

AR 226-1813

3M MEDICAL DEPARTMENT, CORPORATE TOXICOLOGY

Title: Comparative Molecular Biology of Perfluorooctanesulfonate (PFOS, T-6295), N-ethyl perfluorooctanesulfonamido ethanol (N-EtFOSE, T-6316), N-Ethyl perfluorooctanesulfonamide (N-EtFOSA, T-6868), Perfluorooctanesulfonamido acetate (FOSAA, T-7071), and/or Perfluorooctanesulfonamide (FOSA, T-7132) of in Rats and Guinea Pigs following Oral dosing.

AMENDED Final Report

Date: July, 16 2004

Study Numbers. T-6295.8, T-6316.4, T-6868.2, T-7071.1, T-7132.1
Strategic Toxicology Study Number: DT-15-B

Sponsor: 3M Specialty Chemicals Division
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Study Location(s): 1. 3M Strategic Alternative Toxicology Laboratory
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In-Life Start Date	Protocol:	11/16/1998
In-Life End Date	Protocol:	11/20/1998
In-Life Start Date	Amendment #1:	03/01/1999
In-Life End Date	Amendment #1:	03/05/1999
In-Life Start Date	Amendment #2:	02/19/2001
In-Life End Date	Amendment #2:	02/23/2001

The purpose for this amendment was to change the conclusion drawn in the summary that Lin Xu and Drag Anders wrote in Appendix 6E (page 59) which read:

The original statement:

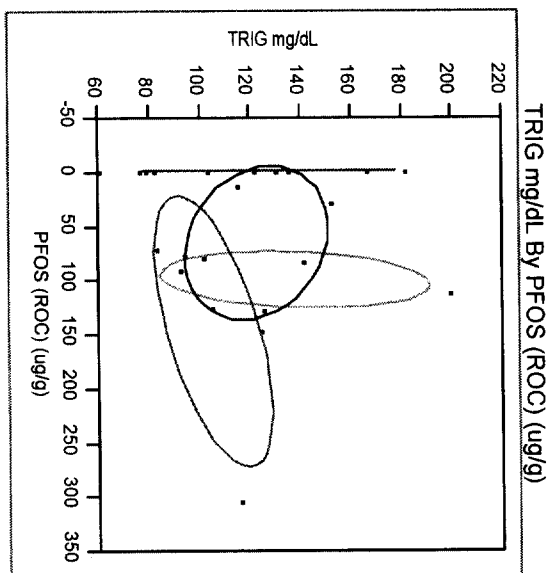
"These data showed that FOSA was consistently identified as a metabolite of PFOS, whether PFOS was administered directly or formed as a metabolite, but the source of the amino group is not readily apparent."

Was changed to read:

"These data showed that FOSA was consistently identified as a metabolite of PFOS, but that the source of the amino group is not readily apparent."

The rationale for these changes were that the original conclusion was not directly supported by the data presented in this study. That conclusion was not entirely correct because all of the administered compounds, besides PFOS, have a sulfonamido group within the parent compound. Therefore, the likely source of the amino group for sulfonamide moiety containing compounds would be the parent compound itself. There is quantitative evidence in these studies suggesting that the perfluorooctanesulfonamides were metabolized first to PFOS followed by metabolism of that PFOS back to a sulfonamide. The only evidence for a possible metabolism of PFOS to FOSA came from the PFOS dose groups, not the perfluorooctanesulfonamido derivative parent compound dose groups. Furthermore, in two out of three independent studies analyzed at Rochester, the control group liver samples had a high background of FOSA, suggesting possible contamination of the liver samples. Therefore, as stated, the conversion of PFOS to PFOSA would have to be verified by further studies.

Dr Anders is in agreement with the amended conclusion stated above.



— Bivariate Normal Ellipse P=0.500 Dose Group (mg/Kg/da=CONT
 - - - Bivariate Normal Ellipse P=0.500 Dose Group (mg/Kg/da=N-ETFOSEA40mkd
 . . . Bivariate Normal Ellipse P=0.500 Dose Group (mg/Kg/da=N-ETFOSEA40mkd
 - . - Bivariate Normal Ellipse P=0.500 Dose Group (mg/Kg/da=PFOS40mkd

Variable	Bivariate Dose Group (mg/Kg/da=CONT				Signif. Prob	Number
	Mean	Std Dev	Correlation			
PFOS (ROC) (ug/g)	0.016667	0.040825	-0.50738		0.3042	6
TRIG mg/dL	128.1667	42.64466				

Variable	Bivariate Dose Group (mg/Kg/da=N-ETFOSEA40mkd				Signif. Prob	Number
	Mean	Std Dev	Correlation			
PFOS (ROC) (ug/g)	100.4	22.36172	0.198189		0.8018	4
TRIG mg/dL	138.5	45.38355				

Variable	Bivariate Dose Group (mg/Kg/da=N-ETFOSEA40mkd				Signif. Prob	Number
	Mean	Std Dev	Correlation			
PFOS (ROC) (ug/g)	66.55	60.59266	-0.19932		0.8007	4
TRIG mg/dL	123.25	24.04683				

Variable	Bivariate Dose Group (mg/Kg/da=PFOS40mkd				Signif. Prob	Number
	Mean	Std Dev	Correlation			
PFOS (ROC) (ug/g)	148.2	106.5116	0.595121		0.4049	4
TRIG mg/dL	106	19.74842				

SRPT T-6295.8, T-6316.4, T-6868.2, T-7071.1, T-7132.1
DT15-B

Signatures:

Final Report Amendment #1 prepared by,

Andrew M. Seacat

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Study Director

7/16/04

Date