Section A6.1.4s Acute Dermal Irritation - Rabbit

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			Official use only
		1 REFERENCE	
1.1	**Reference**	[redacted]; Acute Dermal Irritation / Corrosion Study in the Rabbit of CGA 55186 Tech. (cypermethrin); [redacted] Dates of experimental work: 16 February 1984 – 27 February 1984	
1.2	**Data protection**	Yes	
1.2.1	Data owner	Chimac-Agriphar s.a.	
1.2.2			
1.2.3	Criteria for data protection	Data submitted to the MS after 13 May 2000 on existing a.s. for the purpose of its entry into Annex I authorisation.	
		2 GUIDELINES AND QUALITY ASSURANCE	
2.1	**Guideline study**	Yes, protocol based on OECD guideline no. 404 (adopted 12 May 1981), corresponding method B.4. of Directive 92/69/EEC.	
2.2	**GLP**	No GLP was not compulsory at the time the study was performed.	
2.3	**Deviations**	No	
		3 MATERIALS AND METHODS	
3.1	**Test material**	CGA 55186 tech (cypermethrin cis:trans/40:60).	
3.1.1	Lot/Batch number	307046	
3.1.2	Specification	Deviating from specification given in section 2 as follows	
3.1.2.1	Description	Liquid, viscous	
3.1.2.2	Purity	92.6%	
3.1.2.3	Stability	Stable	X
3.2	**Test Animals**		
3.2.1	Species	Rabbit	
3.2.2	Strain	White New Zealand	
3.2.3	Source	[redacted]	
3.2.4	Sex	Male	
3.2.5	Age/weight at study initiation	12-14 weeks, 1980-2150 g	
3.2.6	Number of animals per group	3	
3.2.7	Control animals	No	
3.3	**Administration/ Exposure**	Dermal	

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3.3.1	Application		
3.3.1.1	Preparation of test substance	Test substance was used as delivered.	
3.3.1.2	Test site and Preparation of Test Site	Not less than 24 hours before treatment, an area of 6 cm² was shaved on the back of each animal.	
3.3.2	Occlusion	Occlusive	
3.3.3	Vehicle	None	
3.3.4	Concentration in vehicle	Not applicable, test item was administered as delivered	
3.3.5	Total volume applied	0.5 ml	
3.3.6	Removal of test substance	Dressing removed and skin observed. Removal of test substance not mentioned in report.	
3.3.7	Duration of exposure	4 hours	
3.3.8	Postexposure period	7 days	
3.3.9	Controls	None	
3.4	**Examinations**		
3.4.1	Clinical signs	No	
3.4.2	Dermal examination	Yes	
3.4.2.1	scoring system	Scoring system for erythema and eschar formation: No erythema = 0 Very slight erythema = 1 Well defined erythema = 2 Moderate or severe erythema = 3 Severe erythema to slight eschar formation = 4 Corrosion = + Scoring system for oedema formation: No oedema = 0 Very slight oedema = 1 Slight oedema = 2 Moderate oedema (approx. 1 mm) = 3 Severe oedema (> 1 mm, extended beyond applic. site)) = 4	X
3.4.2.2	Examination time points	1h, 24h, 48h, 72h, 7 days	
3.4.3	Other examinations	Bodyweight	
3.5	**Further remarks**	-	

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		4 RESULTS AND DISCUSSION	
4.1	**Average score**		
4.1.1	Erythema	Average score (EU methodology) for all animals at 24, 48, 72 h = 1.3, 1.3, 2	X
4.1.2	Edema	Average score (EU methodology) for all animals at 24, 48, 72 h = 1, 1, 2	X
4.2	**Reversibility**	Yes All animals recovered after the 7 day observation period	
4.3	**Other examinations**	All animals showed normal body weight development. No other reactions observed.	
4.4	**Overall result**	Cypermethrin is slightly irritant and not corrosive	
		5 APPLICANT'S SUMMARY AND CONCLUSION	
5.1	**Materials and methods**	Acute (4h) irritation/corrosion to rabbit skin using 0.5 ml cypermethrin. Study in compliance with EC method B.4 of Directive 92/69/EEC (corresponding OECD guideline 404)	
5.2	**Results and discussion**	Animals showed very slight to well defined erythema and very slight oedema. All animals recovered within 7 days.	
5.3	**Conclusion**		
5.3.1	Reliability	1	
5.3.2	Deficiencies	No. Study evaluated and accepted under Directive 91/414/EC.	

	Evaluation by Competent Authorities
	Use separate "evaluation boxes" to provide transparency as to the comments and views submitted
	EVALUATION BY RAPPORTEUR MEMBER STATE
Date	March, 2007.

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Materials and Methods	The applicant's version is acceptable with the following amendments: *3.1.2.3 Stability*: data no shown *3.4.2.1 Scoring system, mode of calculation and interpretation*: **The index of primary cutaneous irritation** was calculated as follows: The scores obtained for erythema and oedema at 1, 24, 48, and 72 hours after removal of the patch on the 3 rabbits examined are summed up. The sums of total oedema and erythema were divided by 12 (when all the animals survived). The score obtained is defined as the index of primary cutaneous irritation. Interpretation: Index < 0.5 : not irritant 0.6 – 3.0 : slightly irritant 3.1 – 5.0 : irritant 5.1 – 8.0 : severely irritant
Results and discussion	Revision of results: See table A6_1_4S-1. The calculated dermal irritation index was 2.0.
Conclusion	Cypermethrin is slightly irritant and not corrosive when applied to the rabbit skin. Cypermethrin does not require classification.
Reliability	1
Acceptability	acceptable
Remarks	-
	COMMENTS FROM ...
Date	*Give date of comments submitted*
Materials and Methods	*Discuss additional relevant discrepancies referring to the (sub)heading numbers and to applicant's summary and conclusion.* *Discuss if deviating from view of rapporteur member state*
Results and discussion	*Discuss if deviating from view of rapporteur member state*
Conclusion	*Discuss if deviating from view of rapporteur member state*
Reliability	*Discuss if deviating from view of rapporteur member state*
Acceptability	*Discuss if deviating from view of rapporteur member state*
Remarks	

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Table A6_1_4S-1. Table for skin irritation study

score	time	Erythema	Edema
average score (EU methology)/ animal	***24h, 48h, 72h***	***1.33/1.33/2.0***	***0.33/0.33/0.67***
Individual score (Draize scores)	60 min	0/2/2	0/1/1
	24 h	2/2/2	1/1/1
	48 h	1/1/2	0/0/1
	72 h	1/1/2	0/0/0
other times	7 days	0/0/0	0/0/0
reversibility: *		c	c
average time for reversibility		7 days	72 h
Index of primary cutaneous irritation		2.0	

* c : completely reversible
n c : not completely reversible
n : not reversible