Pro Val Ala Ala Val Thr
 355 360 365
 Val Ser Tyr Pro Lys Glu Ala Ile Arg Lys Glu Cys Leu Ile Asp Gly
 370 375 380
 Glu Leu Gln Gly Phe Gly Gln Leu His Pro Arg Ser Gln Gly Val Glu
 385 390 395 400
 Thr Leu Gly Thr Ile Tyr Ser Ser Ser Leu Phe Pro Asn Arg Ala Pro
 405 410 415
 Ala Gly Arg Val Leu Leu Leu Asn Tyr Ile Gly Gly Ser Thr Asn Thr
 420 425 430
 Gly Ile Val Ser Lys Thr Glu Ser Glu Leu Val Glu Ala Val Asp Arg
 435 440 445
 Asp Leu Arg Lys Met Leu Ile Asn Pro Lys Ala Val Asp Pro Leu Val
 450 455 460
 Leu Gly Val Arg Val Trp Pro Gln Ala Ile Pro Gln Phe Leu Ile Gly
 465 470 475 480

eol f-seql

His Leu Asp His Leu Glu Ala Ala Lys Ser Ala Leu Gly Lys Gly Gly
485 490 495

Tyr Asp Gly Leu Phe Leu Gly Gly Asn Tyr Val Ala Gly Val Ala Leu
500 505 510

Gly Arg Cys Val Glu Gly Ala Tyr Glu Ser Ala Ser Gln Ile Ser Asp
515 520 525

Tyr Leu Thr Lys Tyr Ala Tyr Lys
530 535

<210> 33
<211> 1518
<212> DNA
<213> Amaranthus tuberculatus

<400> 33
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ttgtttgaag ctgattctag agctggaggc aaacttaaaa ctgttaaaaa agatggtttt 180
atttgggatg agggggcaaa tactatgaca gaaagtgagg cagaagtctc gagtttgatc 240
gatgatcttg ggcttcgtga gaagcaacag ttgccaatTT cacaaaataa aagatacata 300
gctagagatg gtcttcctgt gctactacct tcaaatcccg ctgcactgct cacgagcaat 360
atcctttcag caaaatcaaa gctgcaaatt atgttgaac catTTTTctg gagaaaacac 420
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catTTTggga aagagtttgt tgattatgtt attgaccctt ttgttgcggg tacatgtgggt 540
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aggTTTggct ctgtgtttgc tggactaatt caatcaacat tgttatctaa gaaggaaaag 660
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ctccagtgtg aggtgctgtc cttgtcatac aaccagaagg ggatcccttc attagggaaT 840
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gtcactgctc caattcgcaa tgtcaaagaa atgaagatta tgaaattcgg aaatccattt 960
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caacataatg gactgaagac tcttggTact ttattttcct ccatgatgtt tcccgatcgt 1140
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gcaaacgctt caacggatga attgaagcaa atagtttctt ctgaccttca gcagctgttg 1260
ggcactgagg acgaaccttc atttgtcaat catctctttt ggagcaacgc attcccgttg 1320

eof-seql

tatggacaca attacgattc tgttttgaga gccatagaca agatggaaaa ggatcttcct	1380
ggatTTTTTT atgcaggtaa ccataagggt ggactttcag tgggaaaagc gatggcctcc	1440
ggatgcaagg ctgcggaact tgtaatatcc tatctggact ctcatatata tgtgaagatg	1500
gatgagaaga ccgcgtaa	1518

<210> 34
 <211> 505
 <212> PRT
 <213> Amaranthus

<400> 34

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Ala Val Val Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Lys Leu	20 25 30
Lys Ser His Gly Leu Asn Val Thr Leu Phe Glu Ala Asp Ser Arg Ala	35 40 45
Gly Gly Lys Leu Lys Thr Val Lys Lys Asp Gly Phe Ile Trp Asp Glu	50 55 60
Gly Ala Asn Thr Met Thr Glu Ser Glu Ala Glu Val Ser Ser Leu Ile	65 70 75 80
Asp Asp Leu Gly Leu Arg Glu Lys Gln Gln Leu Pro Ile Ser Gln Asn	85 90 95
Lys Arg Tyr Ile Ala Arg Asp Gly Leu Pro Val Leu Leu Pro Ser Asn	100 105 110
Pro Ala Ala Leu Leu Thr Ser Asn Ile Leu Ser Ala Lys Ser Lys Leu	115 120 125
Gln Ile Met Leu Glu Pro Phe Phe Trp Arg Lys His Asn Ala Thr Glu	130 135 140
Leu Ser Asp Glu His Val Gln Glu Ser Val Gly Glu Phe Phe Glu Arg	145 150 155 160
His Phe Gly Lys Glu Phe Val Asp Tyr Val Ile Asp Pro Phe Val Ala	165 170 175
Gly Thr Cys Gly Gly Asp Pro Gln Ser Leu Ser Met His His Thr Phe	180 185 190
Pro Glu Val Trp Asn Ile Glu Lys Arg Phe Gly Ser Val Phe Ala Gly	195 200 205

Leu Ile Gln Ser Thr Leu Leu Ser Lys Lys Glu Lys Gly Gly Gly Gly
 210 215 220
 Asn Ala Ser Ile Lys Lys Pro Arg Val Arg Gly Ser Phe Ser Phe His
 225 230 235 240
 Gly Gly Met Gln Thr Leu Val Asp Thr Ile Cys Lys Gln Leu Gly Glu
 245 250 255
 Asp Glu Leu Lys Leu Gln Cys Glu Val Leu Ser Leu Ser Tyr Asn Gln
 260 265 270
 Lys Gly Ile Pro Ser Leu Gly Asn Trp Ser Val Ser Ser Met Ser Asn
 275 280 285
 Asn Thr Ser Glu Asp Gln Ser Tyr Asp Ala Val Val Val Thr Ala Pro
 290 295 300
 Ile Arg Asn Val Lys Glu Met Lys Ile Met Lys Phe Gly Asn Pro Phe
 305 310 315 320
 Ser Leu Asp Phe Ile Pro Glu Val Ser Tyr Val Pro Leu Ser Val Met
 325 330 335
 Ile Thr Ala Phe Lys Lys Asp Lys Val Lys Arg Pro Leu Glu Gly Phe
 340 345 350
 Gly Val Leu Ile Pro Ser Lys Glu Gln His Asn Gly Leu Lys Thr Leu
 355 360 365
 Gly Thr Leu Phe Ser Ser Met Met Phe Pro Asp Arg Ala Pro Ser Asp
 370 375 380
 Met Cys Leu Phe Thr Thr Phe Val Gly Gly Ser Arg Asn Arg Lys Leu
 385 390 395 400
 Ala Asn Ala Ser Thr Asp Glu Leu Lys Gln Ile Val Ser Ser Asp Leu
 405 410 415
 Gln Gln Leu Leu Gly Thr Glu Asp Glu Pro Ser Phe Val Asn His Leu
 420 425 430
 Phe Trp Ser Asn Ala Phe Pro Leu Tyr Gly His Asn Tyr Asp Ser Val
 435 440 445
 Leu Arg Ala Ile Asp Lys Met Glu Lys Asp Leu Pro Gly Phe Phe Tyr
 450 455 460
 Ala Gly Asn His Lys Gly Gly Leu Ser Val Gly Lys Ala Met Ala Ser
 465 470 475 480

Gly Cys Lys Ala Ala Glu Leu Val Ile Ser Tyr Leu Asp Ser His Ile
 485 490 495

Tyr Val Lys Met Asp Glu Lys Thr Ala
 500 505

<210> 35
 <211> 1521
 <212> DNA
 <213> Arabidopsis thaliana

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 accgtcggat cttcaaaaat cgaaggcggg ggaggcacca ccatcacgac ggattgtgtg 180
 attgtcggcg gaggtattag tggctcttgc atcgtcagg cgcttgctac taagcatcct 240
 gatgctgctc cgaatttaat tgtgaccgag gctaaggatc gtgttgagg caacattatc 300
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 cctatgctca ctatggtggt agatagtggg ttgaaggatg atttggtgtt gggagatcct 420
 actgcgcaa ggtttgtgtt gtggaatggg aaattgaggc cggttccatc gaagctaaca 480
 gacttaccgt tctttgattt gatgagtatt ggtgggaaga ttagagctgg ttttgggtgca 540
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 aacctcgtg atgaggtttt tgagcgcctg attgaaccgt tttgttcagg tgtttatgct 660
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 aatggtggaa gcataatagg tgggtactttt aaggcaattc aggagaggaa aaacgctccc 780
 aaggcagaac gagacccgag cctgccaaaa ccacagggcc aaacagttgg ttctttcagg 840
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 tattaccac cagttgcagc agtatctatc tcgtaccga aagaagcaat ccgaacagaa 1140
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<210> 36

eof-seql

<211> 506
 <212> PRT
 <213> Arabidopsis

<400> 36

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Phe Ser Lys Pro Asn Leu Arg Leu Asn Val Tyr Lys Pro Leu Arg Leu
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Arg Cys Ser Val Ala Gly Gly Pro Thr Val Gly Ser Ser Lys Ile Glu
 35 40 45

Gly Gly Gly Gly Thr Thr Ile Thr Thr Asp Cys Val Ile Val Gly Gly
 50 55 60

Gly Ile Ser Gly Leu Cys Ile Ala Gln Ala Leu Ala Thr Lys His Pro
 65 70 75 80

Asp Ala Ala Pro Asn Leu Ile Val Thr Glu Ala Lys Asp Arg Val Gly
 85 90 95

Gly Asn Ile Ile Thr Arg Glu Glu Asn Gly Phe Leu Trp Glu Glu Gly
 100 105 110

Pro Asn Ser Phe Gln Pro Ser Asp Pro Met Leu Thr Met Val Val Asp
 115 120 125

Ser Gly Leu Lys Asp Asp Leu Val Leu Gly Asp Pro Thr Ala Pro Arg
 130 135 140

Phe Val Leu Trp Asn Gly Lys Leu Arg Pro Val Pro Ser Lys Leu Thr
 145 150 155 160

Asp Leu Pro Phe Phe Asp Leu Met Ser Ile Gly Gly Lys Ile Arg Ala
 165 170 175

Gly Phe Gly Ala Leu Gly Ile Arg Pro Ser Pro Pro Gly Arg Glu Glu
 180 185 190

Ser Val Glu Glu Phe Val Arg Arg Asn Leu Gly Asp Glu Val Phe Glu
 195 200 205

Arg Leu Ile Glu Pro Phe Cys Ser Gly Val Tyr Ala Gly Asp Pro Ser
 210 215 220

Lys Leu Ser Met Lys Ala Ala Phe Gly Lys Val Trp Lys Leu Glu Gln
 225 230 235 240

Asn Gly Gly Ser Ile Ile Gly Gly Thr Phe Lys Ala Ile Gln Glu Arg
 245 250 255

eof-seql

Lys Asn Ala Pro Lys Ala Glu Arg Asp Pro Arg Leu Pro Lys Pro Glu
 260 265 270
 Gly Gln Thr Val Gly Ser Phe Arg Lys Gly Leu Arg Met Leu Pro Glu
 275 280 285
 Ala Ile Ser Ala Arg Leu Gly Ser Lys Val Lys Leu Ser Trp Lys Leu
 290 295 300
 Ser Gly Ile Thr Lys Leu Glu Ser Gly Gly Tyr Asn Leu Thr Tyr Glu
 305 310 315 320
 Thr Pro Asp Gly Leu Val Ser Val Gln Ser Lys Ser Val Val Met Thr
 325 330 335
 Val Pro Ser His Val Ala Ser Gly Leu Leu Arg Pro Leu Ser Glu Ser
 340 345 350
 Ala Ala Asn Ala Leu Ser Lys Leu Tyr Tyr Pro Pro Val Ala Ala Val
 355 360 365
 Ser Ile Ser Tyr Pro Lys Glu Ala Ile Arg Thr Glu Cys Leu Ile Asp
 370 375 380
 Gly Glu Leu Lys Gly Phe Gly Gln Leu His Pro Arg Thr Gln Gly Val
 385 390 395 400
 Glu Thr Leu Gly Thr Ile Tyr Ser Ser Ser Leu Phe Pro Asn Arg Ala
 405 410 415
 Pro Pro Gly Arg Ile Leu Leu Leu Asn Tyr Ile Gly Gly Ser Thr Asn
 420 425 430
 Thr Gly Ile Leu Ser Lys Ser Glu Gly Glu Leu Val Glu Ala Phe Leu
 435 440 445
 Val Gly His Phe Asp Ile Leu Asp Thr Ala Lys Ser Ser Leu Thr Ser
 450 455 460
 Ser Gly Tyr Glu Gly Leu Phe Leu Gly Gly Asn Tyr Val Ala Gly Val
 465 470 475 480
 Ala Leu Gly Arg Cys Val Glu Gly Ala Tyr Glu Thr Ala Ile Glu Val
 485 490 495
 Asn Asn Phe Met Ser Arg Tyr Ala Tyr Lys
 500 505

<210> 37
 <211> 1515

eol f-seql

<212> DNA
<213> Nicotiana tabacum

<400> 37
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gtatttgaag cagaagggaa agctggaggg aagttacgta gcgtgagcca agatggcctg 180
atatgggatg aaggggcaaa tactatgact gaaagtgaag gtgatgttac atttttgatt 240
gattctcttg gactccgaga aaagcaacaa tttccacttt cacaaaacaa gcgctacatt 300
gccagaaatg gtactcctgt actgttacct tcaaatacaa ttgatctgat caaaagcaat 360
tttctttcca ctggatcaaa gcttcagatg cttctggaac caatattatg gaagaataaa 420
aagctctccc aggtgtctga ctcacatgaa agtgtcagtg gattcttcca gcgtcatttt 480
ggaaaggagg ttgttgacta tctaattgac cttttgttg ctggaacgtg tgggtggtgat 540
cctgactcgc ttcaatgca ccattcattt ccagagttgt ggaatttaga gaaaagggtt 600
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gggccaccca aaacttcagc aaataagaag cgccagcggg gatctttttc ctttttgggc 720
ggaatgcaaa cacttactga tgcaatatgc aaagatctca gagaagatga acttagacta 780
aactctagag ttctggaatt atctttagc tgtactgagg actctgcgat agatagctgg 840
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aaaagacatt gctga 1515

<210> 38
<211> 504
<212> PRT
<213> Nicotiana

<400> 38

Met Ala Pro Ser Ala Gly Glu Asp Lys His Ser Ser Ala Lys Arg Val
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eol f-seq1

Ala Val Ile Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Lys Leu
20 25 30

Lys Ile His Gly Leu Asn Val Thr Val Phe Glu Ala Glu Gly Lys Ala
35 40 45

Gly Gly Lys Leu Arg Ser Val Ser Gln Asp Gly Leu Ile Trp Asp Glu
50 55 60

Gly Ala Asn Thr Met Thr Glu Ser Glu Gly Asp Val Thr Phe Leu Ile
65 70 75 80

Asp Ser Leu Gly Leu Arg Glu Lys Gln Gln Phe Pro Leu Ser Gln Asn
85 90 95

Lys Arg Tyr Ile Ala Arg Asn Gly Thr Pro Val Leu Leu Pro Ser Asn
100 105 110

Pro Ile Asp Leu Ile Lys Ser Asn Phe Leu Ser Thr Gly Ser Lys Leu
115 120 125

Gln Met Leu Leu Glu Pro Ile Leu Trp Lys Asn Lys Lys Leu Ser Gln
130 135 140

Val Ser Asp Ser His Glu Ser Val Ser Gly Phe Phe Gln Arg His Phe
145 150 155 160

Gly Lys Glu Val Val Asp Tyr Leu Ile Asp Pro Phe Val Ala Gly Thr
165 170 175

Cys Gly Gly Asp Pro Asp Ser Leu Ser Met His His Ser Phe Pro Glu
180 185 190

Leu Trp Asn Leu Glu Lys Arg Phe Gly Ser Val Ile Leu Gly Ala Ile
195 200 205

Arg Ser Lys Leu Ser Pro Lys Asn Glu Lys Lys Gln Gly Pro Pro Lys
210 215 220

Thr Ser Ala Asn Lys Lys Arg Gln Arg Gly Ser Phe Ser Phe Leu Gly
225 230 235 240

Gly Met Gln Thr Leu Thr Asp Ala Ile Cys Lys Asp Leu Arg Glu Asp
245 250 255

Glu Leu Arg Leu Asn Ser Arg Val Leu Glu Leu Ser Cys Ser Cys Thr
260 265 270

Glu Asp Ser Ala Ile Asp Ser Trp Ser Ile Ile Ser Ala Ser Pro His
275 280 285

eof-seq1

Lys Arg Gln Ser Glu Glu Glu Ser Phe Asp Ala Val Ile Met Thr Ala
290 295 300

Pro Leu Cys Asp Val Lys Ser Met Lys Ile Ala Lys Arg Gly Asn Pro
305 310 315 320

Phe Leu Leu Asn Phe Ile Pro Glu Val Asp Tyr Val Pro Leu Ser Val
325 330 335

Val Ile Thr Thr Phe Lys Arg Glu Asn Val Lys Tyr Pro Leu Glu Gly
340 345 350

Phe Gly Val Leu Val Pro Ser Lys Glu Gln Gln His Gly Leu Lys Thr
355 360 365

Leu Gly Thr Leu Phe Ser Ser Met Met Phe Pro Asp Arg Ala Pro Asn
370 375 380

Asn Val Tyr Leu Tyr Thr Thr Phe Val Gly Gly Ser Arg Asn Arg Glu
385 390 395 400

Leu Ala Lys Ala Ser Arg Thr Glu Leu Lys Glu Ile Val Thr Ser Asp
405 410 415

Leu Lys Gln Leu Leu Gly Ala Glu Gly Glu Pro Thr Tyr Val Asn His
420 425 430

Leu Tyr Trp Ser Lys Ala Phe Pro Leu Tyr Gly His Asn Tyr Asp Ser
435 440 445

Val Leu Asp Ala Ile Asp Lys Met Glu Lys Asn Leu Pro Gly Leu Phe
450 455 460

Tyr Ala Gly Asn His Arg Gly Gly Leu Ser Val Gly Lys Ala Leu Ser
465 470 475 480

Ser Gly Cys Asn Ala Ala Asp Leu Val Ile Ser Tyr Leu Glu Ser Val
485 490 495

Ser Thr Asp Ser Lys Arg His Cys
500

<210> 39

<211> 1509

<212> DNA

<213> Glycine max

<400> 39

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gctggggtaa gtgggcttgc tgcggcttac aaattgaaat cacatggtct ggatgtcact 120

gtatttgaag ctgagggaag agctggaggg aggttgagaa gtgtttctca ggatggtcta 180

eof-seq1

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 gtgaaaaatg gggcaccact tctggtaccc acaaatcctg ctgcactact gaagagtaaa 360
 ctgctttctg cacaatcaaa gatccatctc atttttgaac cttttatgtg gaaaagaagt 420
 gacccctcta atgtgtgtga tgaaaattct gtggaaagtg taggcagggtt ctttgaacgt 480
 cattttggaa aagaggttgt ggactatctg attgatcctt ttgttggggg cactagtgca 540
 gcagatcctg aatctctctc tatgcgccat tctttcccag agctatggaa tttggagaaa 600
 aggtttggct ccattatagc cggggcattg caatctaagt tattcgccaa aagggaaaaa 660
 actggagaaa ataggactgc actaagaaaa aacaaacaca agcgtggttc gttttctttc 720
 cagggtggga tgcagacact gacagataca ttgtgcaaag agcttggcaa agacgacctt 780
 aaattaaatg aaaaggtttt gacattagct tatggtcatg atggaagttc ctcttcacaa 840
 aactggctta ttactagtgc ttctaacca agtacacaag atgttgatgc agtaatcatg 900
 acggctcctc tatataatgt caaggacatc aagatcacia aaaggggaac tccctttcca 960
 cttaattttc ttcccagggt aagctacgtg ccaatctcag tcatgattac taccttcaaa 1020
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 aaaaatggtt taaaaaccct tggtagactt ttttcctcta tgatgttccc agatcgtgca 1140
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 ggacgtaact atgggtcagt tcttcaagca attgataaga tagaaaaaga tcttcccgga 1380
 tttttctttg caggtaacta caaagggtgga ctctcagttg gcaaagcaat agcctcaggc 1440
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 gataaatga 1509

<210> 40
 <211> 502
 <212> PRT
 <213> Glycine
 <400> 40

Met Ala Ser Ser Ala Thr Asp Asp Asn Pro Arg Ser Val Lys Arg Val
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 Ala Val Val Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Lys Leu
 20 25 30
 Lys Ser His Gly Leu Asp Val Thr Val Phe Glu Ala Glu Gly Arg Ala
 35 40 45
 Gly Gly Arg Leu Arg Ser Val Ser Gln Asp Gly Leu Ile Trp Asp Glu
 50 55 60

eof-seql

Gly Ala Asn Thr Met Thr Glu Ser Glu Ile Glu Val Lys Gly Leu Ile
 65 70 75 80
 Asp Ala Leu Gly Leu Gln Glu Lys Gln Gln Phe Pro Ile Ser Gln His
 85 90 95
 Lys Arg Tyr Ile Val Lys Asn Gly Ala Pro Leu Leu Val Pro Thr Asn
 100 105 110
 Pro Ala Ala Leu Leu Lys Ser Lys Leu Leu Ser Ala Gln Ser Lys Ile
 115 120 125
 His Leu Ile Phe Glu Pro Phe Met Trp Lys Arg Ser Asp Pro Ser Asn
 130 135 140
 Val Cys Asp Glu Asn Ser Val Glu Ser Val Gly Arg Phe Phe Glu Arg
 145 150 155 160
 His Phe Gly Lys Glu Val Val Asp Tyr Leu Ile Asp Pro Phe Val Gly
 165 170 175
 Gly Thr Ser Ala Ala Asp Pro Glu Ser Leu Ser Met Arg His Ser Phe
 180 185 190
 Pro Glu Leu Trp Asn Leu Glu Lys Arg Phe Gly Ser Ile Ile Ala Gly
 195 200 205
 Ala Leu Gln Ser Lys Leu Phe Ala Lys Arg Glu Lys Thr Gly Glu Asn
 210 215 220
 Arg Thr Ala Leu Arg Lys Asn Lys His Lys Arg Gly Ser Phe Ser Phe
 225 230 235 240
 Gln Gly Gly Met Gln Thr Leu Thr Asp Thr Leu Cys Lys Glu Leu Gly
 245 250 255
 Lys Asp Asp Leu Lys Leu Asn Glu Lys Val Leu Thr Leu Ala Tyr Gly
 260 265 270
 His Asp Gly Ser Ser Ser Ser Gln Asn Trp Ser Ile Thr Ser Ala Ser
 275 280 285
 Asn Gln Ser Thr Gln Asp Val Asp Ala Val Ile Met Thr Ala Pro Leu
 290 295 300
 Tyr Asn Val Lys Asp Ile Lys Ile Thr Lys Arg Gly Thr Pro Phe Pro
 305 310 315 320
 Leu Asn Phe Leu Pro Glu Val Ser Tyr Val Pro Ile Ser Val Met Ile
 325 330 335

eof-seq1

Thr Thr Phe Lys Lys Glu Asn Val Lys Arg Pro Leu Glu Gly Phe Gly
340 345 350

Val Leu Val Pro Ser Lys Glu Gln Lys Asn Gly Leu Lys Thr Leu Gly
355 360 365

Thr Leu Phe Ser Ser Met Met Phe Pro Asp Arg Ala Pro Ser Asp Leu
370 375 380

Tyr Leu Tyr Thr Thr Phe Ile Gly Gly Thr Gln Asn Arg Glu Leu Ala
385 390 395 400

Gln Ala Ser Thr Asp Glu Leu Arg Lys Ile Val Thr Ser Asp Leu Arg
405 410 415

Lys Leu Leu Gly Ala Glu Gly Glu Pro Thr Phe Val Asn His Phe Tyr
420 425 430

Trp Ser Lys Gly Phe Pro Leu Tyr Gly Arg Asn Tyr Gly Ser Val Leu
435 440 445

Gln Ala Ile Asp Lys Ile Glu Lys Asp Leu Pro Gly Phe Phe Phe Ala
450 455 460

Gly Asn Tyr Lys Gly Gly Leu Ser Val Gly Lys Ala Ile Ala Ser Gly
465 470 475 480

Cys Lys Ala Ala Asp Leu Val Ile Ser Tyr Leu Asn Ser Ala Ser Asp
485 490 495

Asn Thr Val Pro Asp Lys
500

<210> 41
<211> 1205
<212> DNA
<213> Cucumis sativus

<400> 41
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ttagttctgg gagaccaga tgcacctcga tttgtattgt ggaatggaaa gctcagacca 120
gtgcctgcga aacctaata tctaccttc tttgacctga tgagcattgg tggaaaaatc 180
agagcaggct ttggtgccct gggcattcgc cctcctcctc caggctcgaga ggaatcagtt 240
gaagaatttg tccgtcggaa ccttggcaat gaagtttttg aacgtttgat agagccattt 300
tgttctggtg tatacgtgg tgacccttca aagctaagca tgaaagcagc ttttggttaag 360
gtttggaggc tagagcaaaa tgggtgtagt attattggtg ggactttcaa agcacttcaa 420
gaaaggaata aaactaccaa accaccaaga gatccgcgtc taccaaagcc taagggccaa 480

eol f-seql

actgttggat cttttcggaa aggacttacc atgttgccaa atgctatttc tacttgtttg 540
 gggagtaaag taaaagtatc ttggaagcta tctagtatca gtaaagtgga tgacggaggt 600
 tatagtttga catacgaac accagaagga ctagtctcca tactaagcag aagtgtcatc 660
 atgacggttc cttcttatat tgctggcact ctgttgcgtc caatctcggg gaaagctgca 720
 gatgcacttt caaaatttta ttatccacca gttgcatcag tgaccatatac atatccaaaa 780
 ggagcaatta ggaaagaatg cttgattgat ggtgaactaa agggggtttg tcaattgcac 840
 cctcgtagcc aggggggtgac tactttggga actatataca gctcatcact ttttcctaata 900
 cgagcgccag atggaaggggt attgctcttg aactacattg gaggggctac taatactgga 960
 attctttctc agacagagag cgagctcata gaagtagttg atcgggattt aagaaaaatc 1020
 ctcataaacc caaacgcaga ggatcctcta ccattgagcg tgaggggtgtg gccacaagcc 1080
 attccacagt tcttgattgg ccattctgat gttctagaca ccgccaaggc cggactgaga 1140
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 agatg 1205

<210> 42
 <211> 401
 <212> PRT
 <213> Cucumi s

<400> 42

Ser Phe Gln Pro Ser Asp Pro Ile Leu Thr Met Val Val Asp Ser Gly
 1 5 10 15

Leu Lys Asp Asp Leu Val Leu Gly Asp Pro Asp Ala Pro Arg Phe Val
 20 25 30

Leu Trp Asn Gly Lys Leu Arg Pro Val Pro Ala Lys Pro Asn Asp Leu
 35 40 45

Pro Phe Phe Asp Leu Met Ser Ile Gly Gly Lys Ile Arg Ala Gly Phe
 50 55 60

Gly Ala Leu Gly Ile Arg Pro Pro Pro Pro Gly Arg Glu Glu Ser Val
 65 70 75 80

Glu Glu Phe Val Arg Arg Asn Leu Gly Asn Glu Val Phe Glu Arg Leu
 85 90 95

Ile Glu Pro Phe Cys Ser Gly Val Tyr Ala Gly Asp Pro Ser Lys Leu
 100 105 110

Ser Met Lys Ala Ala Phe Gly Lys Val Trp Arg Leu Glu Gln Asn Gly
 115 120 125

Gly Ser Ile Ile Gly Gly Thr Phe Lys Ala Leu Gln Glu Arg Asn Lys
 130 135 140

eof-seql

Thr Thr Lys Pro Pro Arg Asp Pro Arg Leu Pro Lys Pro Lys Gly Gln
145 150 155 160

Thr Val Gly Ser Phe Arg Lys Gly Leu Thr Met Leu Pro Asn Ala Ile
165 170 175

Ser Thr Cys Leu Gly Ser Lys Val Lys Val Ser Trp Lys Leu Ser Ser
180 185 190

Ile Ser Lys Val Asp Asp Gly Gly Tyr Ser Leu Thr Tyr Glu Thr Pro
195 200 205

Glu Gly Leu Val Ser Ile Leu Ser Arg Ser Val Ile Met Thr Val Pro
210 215 220

Ser Tyr Ile Ala Gly Thr Leu Leu Arg Pro Ile Ser Gly Lys Ala Ala
225 230 235 240

Asp Ala Leu Ser Lys Phe Tyr Tyr Pro Pro Val Ala Ser Val Thr Ile
245 250 255

Ser Tyr Pro Lys Gly Ala Ile Arg Lys Glu Cys Leu Ile Asp Gly Glu
260 265 270

Leu Lys Gly Phe Gly Gln Leu His Pro Arg Ser Gln Gly Val Thr Thr
275 280 285

Leu Gly Thr Ile Tyr Ser Ser Ser Leu Phe Pro Asn Arg Ala Pro Asp
290 295 300

Gly Arg Val Leu Leu Leu Asn Tyr Ile Gly Gly Ala Thr Asn Thr Gly
305 310 315 320

Ile Leu Ser Gln Thr Glu Ser Glu Leu Ile Glu Val Val Asp Arg Asp
325 330 335

Leu Arg Lys Ile Leu Ile Asn Pro Asn Ala Glu Asp Pro Leu Pro Leu
340 345 350

Ser Val Arg Val Trp Pro Gln Ala Ile Pro Gln Phe Leu Ile Gly His
355 360 365

Leu Asp Val Leu Asp Thr Ala Lys Ala Gly Leu Arg Glu Ala Gly Met
370 375 380

Glu Gly Leu Phe Leu Gly Gly Asn Tyr Val Cys Gly Val Ala Leu Gly
385 390 395 400

Arg

eol f-seql

<210> 43
 <211> 1521
 <212> DNA
 <213> Oryza sativa

<400> 43
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 gcggccgaca gggcggttg gaagatacgg accaactccg agggcggtt catctgggac 180
 gaaggggcca acaccatgac agagagtga tggaggcaa gcaggcttat tgacgatctt 240
 ggctacaag gcaaacagca gtatcctaac tcacaacaca agcgttacat tgtcaaagat 300
 ggagcaccaa cactgattcc ctcatatccc attgcgtca tgaaaagcac tgttctttct 360
 acaaaatcaa agctcaagct atttctggaa ccatttctct atgagaaatc tagcagaagg 420
 acctcgggaa aagtgtctga tgaacattta agtgagagt tgatttttct gtgtatatgt 480
 agagataatc aggttgttga ttatcttatt gatccatttg tggctggaac aagcggagga 540
 gatcctgagt cattatcaat tcgtcatgca tttccagcat tatggaattt ggagaataag 600
 tatggctctg tcattgctgg tgccatcttg tccaaactat ccactaaggg tgattcagtg 660
 aagacaggag gtgcttcgcc agggaaagga aggaataaac gtgtgtcatt ttcatttcat 720
 ggtggaatgc agtactaat agatgcactt cacaatgaag ttggagatgg taacgtgaag 780
 ctgtgtacag aagtgtgtc attggcatgt tgctgtgat gagtctcttc ttctggtggt 840
 tggtaattt ctgttgattc aaaagatgct aaagggaag atctcagaaa gaaccaatct 900
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 ataccctata aggaacagca aaagcatggt ctcaaaaccc ttgggaccct cttctctcgc 1140
 atgatgtttc cagatcgagc tcctaattgat caatatctat atacatcttt cattgggggg 1200
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 gacctaaaga agctcttggg tgttgaggga caacctact ttgtgaagca tgtacattgg 1320
 agaaatgctt ttcctttata tggccagaat tatgatctgg tactggaagc tatagcaaaa 1380
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 tgcacagatc aggacaatta g 1521

<210> 44
 <211> 506
 <212> PRT
 <213> Oryza sativa

<400> 44

eol f-seql

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1 5 10 15

Gly Ala Gly Val Ser Gly Leu Ala Ala Ala Tyr Arg Leu Arg Lys Arg
20 25 30

Gly Val Gln Val Thr Val Phe Glu Ala Ala Asp Arg Ala Gly Gly Lys
35 40 45

Ile Arg Thr Asn Ser Glu Gly Gly Phe Ile Trp Asp Glu Gly Ala Asn
50 55 60

Thr Met Thr Glu Ser Glu Leu Glu Ala Ser Arg Leu Ile Asp Asp Leu
65 70 75 80

Gly Leu Gln Gly Lys Gln Gln Tyr Pro Asn Ser Gln His Lys Arg Tyr
85 90 95

Ile Val Lys Asp Gly Ala Pro Thr Leu Ile Pro Ser Asp Pro Ile Ala
100 105 110

Leu Met Lys Ser Thr Val Leu Ser Thr Lys Ser Lys Leu Lys Leu Phe
115 120 125

Leu Glu Pro Phe Leu Tyr Glu Lys Ser Ser Arg Arg Thr Ser Gly Lys
130 135 140

Val Ser Asp Glu His Leu Ser Glu Ser Val Ile Phe Leu Cys Ile Cys
145 150 155 160

Arg Asp Asn Gln Val Val Asp Tyr Leu Ile Asp Pro Phe Val Ala Gly
165 170 175

Thr Ser Gly Gly Asp Pro Glu Ser Leu Ser Ile Arg His Ala Phe Pro
180 185 190

Ala Leu Trp Asn Leu Glu Asn Lys Tyr Gly Ser Val Ile Ala Gly Ala
195 200 205

Ile Leu Ser Lys Leu Ser Thr Lys Gly Asp Ser Val Lys Thr Gly Gly
210 215 220

Ala Ser Pro Gly Lys Gly Arg Asn Lys Arg Val Ser Phe Ser Phe His
225 230 235 240

Gly Gly Met Gln Ser Leu Ile Asp Ala Leu His Asn Glu Val Gly Asp
245 250 255

Gly Asn Val Lys Leu Gly Thr Glu Val Leu Ser Leu Ala Cys Cys Cys
260 265 270

Asp Gly Val Ser Ser Ser Gly Gly Trp Ser Ile Ser Val Asp Ser Lys
 275 280 285

Asp Ala Lys Gly Lys Asp Leu Arg Lys Asn Gl n Ser Phe Asp Ala Val
 290 295 300

Ile Met Thr Ala Pro Leu Ser Asn Val Gl n Arg Met Lys Phe Thr Lys
 305 310 315 320

Gly Gly Val Pro Phe Val Leu Asp Phe Leu Pro Lys Val Asp Tyr Leu
 325 330 335

Pro Leu Ser Leu Met Val Thr Ala Phe Lys Lys Gl u Asp Val Lys Lys
 340 345 350

Pro Leu Gl u Gly Phe Gly Ala Leu Ile Pro Tyr Lys Gl u Gl n Gl n Lys
 355 360 365

His Gly Leu Lys Thr Leu Gly Thr Leu Phe Ser Ser Met Met Phe Pro
 370 375 380

Asp Arg Ala Pro Asn Asp Gl n Tyr Leu Tyr Thr Ser Phe Ile Gly Gly
 385 390 395 400

Ser His Asn Arg Asp Leu Ala Gly Ala Pro Thr Ala Ile Leu Lys Gl n
 405 410 415

Leu Val Thr Ser Asp Leu Arg Lys Leu Leu Gly Val Gl u Gly Gl n Pro
 420 425 430

Thr Phe Val Lys His Val His Trp Arg Asn Ala Phe Pro Leu Tyr Gly
 435 440 445

Gl n Asn Tyr Asp Leu Val Leu Gl u Ala Ile Ala Lys Met Gl u Asn Asn
 450 455 460

Leu Pro Gly Phe Phe Tyr Ala Gly Asn Asn Lys Asp Gly Leu Ala Val
 465 470 475 480

Gly Asn Val Ile Ala Ser Gly Ser Lys Ala Ala Asp Leu Val Ile Ser
 485 490 495

Tyr Leu Gl u Ser Cys Thr Asp Gl n Asp Asn
 500 505

<210> 45
 <211> 1725
 <212> DNA
 <213> Oryza sativa

<400> 45
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eol f-seql

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ggcgccggcg	tcagtgggct	cgcggcggcg	tacaggctga	ggaagcgcg	cgtgcagggtg	240
acgggtgttcg	aggcggccga	cagggcggggt	gggaagatac	ggaccaactc	cgagggcgggg	300
ttcatctggg	acgaaggggc	caacaccatg	acagagagtg	aattggaggc	aagcaggctt	360
attgacgatc	ttggcctaca	aggcaaacag	cagtatccta	actcacaaca	caagcgttac	420
attgtcaaag	atggagcacc	aacactgatt	ccctcagatc	ccattgcgct	catgaaaagc	480
actgttcttt	ctacaaaatc	aaagctcaag	ctatttctgg	aaccatttct	ctatgagaaa	540
tctagcagaa	ggacctcggg	aaaagtgtct	gatgaacatt	taagtgagag	tgttgcaagt	600
ttctttgaac	gccactttgg	aaaagagggt	gttgattatc	ttattgatcc	atttgtggct	660
ggaacaagcg	gaggagatcc	tgagtcatta	tcaattcgtc	atgcatttcc	agcattatgg	720
aatttgagaa	ataagtatgg	ctctgtcatt	gctggtgcca	tcttgtccaa	actatccact	780
aagggtgatt	cagtgaagac	aggaggtgct	tcgccaggga	aaggaaggaa	taaacgtgtg	840
tcattttcat	ttcatggtgg	aatgcagtca	ctaatagatg	cacttcacaa	tgaagttgga	900
gatggtaacg	tgaagcttgg	tacagaagtg	ttgtcattgg	catgttgctg	tgatggagtc	960
tcttcttctg	gtggttggtc	aatttctgtt	gattcaaaag	atgctaaagg	gaaagatctc	1020
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ggatttggtg	ccttgatacc	ctataaggaa	cagcaaaagc	atggtctcaa	aacccttggt	1260
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ctatatacat	ctttcattgg	ggggagccat	aatagagacc	tcgctggggc	tccaacggct	1440
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<210> 46
 <211> 574
 <212> PRT
 <213> Oryza sativa

<400> 46

Met	Leu	Ser	Pro	Ala	Thr	Thr	Phe	Ser	Ser	Ser	Ser	Ser	Ser	Ser
1				5				10					15	

eol f-seql

Pro Ser Arg Ala His Ala Arg Ala Pro Thr Arg Phe Ala Val Ala Ala
20 25 30

Ser Ala Arg Ala Ala Arg Phe Arg Pro Ala Arg Ala Met Ala Ala Ser
35 40 45

Asp Asp Pro Arg Gly Gly Arg Ser Val Ala Val Val Gly Ala Gly Val
50 55 60

Ser Gly Leu Ala Ala Ala Tyr Arg Leu Arg Lys Arg Gly Val Gln Val
65 70 75 80

Thr Val Phe Glu Ala Ala Asp Arg Ala Gly Gly Lys Ile Arg Thr Asn
85 90 95

Ser Glu Gly Gly Phe Ile Trp Asp Glu Gly Ala Asn Thr Met Thr Glu
100 105 110

Ser Glu Leu Glu Ala Ser Arg Leu Ile Asp Asp Leu Gly Leu Gln Gly
115 120 125

Lys Gln Gln Tyr Pro Asn Ser Gln His Lys Arg Tyr Ile Val Lys Asp
130 135 140

Gly Ala Pro Thr Leu Ile Pro Ser Asp Pro Ile Ala Leu Met Lys Ser
145 150 155 160

Thr Val Leu Ser Thr Lys Ser Lys Leu Lys Leu Phe Leu Glu Pro Phe
165 170 175

Leu Tyr Glu Lys Ser Ser Arg Arg Thr Ser Gly Lys Val Ser Asp Glu
180 185 190

His Leu Ser Glu Ser Val Ala Ser Phe Phe Glu Arg His Phe Gly Lys
195 200 205

Glu Val Val Asp Tyr Leu Ile Asp Pro Phe Val Ala Gly Thr Ser Gly
210 215 220

Gly Asp Pro Glu Ser Leu Ser Ile Arg His Ala Phe Pro Ala Leu Trp
225 230 235 240

Asn Leu Glu Asn Lys Tyr Gly Ser Val Ile Ala Gly Ala Ile Leu Ser
245 250 255

Lys Leu Ser Thr Lys Gly Asp Ser Val Lys Thr Gly Gly Ala Ser Pro
260 265 270

Gly Lys Gly Arg Asn Lys Arg Val Ser Phe Ser Phe His Gly Gly Met
275 280 285

eol f-seql

Gln Ser Leu Ile Asp Ala Leu His Asn Glu Val Gly Asp Gly Asn Val
290 295 300

Lys Leu Gly Thr Glu Val Leu Ser Leu Ala Cys Cys Cys Asp Gly Val
305 310 315 320

Ser Ser Ser Gly Gly Trp Ser Ile Ser Val Asp Ser Lys Asp Ala Lys
325 330 335

Gly Lys Asp Leu Arg Lys Asn Gln Ser Phe Asp Ala Val Ile Met Thr
340 345 350

Ala Pro Leu Ser Asn Val Gln Arg Met Lys Phe Thr Lys Gly Gly Val
355 360 365

Pro Phe Val Leu Asp Phe Leu Pro Lys Val Asp Tyr Leu Pro Leu Ser
370 375 380

Leu Met Val Thr Ala Phe Lys Lys Glu Asp Val Lys Lys Pro Leu Glu
385 390 395 400

Gly Phe Gly Ala Leu Ile Pro Tyr Lys Glu Gln Gln Lys His Gly Leu
405 410 415

Lys Thr Leu Gly His Pro Ala Ser Cys Ile Glu Leu Asn Ile Gln Ile
420 425 430

Asn Leu Ala Thr Leu Leu Tyr Phe Phe Ser Gly Thr Leu Phe Ser Ser
435 440 445

Met Met Phe Pro Asp Arg Ala Pro Asn Asp Gln Tyr Leu Tyr Thr Ser
450 455 460

Phe Ile Gly Gly Ser His Asn Arg Asp Leu Ala Gly Ala Pro Thr Ala
465 470 475 480

Ile Leu Lys Gln Leu Val Thr Ser Asp Leu Arg Lys Leu Leu Gly Val
485 490 495

Glu Gly Gln Pro Thr Phe Val Lys His Val His Trp Arg Asn Ala Phe
500 505 510

Pro Leu Tyr Gly Gln Asn Tyr Asp Leu Val Leu Glu Ala Ile Ala Lys
515 520 525

Met Glu Asn Asn Leu Pro Gly Phe Phe Tyr Ala Gly Asn Asn Lys Asp
530 535 540

Gly Leu Ala Val Gly Asn Val Ile Ala Ser Gly Ser Lys Ala Ala Asp
545 550 555 560

Leu Val Ile Ser Tyr Leu Glu Ser Cys Thr Asp Glu Asp Asn
 565 570

eol f-seql

<210> 47
 <211> 1725
 <212> DNA
 <213> *Amaranthus tuberculatus*
 <400> 47
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 cccgcgcgcg ccatggccgc ctccgacgac cccgcggcg ggaggtccgt cgccgtcgtc 180
 ggcgcggcg tcagtgggt cgcggcgcg tacaggctga ggaagcgcg cgtgcagggtg 240
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 aatttgaga ataagtatgg ctctgtcatt gctggtgcc tcttgtccaa actatccact 780
 aagggtgatt cagtgaagac aggaggtgct tcgccaggga aaggaaggaa taaacgtgtg 840
 tcattttcat ttcatggtgg aatgcagtca ctaatagatg cacttcacaa tgaagttgga 900
 gatggtaacg tgaagcttgg tacagaagtg ttgtcattgg catgttgctg tgatggagtc 960
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 atgaagtta caaaagggtg agttcccttt gtgctagact ttcttcctaa ggctcgattat 1140
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 ttctcaggga ccctcttctc ctcatgatg tttccagatc gagctcctaa tgatcaatat 1380
 ctatatacat ctttcattgg ggggagccat aatagagacc tcgctggggc tccaacggct 1440
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 aataacaagg atgggttggc tgttggaat gttatagctt caggaagcaa ggctgctgac 1680
 cttgtgatct cttatcttga atcttgaca gatcaggaca attag 1725

eof-seql

<210> 48
 <211> 550
 <212> PRT
 <213> *Amaranthus tuberculatus*

<400> 48

Met Ser Ala Met Ala Leu Ser Ser Ser Ile Leu Gln Cys Pro Pro His
 1 5 10 15

Ser Asp Ile Ser Phe Arg Phe Phe Ala His Thr Arg Thr Gln Pro Pro
 20 25 30

Ile Phe Phe Gly Arg Pro Arg Lys Leu Ser Tyr Ile His Cys Ser Thr
 35 40 45

Ser Ser Ser Ser Thr Ala Asn Tyr Gln Asn Thr Ile Thr Ser Gln Gly
 50 55 60

Glu Gly Asp Lys Val Leu Asp Cys Val Ile Val Gly Ala Gly Ile Ser
 65 70 75 80

Gly Leu Cys Ile Ala Gln Ala Leu Ser Thr Lys His Ile Gln Ser Asn
 85 90 95

Leu Asn Phe Ile Val Thr Glu Ala Lys His Arg Val Gly Gly Asn Ile
 100 105 110

Thr Thr Met Glu Ser Asp Gly Tyr Ile Trp Glu Glu Gly Pro Asn Ser
 115 120 125

Phe Gln Pro Ser Asp Pro Val Leu Thr Met Ala Val Asp Ser Gly Leu
 130 135 140

Lys Asp Asp Leu Val Leu Gly Asp Pro Asn Ala Pro Arg Phe Val Leu
 145 150 155 160

Trp Asn Gly Lys Leu Arg Pro Val Pro Ser Lys Pro Thr Asp Leu Pro
 165 170 175

Phe Phe Asp Leu Met Ser Phe Pro Gly Lys Ile Arg Ala Gly Leu Gly
 180 185 190

Ala Leu Gly Leu Arg Pro Pro Pro Pro Ser Tyr Glu Glu Ser Val Glu
 195 200 205

Glu Phe Val Arg Arg Asn Leu Gly Asp Glu Val Phe Glu Arg Leu Ile
 210 215 220

Glu Pro Phe Cys Ser Gly Val Tyr Ala Gly Asp Pro Ala Lys Leu Ser
 225 230 235 240

Met Lys Ala Ala Phe Gly Lys Val Trp Thr Leu Glu Gln Lys Gly Gly
 245 250 255
 Ser Ile Ile Ala Gly Thr Leu Lys Thr Ile Gln Glu Arg Lys Asn Asn
 260 265 270
 Pro Pro Pro Pro Arg Asp Pro Arg Leu Pro Lys Pro Lys Gly Gln Thr
 275 280 285
 Val Gly Ser Phe Arg Lys Gly Leu Ile Met Leu Pro Thr Ala Ile Ala
 290 295 300
 Ala Arg Leu Gly Ser Lys Val Lys Leu Ser Trp Thr Leu Ser Asn Ile
 305 310 315
 Asp Lys Ser Leu Asn Gly Glu Tyr Asn Leu Thr Tyr Gln Thr Pro Asp
 325 330 335
 Gly Pro Val Ser Val Arg Thr Lys Ala Val Val Met Thr Val Pro Ser
 340 345 350
 Tyr Ile Ala Ser Ser Leu Leu Arg Pro Leu Ser Asp Val Ala Ala Asp
 355 360 365
 Ser Leu Ser Lys Phe Tyr Tyr Pro Pro Val Ala Ala Val Ser Leu Ser
 370 375 380
 Tyr Pro Lys Glu Ala Ile Arg Pro Glu Cys Leu Ile Asp Gly Glu Leu
 385 390 395 400
 Lys Gly Phe Gly Gln Leu His Pro Arg Ser Gln Gly Val Glu Thr Leu
 405 410 415
 Gly Thr Ile Tyr Ser Ser Ser Leu Phe Pro Gly Arg Ala Pro Pro Gly
 420 425 430
 Arg Thr Leu Ile Leu Ser Tyr Ile Gly Gly Ala Thr Asn Leu Gly Ile
 435 440 445
 Leu Gln Lys Ser Glu Asp Glu Leu Ala Glu Thr Val Asp Lys Asp Leu
 450 455 460
 Arg Lys Ile Leu Ile Asn Pro Asn Ala Lys Gly Ser Arg Val Leu Gly
 465 470 475 480
 Val Arg Val Trp Pro Lys Ala Ile Pro Gln Phe Leu Val Gly His Phe
 485 490 495
 Asp Val Leu Asp Ala Ala Lys Ala Gly Leu Ala Asn Ala Gly Gln Lys
 500 505 510

Gly Leu Phe Leu Gly Gly Asn Tyr Val Ser Gly Val Ala Leu Gly Arg
 515 520 525

Cys Ile Glu Gly Ala Tyr Asp Ser Ala Ser Glu Val Val Asp Phe Leu
 530 535 540

Ser Gln Tyr Lys Asp Lys
 545 550

<210> 49
 <211> 47
 <212> PRT
 <213> Zea mays

<400> 49

Met Leu Ala Leu Thr Ala Ser Ala Ser Ser Ala Ser Ser His Pro Tyr
 1 5 10 15

Arg His Ala Ser Ala His Thr Arg Arg Pro Arg Leu Arg Ala Val Leu
 20 25 30

Ala Met Ala Gly Ser Asp Asp Pro Arg Ala Ala Pro Ala Arg Ser
 35 40 45

<210> 50
 <211> 47
 <212> PRT
 <213> Sorghum

<400> 50

Met Leu Ala Arg Thr Ala Thr Val Ser Ser Thr Ser Ser His Ser His
 1 5 10 15

Pro Tyr Arg Pro Thr Ser Ala Arg Ser Leu Arg Leu Arg Pro Val Leu
 20 25 30

Ala Met Ala Gly Ser Asp Asp Ser Arg Ala Ala Pro Ala Arg Ser
 35 40 45

<210> 51
 <211> 57
 <212> PRT
 <213> Zea mays

<400> 51

Met Val Ala Ala Thr Ala Thr Ala Thr Ala Met Ala Thr Ala Ala Ser
 1 5 10 15

Pro Leu Leu Asn Gly Thr Arg Ile Pro Ala Arg Leu Arg His Arg Gly
 20 25 30

Leu Ser Val Arg Cys Ala Ala Val Ala Gly Gly Ala Ala Glu Ala Pro
 35 40 45

eol f-seql

Ala Ser Thr Gly Ala Arg Leu Ser Ala
50 55

<210> 52
<211> 56
<212> PRT
<213> Sorghum bicolor

<400> 52

Met Val Ala Ala Ala Met Ala Thr Ala Ala Ser Ala Ala Ala Pro
1 5 10 15

Leu Leu Asn Gly Thr Arg Arg Pro Ala Arg Leu Arg Arg Arg Gly Leu
20 25 30

Arg Val Arg Cys Ala Ala Val Ala Gly Gly Ala Ala Glu Ala Pro Ala
35 40 45

Ser Thr Gly Ala Arg Leu Ser Ala
50 55

<210> 53
<211> 51
<212> PRT
<213> Silene pratensis

<400> 53

Met Ala Ser Thr Leu Ser Thr Leu Ser Val Ser Ala Ser Leu Leu Pro
1 5 10 15

Lys Gln Gln Pro Met Val Ala Ser Ser Leu Pro Thr Asn Met Gly Gln
20 25 30

Ala Leu Phe Gly Leu Lys Ala Gly Ser Arg Gly Arg Val Thr Ala Met
35 40 45

Ala Thr Tyr
50

<210> 54
<211> 153
<212> DNA
<213> Silene pratensis

<400> 54

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atggtcgcct catcgctacc aactaatatg ggccaagcct tgtttggact gaaagccggt 120
tctcgtggca gactgactgc aatggccaca tac 153

<210> 55
<211> 153
<212> DNA
<213> Silene pratensis

eol f-seqI

<400> 55

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atggtaggcta gctcactccc tactaatatg ggtaggctc ttttcggact taaggctgga 120

tctaggggta gagttactgc tatggctacc tac 153