

Appendix to:

EFSA (European Food Safety Authority), 2020. Conclusion on the peer review of the pesticide risk assessment of the active substance prosulfuron. EFSA Journal 2020;18(7):6181, 35 pp. doi:10.2903/j.efsa.2020.6181

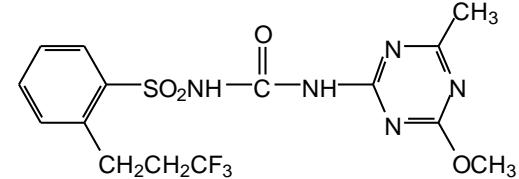
© European Food Safety Authority, 2020

## **Appendix A – List of end points for the active substance and the representative formulation**

### **Identity, Physical and Chemical Properties, Details of Uses, Further Information**

Active substance (ISO Common Name) ‡	Prosulfuron
Function ( <i>e.g.</i> fungicide)	Herbicide
Rapporteur Member State	France
Co-rapporteur Member State	Slovakia
<b>Identity (Annex II A, point 1)</b>	
Chemical name (IUPAC) ‡	<i>N</i> -(4-methoxy-6-methyl-1,3,5-triazin-2-yl)carbamoyl]-2-(3,3,3-trifluoropropyl)benzenesulfonamide or 1-(4-methoxy-6-methyl-1,3,5-triazin-2-yl)-3-[2-(3,3,3-trifluoropropyl)phenylsulfonyl]urea
Chemical name (CA) ‡	<i>N</i> -[(4-methoxy-6-methyl-1,3,5-triazin-2-yl)amino]carbonyl]-2-(3,3,3-trifluoropropyl)benzenesulfonamide
CIPAC No ‡	579
CAS No ‡	94125-34-5
EC No (EINECS or ELINCS) ‡	Not available
FAO Specification (including year of publication) ‡	/
Minimum purity of the active substance as manufactured ‡	950 g/kg
Identity of relevant impurities (of toxicological, ecotoxicological and/or environmental concern) in the active substance as manufactured	CGA 159902 2-(3,3,3-trifluoropropyl)benzenesulfonamide Maximum content 10 g/kg
Molecular formula ‡	<chem>C15H16F3N5O4S</chem>
Molar mass ‡	419.4 g/mol

Structural formula ‡



## Physical and chemical properties (Annex IIA, point 2)

Melting point (state purity) ‡	155°C with decomposition (99.5%)
Boiling point (state purity) ‡	Not measurable due to thermal decomposition
Temperature of decomposition (state purity)	Decomposition begins at about 150°C
Appearance (state purity) ‡	Light beige powder (technical a.i.: 95.1% purity) White powder (pure a.i. : 99.4% purity)
Vapour pressure (state temperature, state purity) ‡	< 3.5 × 10 <sup>-6</sup> Pa (25°C, 99.5 % purity)
Henry's law constant ‡	< 3 × 10 <sup>-4</sup> Pa.m <sup>3</sup> mol <sup>-1</sup>
Solubility in water (state temperature, state purity and pH) ‡	pH 5 : 87 mg/L (25°C, 99.5 % purity) pH 6.8 : 4 g/L (25°C) pH 7.7 : 43 g/L (25°C)
Solubility in organic solvents ‡ (state temperature, state purity)	At 20°C, 99.5% purity: hexane : 6.4 mg/L - toluene : 6.1 g/L octanol : 1.4 g/L - ethanol : 8.4 g/L ethyl acetate : 56 g/L - acetone : 160 g/L dichloromethane : 180 g/L
Surface tension ‡ (state concentration and temperature, state purity)	63 mN/m (20°C, 10.0 g/L suspension in water, 99.5 % purity)
Partition co-efficient ‡ (state temperature, pH and purity)	pH 5 : 1.5 (25°C, 99.5 % purity) pH 6 : - 0,21 (25°C, 99.5 % purity) pH 9 : - 0,76 (25°C, 99.5 % purity)
Dissociation constant (state purity) ‡	pKa = 3.76 (20°C, 99.5 % purity)
UV/VIS absorption (max.) incl. ε ‡ (state purity, pH)	λ max = 227.5 nm (98.4 % purity) ε : 21645 1.mol <sup>-1</sup> .cm <sup>-1</sup>
Flammability ‡ (state purity)	Not highly flammable 99.5 % purity
Explosive properties ‡ (state purity)	Not explosive 99.5 % purity
Oxidising properties ‡ (state purity)	No oxidizing properties 99.5 % purity





and LOQ)

very toxic

**Classification and proposed labelling with regard to physical and chemical data (Annex IIA, point 10)**

Active substance

RMS/peer review proposal

/

**Groundwater metabolites: Screening for biological activity (SANCO/221/2000-rev.10-final Step 3 a Stage 1)**

Activity against target organism

CGA1 50829	CGA1 59902	CGA3 00406	SYN5 42604	CGA3 49707	CGA3 25025	SYN5 47308
no						

## Impact on Human and Animal Health

### Other toxicological studies (Annex IIA, point 5.8)

Studies performed on metabolites or impurities

‡

- CGA159902 (CA1118A):

Acute oral LD<sub>50</sub> > 2000 mg/kg bw

Acute dermal LD<sub>50</sub> > 2000 mg/kg bw

Not a skin or eye irritant

Sensitiser (M&K test) – H317

*In vitro* genotoxicity tests - Ames test: negative; mouse lymphoma assay: positive (small colonies); cytogenetic assay in human lymphocytes: positive

*In vivo* genotoxicity tests – UDS assay: negative; mouse bone marrow micronucleus test: negative

Unlikely to be genotoxic *in vivo*.

- CGA150829 (triazine amine):

Acute oral LD<sub>50</sub> > 2000 mg/kg bw (M); =1000 mg/kg bw (F) – R22

Acute dermal LD<sub>50</sub> > 2000 mg/kg bw

Acute inhalation LC<sub>50</sub> > 5.2 mg/L

Not a skin or eye irritant

Not a sensitiser (M&K test)

Genotoxic potential cannot be concluded.

- CGA325025:

*In vitro* genotoxicity tests - Ames test, mouse lymphoma assay: negative; cytogenetic assay in human lymphocytes: equivocal.

Genotoxic potential cannot be excluded.

- CGA349707:

*In vitro* genotoxicity tests - Ames test, mouse lymphoma assay and cytogenetic assay in human lymphocytes: negative.

ADI of 0.001 mg/kg per day.

- SYN547308:

*In vitro* genotoxicity tests - Ames test, mouse lymphoma assay: negative; cytogenetic assay in human lymphocytes: positive.

*In vivo* genotoxicity tests – mouse bone marrow micronucleus test: negative

Genotoxic *in vitro*, not *in vivo*.

- SYN542604:

*In vitro* genotoxicity tests - Ames test, mouse lymphoma assay and cytogenetic assay in human lymphocytes: negative.

Unlikely to be genotoxic.

- CGA300406:

*In vitro* genotoxicity tests - Ames test, mouse lymphoma assay: negative; cytogenetic assay in human lymphocytes: positive.

*In vivo* genotoxicity tests – mouse bone marrow micronucleus test: negative

Genotoxic *in vitro*, not *in vivo*.

**Metabolism in plants (Annex IIA, point 6.1 and 6.7, Annex IIIA, point 8.1 and 8.6)**

Plant groups covered	Cereals (maize)
Rotational crops	Radish, spinach, wheat, lettuce
Metabolism in rotational crops similar to metabolism in primary crops?	Yes
Processed commodities	Not necessary
Residue pattern in processed commodities similar to residue pattern in raw commodities?	-
Plant residue definition for monitoring	Prosulfuron (Draft subject to the data gap on the genotoxicity of CGA150829)
Plant residue definition for risk assessment	Prosulfuron (Draft subject to the data gap on the genotoxicity of CGA150829)
Conversion factor (monitoring to risk assessment)	-

**Metabolism in livestock (Annex IIA, point 6.2 and 6.7, Annex IIIA, point 8.1 and 8.6)**

Animals covered	Goat, hen
Time needed to reach a plateau concentration in milk and eggs	Milk: n.a. Eggs : 2 days (egg white), 6 days (egg yolk)
Animal residue definition for monitoring	Prosulfuron
Animal residue definition for risk assessment	Prosulfuron
Conversion factor (monitoring to risk assessment)	-
Metabolism in rat and ruminant similar (yes/no)	Yes
Fat soluble residue: (yes/no)	No ( $\log P_{ow} = -0.21$ at pH 6)

**Residues in succeeding crops (Annex IIA, point 6.6, Annex IIIA, point 8.5)**

No accumulation in soil.  
No uptake of soil specific metabolites.

**Stability of residues (Annex IIA, point 6 introduction, Annex IIIA, point 8 Introduction)**

Residues of prosulfuron are stable under freezer storage for at least 25 months (maize grain and forage; beef muscle, beef liver, and milk), 12 months in maize oil and 16 months in eggs

**Residues from livestock feeding studies (Annex II A, point 6.4, Annex III A, point 8.3)**

	Ruminant	Poultry	Pig
<b>Conditions of requirement of feeding studies</b>			
Expected intakes by livestock $\geq 0.1$ mg/kg diet (dry weight basis) (yes/no - If yes, specify the level)	No (but study available)	No (but study available)	No
Potential for accumulation (yes/no)	No	No	No
Metabolism studies indicate potential level of residues $\geq 0.01$ mg/kg in edible tissues (yes/no)	No	No	No
Feeding studies (feeding study on lactating cows 5, 15, 50 ppm in the feed; feeding study on laying hens 0.1, 0.3, 1 ppm in the feed). Residue levels in matrices : Mean (max) mg/kg			
Muscle	<0.05 mg/kg whatever the feeding dose	<0.05 mg/kg whatever the feeding dose	-
Liver	<0.05 mg/kg whatever the feeding dose	<0.05 mg/kg whatever the feeding dose	-
Kidney	<0.05 mg/kg whatever the feeding dose	<0.05 mg/kg whatever the feeding dose	-
Fat	<0.05 mg/kg whatever the feeding dose	<0.05 mg/kg whatever the feeding dose	-
Milk	<0.01 mg/kg whatever the feeding dose		
Eggs		<0.05 mg/kg whatever the feeding dose	

**Summary of residues data according to the representative uses on raw agricultural commodities and feedingstuffs (Annex II A, point 6.3, Annex III A, point 8.2)**

Crop	Northern or Mediterranean Region, field or glasshouse, and any other useful information	Trials results relevant to the representative uses (a)	Recommendation/comments	MRL estimated from trials according to the representative use	HR (c)	STMR (b)
Maize	Northern EU	8×<0.01; 4×<0.02	EU intended GAP : foliar treatment, 20 g as/ha, PHI 90 days (grain)	0.02*	<0.02	<0.01
	Southern EU	4×<0.01; 3×<0.02			<0.02	<0.01
Sweet corn	Northern EU	11×<0.01	EU intended GAP : foliar treatment, 20 g as/ha, PHI 90 days (grain)	0.02*	<0.01	<0.01
	Southern EU	4×<0.01			<0.01	<0.01

(a) Numbers of trials in which particular residue levels were reported *e.g.* 3 × <0.01, 1 × 0.01, 6 × 0.02, 1 × 0.04, 1 × 0.08, 2 × 0.1, 2 × 0.15, 1 × 0.17

(b) Supervised Trials Median Residue *i.e.* the median residue level estimated on the basis of supervised trials relating to the representative use

(c) Highest residue

\* The MRL is proposed at the LOQ.

**Consumer risk assessment (Annex II A, point 6.9, Annex III A, point 8.8)**

ADI	0.02 mg/kg bw per day
TMDI (% ADI) according to WHO European diet	0.3 % (WHO cluster diet B)
TMDI (% ADI) according to French diets	0.01 % (FR toddler)
IEDI (WHO European Diet) % ADI)	Not performed as TMDI is below ADI
NEDI (specify diet) (% ADI)	N/A
Factors included in IEDI and NEDI	N/A
ARfD	0.1 mg/kg bw
IESTI (% ARfD)	0.7% (sweet corn) 0.1% (maize)
NESTI (% ARfD) according to national (to be specified) large portion consumption data	N/A
Factors included in IESTI and NESTI	N/A

**Processing factors (Annex II A, point 6.5, Annex III A, point 8.4)**

Not necessary as residue level in RAC<0.1 mg/kg

**Proposed MRLs (Annex II A, point 6.7, Annex III A, point 8.6)**

Maize grain	0.02* mg/kg
Sweet corn	0.02* mg/kg

When the MRL is proposed at the LOQ, this should be annotated by an asterisk after the figure.



























incubation at 25°C, 508 mm  
 Residue prosulfuron 70 - 84 %, metabolites < 12 %  
 (CGA 300406)  
 RA in leachates : 33 - 59 %, mainly prosulfuron,  
 metabolites < 4 % each

2 soils (OC 1.7 - 2.1 %, pH 7.2 - 7.3), 2 labels, 180 d  
 incubation at 20°C, 200 mm  
 Residue : prosulfuron 18 - 31 %, metabolites <10 %  
 each (CGA 159902, 150829, 300406, 349707, M5),  
 CO<sub>2</sub> 10 - 45 %  
 RA in leachates : 0.8 - 12 % of applied to columns  
 (prosulfuron)

#### Lysimeter/ field leaching studies ‡

USA, undisturbed 20 cm diameter soil columns, silt loam (1.94 % OC, pH 5.6) in Kentucky and sand soil (0.3 % OC, pH 4.9) in North Carolina, 2 labels, 44 g as/ha. Overflow occurred in Kentucky.  
 Total residues in drainage water (LOD 0.4 µg/l) soil depth 0.90 m  
 Silt loam:  
 < 0.4 %, mean conc. 0.13 µg/l (phenyl); < 0.1 % (triazine)  
 Sand:  
 mean 0.98 µg/l max. 3 µg/l (phenyl); mean 0.08 µg/l max. 1 µg/l (triazine)  
 Compounds in drainage water from sand soil, NC prosulfuron traces in initial preferential flow soil depth 0.90 m  
 prosulfuron phenyl sulfonamide (CGA159902) max. 2.4 µg/l  
 M5 (derivative of prosulfuron phenyl sulfonamide (CGA159902)) max. 1 µg/l  
 CGA325028 max. 0.74 µg/l  
 O-desmethyl-prosulfuron (CGA300406) max. 0.08 µg/l  
 No realistic mean concentrations can be calculated for individual compounds. Triazine moiety less mobile than phenyl moiety. No prosulfuron triazine amine (CGA150829) in soil after 1 year.

Swiss lysimeter, sandy soil (1.05 % OC, pH 6.1), phenyl label, 28 or 2 x 28 g as/ha (Spring)  
 Total residues (LOD 0.05 µg/l)  
 mean concentrations, soil depth 1.2 m  
 1 appl. 0.23 / 0.12 / 0.07 µg/l (year 1 / 2 / 3)  
 2 appl. 0.24 / 0.31 / 0.22 µg/l (year 1 / 2 / 3)  
 max. concentrations 0.46 / 0.42 µg/l (1 / 2 appl.)

Prosulfuron, prosulfuron phenyl sulfonamide (CGA159902), O-desmethyl-prosulfuron (CGA300406), CGA349707, SYN542604 (M5), demethoxy amino-prosulfosulfuron (CGA 325025) and unknowns < 0.1 µg/l each. Total extractable RA in soil < 2.5 µg/kg after 3 years.

### Route and rate of degradation in water (Annex II A, point 7.2.1)

Hydrolytic degradation of the active substance and metabolites > 10 % ‡

pH 5 (25°C)
DT <sub>50</sub> 5 - 12 d
prosulfuron phenyl sulfonamide (CGA159902) 58 %
prosulfuron triazine amine (CGA150829) 43 %
prosulfuron polyimide (CGA325030) 22-31 %
G28533 16 %
pH 7 (25°C)
DT <sub>50</sub> 424 - 651 d
pH 9 (25°C)
DT <sub>50</sub> 682 - 1690 d
(no major metabolite at pH 7 and 9)

### PEC (ground water) (Annex III A, point 9.2.1)

Method of calculation and type of study (e.g. modelling, field leaching, lysimeter )

Models used: FOCUS-PELMO 5.5.3 and FOCUS-PEARL 4.4.4, FOCUS-MACRO 5.5.4

The following PECgw were calculated in a context of amendment of the approval conditions (to remove the restriction to application once every 3 years) and to respond to Data requirement by EFSA.

Input parameters for prosulfuron:

Molar mass = 419.4 g/mol

Water solubility = 43000 mg/L (20°C), 86000 mg/L (30°C)

Vapour pressure = 0 Pa (20°C), 0 Pa (30°C)

Kfoc = 11.7 mL/g (geometric mean, n=8)

1/n = 0.869

DT<sub>50</sub> = 19.6 days (normalised field geometric mean, pseudo-SFO, n=6). The correct value to be used for further calculations is 18.7 days.

Plant uptake factor = 0.15

Input parameters for prosulfuron triazine amine (CGA150829):

Molar mass = 140.1 g/mol  
 Water solubility = 1000 mg/L (considered conservative)  
 Vapour pressure = 0 Pa  
 $K_{foc} = 45.6 \text{ mL/g (n=27)}$   
 $1/n = 0.9 \text{ (n=27)}$   
 $DT_{50} = 196 \text{ days (normalised laboratory, median, SFO, n= 18). The correct value to be used for further calculations is 216 days}$   
 $ffM = 0.28 \text{ (from prosulfuron).}$   
 Plant uptake factor = 0

Input parameters for prosulfuron phenyl sulfonamide (CGA159902):

Molar mass = 253.2 g/mol  
 Water solubility = 1000 mg/L (considered conservative)  
 Vapour pressure = 0 Pa  
 $K_{foc} = 68.0 \text{ mL/g (geometric mean, n=4); K}_{fom} = 39.4 \text{ mL/g}$   
 $1/n = 0.88$   
 $DT_{50} = 188 \text{ days (normalised laboratory geometric mean, SFO, n=5)}$   
 $ffM = 0.43 \text{ (from prosulfuron)}$   
 Plant uptake factor = 0

Input parameters for O-desmethyl-prosulfuron (CGA300406):

Molar mass = 405.4 g/mol  
 Water solubility = 1000 mg/L (considered conservative)  
 Vapour pressure = 0 Pa  
 $K_{foc} = 46.8 \text{ mL/g (geometric mean, n=3); K}_{fom} = 27.1 \text{ mL/g}$   
 $1/n = 0.90$   
 $DT_{50} = 30.2 \text{ days (maximum laboratory } DT_{50} \text{ representing alkaline conditions, SFO); 2.6 days (minimum laboratory } DT_{50} \text{ representing acidic conditions, SFO)}$   
 $ffM = 0.47 \text{ (from prosulfuron).}$   
 Plant uptake factor = 0

Input parameters for demethoxy amino-prosulfosulfuron (CGA 325025):

Molar mass = 404.4 g/mol  
 Water solubility = 1000 mg/L (considered conservative)

Vapour pressure = 0 Pa Kfoc = 26.2 mL/g (geometric mean, n=4); Kfom = 15.2 mL/g 1/n = 0.973 DT <sub>50</sub> = 62.4 days (normalised laboratory geometric mean, SFO, n=3) ffM = 0.12 (from O-desmethyl-prosulfuron (CGA300406)) Plant uptake factor = 0
<b><u>Input parameters for SYN542604:</u></b>
Molar mass = 381.3 g/mol Water solubility = 1000 mg/L (considered conservative) Vapour pressure = 0 Pa Kfoc = 111 mL/g (geometric mean, n=5); Kfom = 64.4 mL/g 1/n = 0.85 DT <sub>50</sub> = 84.6 days (normalised laboratory geometric mean, SFO, n=8). ffM = 0.88 (from O-desmethyl-prosulfuron (CGA300406)) Plant uptake factor = 0
<b><u>Input parameters for CGA349707:</u></b>
Molar mass = 338.3 g/mol Water solubility = 1000 mg/L (considered conservative) Vapour pressure = 0 Pa Kfoc = 44.0 mL/g (geometric mean, n=3); Kfom = 25.5 mL/g 1/n = 0.96 DT <sub>50</sub> = 153 days (normalised laboratory geometric mean, SFO, n=4) ffM = 0.86 (from SYN542604) Plant uptake factor = 0
<b><u>Input parameters for SYN547308:</u></b>
Molar mass = 449.4 g/mol Water solubility = 1000 mg/L (considered conservative) Vapour pressure = 0 Pa Kfoc = 89.5 mL/g; Kfom = 51.9 mL/g 1/n = 0.929 DT <sub>50</sub> = 67.1 days (normalised laboratory geometric mean, SFO, n=4)



MACRO 5.5.4/maize 20 g/ha	Sev	0.001	0.060	0.042	< 0.001	< 0.001	0.326	0.016	0.004
	Thi	0.008	0.190	0.256	< 0.001	0.004	0.741	0.050	0.050
PEARL 4.4.4/maize 15 g/ha	Scenario	Parent (µg/L)	Metabolite (µg/L)						
			prosulfuron triazine amine CGA15082 9	prosulfuron phenyl sulfonamide CGA15990 2	O-desmethyl-prosulfuron CGA30040 6	SYN54260 4	CGA34970 7	demethoxy amino-prosulfuron CGA325025	SYN54730 8
PEALM 5.5.3/maize 15 g/ha	Cha	0.022	0.132	0.202	<0.001	0.008	0.503	0.044	0.071
	Scenario	Parent (µg/L)	Metabolite (µg/L)						
			prosulfuron triazine amine CGA15082 9	prosulfuron phenyl sulfonamide CGA15990 2	O-desmethyl-prosulfuron CGA30040 6	SYN54260 4	CGA34970 7	demethoxy amino-prosulfuron CGA325025	SYN54730 8
	Cha	0.036	0.129	0.211	0.001	0.016	0.531	0.054	0.100
	Ham	0.086	0.158	0.268	0.004	0.031	0.687	0.080	0.148
	Jok	-	-	-	-	-	-	-	-
	Kre	0.057	0.117	0.200	0.002	0.032	0.473	0.055	0.131
	Oke	0.089	0.112	0.217	0.003	0.039	0.410	0.051	0.162
	Pia	0.018	0.131	0.198	<0.001	0.019	0.549	0.042	0.100
	Por	0.007	0.062	0.100	<0.001	0.004	0.278	0.025	0.043
PELMO 5.5.3/maize 15 g/ha	Sev	<0.00 1	0.051	0.039	<0.001	<0.001	0.261	0.012	0.006
	Thi	0.010	0.166	0.237	<0.001	0.006	0.702	0.048	0.056

M	Scenari	Paren	Metabolite (µg/L)
---	---------	-------	-------------------





	Cha	0.015	0.096	0.144	0.012	0.006	0.374	0.033	0.051
--	-----	-------	-------	-------	-------	-------	-------	-------	-------

### Data gap identified to address potential ground water contamination by soil metabolite M17

#### PEC (gw) from lysimeter / field studies

Parent	1 <sup>st</sup> year	2 <sup>nd</sup> year	3 <sup>rd</sup> year
Annual average ( $\mu\text{g/L}$ )	Not available for separate compounds		

Metabolite X	1 <sup>st</sup> year	2 <sup>nd</sup> year	3 <sup>rd</sup> year
Annual average ( $\mu\text{g/L}$ )	Not available for separate compounds		

#### Residues requiring further assessment

Environmental occurring residues requiring further assessment by other disciplines (toxicology and ecotoxicology) and / or requiring consideration for groundwater exposure.

Soil: prosulfuron, triazine amine (CGA150829), prosulfuron phenyl sulfonamide (CGA159902), O-desmethyl-prosulfuron (CGA300406), demethoxy amino-prosulfuron (CGA325025), SYN542604, CGA349707,

Ground water: prosulfuron, triazine amine (CGA150829), prosulfuron phenyl sulfonamide (CGA159902), O-desmethyl-prosulfuron (CGA300406), demethoxy amino-prosulfuron (CGA325025), SYN542604, CGA349707, M17 (pending identification), M18 ((SYN547308).

Surface water/  
prosulfuron, triazine amine (CGA150829), prosulfuron phenyl sulfonamide (CGA159902), O-desmethyl-prosulfuron (CGA300406), demethoxy amino-prosulfuron (CGA325025), SYN542604, CGA349707

Sediment: Prosulfuron, prosulfuron phenyl sulfonamide (CGA159902), O-desmethyl-prosulfuron (CGA300406)

Air: Prosulfuron

#### Monitoring data, if available (Annex II A, point 7.4)

Soil (indicate location and type of study)

Not available

Surface water (indicate location and type of study)

Not available

Ground water (indicate location and type of study)

Not available

Air (indicate location and type of study)

Not available

**Ecotoxicologically relevant compounds** (consider parent and all relevant metabolites requiring further assessment from the fate section)

Compartment	
soil	prosulfuron
water	prosulfuron
sediment	prosulfuron