



## Stofdocument deel A

CAS-nr: 127-18-4

**Perchloorethyleen**  $\text{CCl}_2=\text{CCl}_2$

VN-nr: 1897

GEVI: 60

**Synoniemen:** tetrachloorethyleen, per, ethyleentetrachloride (Engels: tetrachloroethylene)

Interventiewaarden		10 min.	30 min.	1 uur	2 uur	4 uur	8 uur
Voorlichtingsrichtwaarde	<b>VRW (mg/m<sup>3</sup>)</b>	240	240	240	240	240	240
Alarmeringsgrenswaarde	<b>AGW (mg/m<sup>3</sup>)</b>	1.700	1.700	1.700	1.200	790	540
Levensbedreigende	<b>LBW (mg/m<sup>3</sup>)</b>	35.000	19.000	13.000	8.600	5.800	3.900
Datum vaststelling: 24-09-2009		1 mg/m <sup>3</sup> = 0,145 ppm; 1 ppm = 6,90 mg/m <sup>3</sup>					
<b>Explosiegrens:</b> LEL = n.v.t.			<b>Geur:</b> oplosmiddelachtige / etherische geur				
			<b>LOA:</b> niet afgeleid				

Fysisch-chemische eigenschappen		Overige informatie
<b>Uiterlijk:</b> kleurloze vloeistof	Molecuulmassa: 165,8 g/mol  Zuurgraad: geen data LogKow: 3,4  Wateroplosbaarheid: Niet Verzadigde dampdruk: 18,9 mbar	Publieke grenswaarde: niet afgeleid  MAK: niet afgeleid TLV-TWA: 172 mg/m <sup>3</sup>
<b>Brand:</b> niet brandbaar		
<b>Relatieve dichtheid van verzadigd damp-lucht mengsel:</b> 1,09		Vluchtig Vloeistof wordt makkelijk door intacte huid opgenomen

Toxicologische eigenschappen	
<b>Effecten bij inhalatoire blootstelling</b>  <u>Onder VRW:</u> geen effecten  <u>VRW → AGW:</u> irritatie ogen, huid en luchtwegen, hoesten, misselijkheid, hoofdpijn, duizeligheid, verminderde coördinatie  <u>AGW → LBW:</u> benauwdheid, misselijkheid, bewustzijnsdaling  <u>Boven LBW:</u> ademnood, verstoring van het hartritme, longoedeem, ademstilstand, coma, sterfte	<b>Toxiciteit bij eenmalige, inhalatoire blootstelling</b> <ul style="list-style-type: none"> <li>Perchloorethyleen werkt irriterend op de ogen, huid en luchtwegen.</li> <li>Perchloorethyleen heeft een depressieve werking op het CZS en de ademhaling.</li> <li>Effecten op de lever en nier na blootstelling aan perchloorethyleen zijn beschreven.</li> <li>Blootstelling aan perchloorethyleen kan longoedeem veroorzaken. De verschijnselen hiervan kunnen vertraagd optreden en versterkt worden door lichamelijke inspanning.</li> </ul>
<b>Effecten bij blootstelling aan vloeistof</b>  <u>Huidcontact:</u> irritatie, roodheid, pijn  <u>Oogcontact:</u> irritatie, roodheid, pijn, tranen	<b>Carcinogeniteit</b>  IARC classificatie: 2A CRP: niet afgeleid

Beknopte medische informatie
<b>Ontsmetting damp</b> <i>algemeen:</i> frisse lucht, rust en direct spoedeisende medische hulp inzetten.
<b>Ontsmetting vloeistof</b> <i>huid:</i> verontreinigde kleding uittrekken, minimaal 20 min. spoelen met veel water of douchen, arts raadplegen.  <i>ogen:</i> minimaal 15 min. spoelen met water (evt. contactlenzen verwijderen), dan naar oogarts brengen.  <i>inslikken:</i> mond laten spoelen (uitspugen!), GEEN braken opwekken en direct spoedeisende medische hulp inzetten.
<b>Specifieke behandeling en materialen:</b> geen.
Neem contact op met het NVIC (tel: +31 (0)30 –274 8888) voor informatie met betrekking tot medisch handelen.



## Stofdocument deel B

CAS-nr: 127-18-4

**Tetrachloroethylene**

$\text{CCl}_2=\text{CCl}_2$

UN-nr: 1897

### Basis for the Dutch Intervention Values

**VRW:** AEGL value is adopted, 2h value added

**AGW:** Same point of departure as AEGL values, but different value for n

**LBW:** Different point of departure and different value for n than AEGL values

Date: 24-09-2009

AEGL document: Interim, 2009

### Dutch Intervention Values ( $\text{mg}/\text{m}^3$ )

	10 min	30 min	1 h	2 h	4 h	8 h	End point
<b>VRW</b>	240	240	240	240	240	240	Mild eye irritation in humans
<b>AGW</b>	1,700	1,700	1,700	1,200	790	540	No-effect level for ataxia in rats
<b>LBW</b>	35,000	19,000	13,000	8,600	5,800	3,900	Calculated treshold ( $\text{LC}_{01}$ ) for lethality in animals (mice).

### Derivation of the Dutch Intervention Values

**VRW:** The VRW derivation is based on the exposure of 6 volunteers to 106 ppm ( $730 \text{ mg}/\text{m}^3$ ) for 1 hour. At this level, an apparent non-objectionable odour and eye irritation were noted, and one subject experienced a slight fullness in the head. An interspecies uncertainty factor was not applicable. An intraspecies uncertainty factor of 3 was applied because mucous membrane irritation is caused by a direct effect of the chemical and the response is not expected to vary greatly among individuals. Because irritation is considered a threshold effect which should not vary over time, the VRW value was not scaled across time, but the same value was applied to all times.

**AGW:** The AGW values are based upon the no-effect level for ataxia after first exposure in rats following exposure to 1,150 ppm ( $7,900 \text{ mg}/\text{m}^3$ ) tetrachloroethylene for 4 hours/day, 5 days/week for 2 weeks (the time period of 4 hours) was used for the derivation. Exposure to the next higher concentration of 2,300 ppm ( $16,000 \text{ mg}/\text{m}^3$ ) resulted in reversible ataxia. A total uncertainty factor of 10 is applied. An interspecies uncertainty factor of 3 is applied based on the similarity of effects manifested in rodents compared to humans produced by agents that are CNS depressants. An intraspecies uncertainty factor of 3 is applied based on total range of sensitivity (2-3 fold) between several different groups (newborns, pregnant women and the elderly) for anaesthetic gases. Many organic vapours, particularly those which are strongly lipophilic, produce an anaesthetic effect in exposed humans. On the basis of this knowledge, it is reasonable to assume that the same 2-3 fold difference in sensitivity among individuals would apply for tetrachloroethylene.

Time scaling was performed using the equation  $C^n \times t = k$  and an n-value of 1.8 (based on probit analysis of rat mortality data). The 10- and 30-minute AGW values were set equal to the 1-hour value of 250 ppm ( $1,700 \text{ mg}/\text{m}^3$ ) because a human study demonstrated an exposure to 600 ppm ( $4,100 \text{ mg}/\text{m}^3$ ) for 10 minutes caused significant effects (eye and nose irritation, dizziness, tightness and numbing about the mouth, some loss of inhibitions, and motor coordination required great effort). After applying an uncertainty factor of 3 (for intraspecies variation), the AGW values based upon this study are consistent with the 1- hour AGW value of 250 ppm ( $1,700 \text{ mg}/\text{m}^3$ ).

**LBW:** In contrast to the AEGL, the LBW derivation is not based on one-third of the 4-hour mouse  $\text{LC}_{50}$  value of 5,200 ppm ( $36,000 \text{ mg}/\text{m}^3$ ), resulting in a point of departure of 1,733 ppm ( $12,000 \text{ mg}/\text{m}^3$ ), but on a probit analysis using DoseResp with rat lethality data. This study of Rowe *et al.* tested multiple concentrations and time points. Probit analysis yielded  $\text{LC}_{01}$  values for the 10-, 30min, 1-, 2-, 4- and 8hr exposure durations of 15390, 8231, 5546, 3736, 2517 and 1696 ppm (106,000, 57,000, 38,000, 26,000, 17,000,  $12,000 \text{ mg}/\text{m}^3$ ) and an n-value of 1.8 for time scaling. An interspecies uncertainty factor of 1 was applied based on similar exposure effects in humans compared with animals, and pharmacokinetic data indicating an interspecies uncertainty factor for toxicokinetic differences of less than 1 when using rat data to derive exposure values for humans. An intraspecies uncertainty factor of 3 is applied based on the previously described argument that the sensitivity for volatile anaesthetics should not vary more than



a factor of 2-3-fold. Use of a total uncertainty factor of 10 (3x3) would be too low, because the LBW values are supported by a human study in which the effects noted were milder than those defined by the LBW definition. In this study, humans exposed to 934 ppm (6,400 mg/m<sup>3</sup>) for 95 minutes experienced tightness of the frontal sinuses, increased hand perspiration, nostril irritation, congestion of Eustachian tubes, lassitude, slight mental foginess, stinging eyes, exhilaration, and/or the tip of the nose and lips anesthetized. An animal study in which rats exposed to 2,300 ppm (16,000 mg/m<sup>3</sup>) for 4 hours/day, 5 days/week for two weeks exhibited overt ataxia only following the first 4 hour exposure.

**Additional toxicological information (including relevant results of a general literature search, if any)**

Tetrachloroethylene has low acute inhalation toxicity. The liver, kidneys, blood and central nervous system (CNS) are the target organs for systemic effects. In humans the initial uptake of tetrachloroethylene following inhalation is rapid, with rates levelling off after a few hours of exposure. The available data suggest that a high proportion is absorbed in humans, but actual percentages have not been reported. Based on the effects of various volatile anaesthetics maximal sensitivity is expected in newborns (particularly prematures), pregnant women, and the elderly. The total variation, however, is not more than 2-3 fold. The least sensitive are older infants, toddlers and children compared to normal adults.

A few epidemiological studies have reported reproductive or developmental abnormalities due to exposure to tetrachloroethylene in drinking water. The results of several studies consistently indicated that occupationally exposed women might suffer higher rates of spontaneous abortion. A retrospective time-to-pregnancy study among Finnish women indicated a reduced ability to reproduce among 20 women exposed to tetrachloroethylene (concentration was not clear) 1-4 days a week or daily by inhalation. The relevance of these effects after single exposures is uncertain.

From the available data, it may be concluded that skin absorption may contribute significantly to the systemic effects of tetrachloroethylene.

H351: Suspected of causing cancer.

**Carcinogenicity and derivation of the CRP value**

IARC classification: 2A (probably carcinogenic to humans)

No carcinogenic risk potency (CRP) was derived

IARC concluded that there is evidence for consistently positive associations between exposure to tetrachloroethylene and the risks for oesophageal and cervical cancer and non-Hodgkin lymphoma. These associations appear unlikely to be due to chance, although confounding factors cannot be excluded and the total numbers in the cohort studies combined are relatively small.

**Odour and derivation of the LOA value**

Odour: solvent like, ethereal

No LOA was derived due to the absence of consistent odour perception.

Odour threshold ranging from 2 - 71 ppm (14 – 490 mg/m<sup>3</sup>)

**Other standards and guidelines (1h values in mg/m<sup>3</sup>, unless otherwise indicated)**

<b>VRW level</b> 240	<b>AEGL-1</b> 240	<b>ERPG-1</b> 690	<b>IDLH:</b> 1035 (30 minutes)
<b>AGW level</b> 1,700	<b>AEGL-2</b> 1,600	<b>ERPG-2</b> 1,380	
<b>LBW level</b> 13,000	<b>AEGL-3</b> 8,300	<b>ERPG-3</b> 6,900	