

Table 10

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This is a paper for discussion. This does not represent the views of the Committee and should not be cited.

Table 10. Repeated dose toxicity studies for PFCAs - PFNA

*Derived by contractor; ** calculated according to EFSA. (2012); NR – not reported; NA – not applicable.

Substance / CAS no. / purity / reference	Strain & species / sex / no. of animals	Dose (mg/kg bw/day) / vehicle / route of admin / duration / Guideline (GL) study) / Good Laboratory Practice (GLP) status	PFAS concentration (µg/mL / µg/g	Observed effects at LOAEL (controls vs treated groups). Recovery (controls vs treated groups).	Published NOAEL / LOAEL (mg/kg bw/day)	Study comm
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PFNA		0, 0.0125, 0.25 or 5.			
CAS No. 375-95-1	Wister rats, Male	Corn oil, Gavage, 14 days, 10/dose.	At 0.25 mg/kg bw/day Serum: 29.95.	Males (mean \pm SD): \downarrow mRNA of AKR1C1, Ugt 2B15, Cyp2C11, Cyp1A2 and Cyp2B6.	Males: 0.25 / 5
>97%.		Non-GL study, GLP not stated.	At 5 mg/kg bw/day Serum: 602.	Recovery not assessed.	
Hadrup et al. (2016)					

At high
concentrations
macroscopic
microscopic
pathological
liver as
decreased
and organ
showed
dose is
to the
not sur
the mR
severa
are dov
as seen
Hepatoc
noted w
PFAS c
Notably
these t
occur,
CYP2B6
CYP2C
down-r
PFNA, v
CYP3A2
corresp
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increas
elimina
multipl
metabo
confirm
broad s
specific
enzyme

Males (mean \pm SE):

\uparrow absolute liver weight (g):
 11.73 ± 0.23 vs 13.99 ± 0.33 .

\uparrow relative liver weight (mg/g body weight):
 34.14 ± 0.30 vs 42.12 ± 0.58 .

\downarrow TP (g/dL): 6.6 ± 0.0 vs 6.0 ± 0.1 .

\downarrow globulin (g/dL): 2.2 ± 0.0 vs 1.5 ± 0.0 .

\uparrow albumin/globulin ratio: 2.0 ± 0.0 vs 2.9 ± 0.1 .

\downarrow cholesterol (mg/dL): 117 ± 5 vs 86 ± 2 .

\downarrow TGs (mg/dL): 120 ± 9 vs 59 ± 9 .

\uparrow total bile acids ($\mu\text{mol/L}$):
 13.1 ± 2.4 vs 27.6 ± 3.1 .

\uparrow hepatocytic cytoplasmic alterations: 0 vs 10 (minimal).

\uparrow hepatocyte hypertrophy: 0

PFNA (ammonium salt)	Crl:CD-1 mice.	0, 3, 10, 30, 300 and 3000 ppm in diet equivalent to 0.27, 0.9, 2.7, 9, 27 or 270 **	NR	Males (mean):		Ammonium perfluorooctanoate led to a mice and concentrations 300 ppm weight the low tested, material be more ammonium perfluorooctanoate and the weight terms of require production change magnitude liver weight increased
				↑ absolute liver weight (g): 1.77/2.08 vs 3.01.	↑ relative liver weight (g/100g): 5.4/6.0 vs 8.4.	
CAS No.	Male and female.	Diet.		Females (mean):	Males: NA / 3*. Females: NA / 3*.	
4149-60-4	5/sex/dose.	14 days,		↑ absolute liver weight (g): 1.37/1.61 vs 2.29.		
99%.		Non-GL study,		↑ relative liver weight (g/100g): 5.7/5.7 vs 7.7.		
Kennedy Jr (1987)		GLP not stated.		Recovery not assessed.		