Supporting Information

Comparative uptake and biological distribution of C6 and C8 [¹⁸F]labeled perfluorinated alkyl substances in pregnant mice via different routes of administration.

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*E-mail: <u>lapi@uab.edu</u>. Telephone (205) 975-8689. Address 1824 6th Ave. South, WTI 310F, Birmingham, AL 35244 **Figure S1.** Representative images illustrating [¹⁸F]FDG uptake at (A) 0-5 min post tail vein injection, (B) 55-60 min post tail vein injection, (C) 0-5 min post oral gavage ingestion and (D) 55-60 min post oral gavage ingestion. For each letter, a representative coronal slice view (left of scale) and summed maximal intensity projection (MIP- right of scale) is given. All organs and tissues readily observed are labeled as follows: h = heart; I = liver, p = placenta, b = bladder, k = kidney, f = fetus, s = stomach and int = intestine.

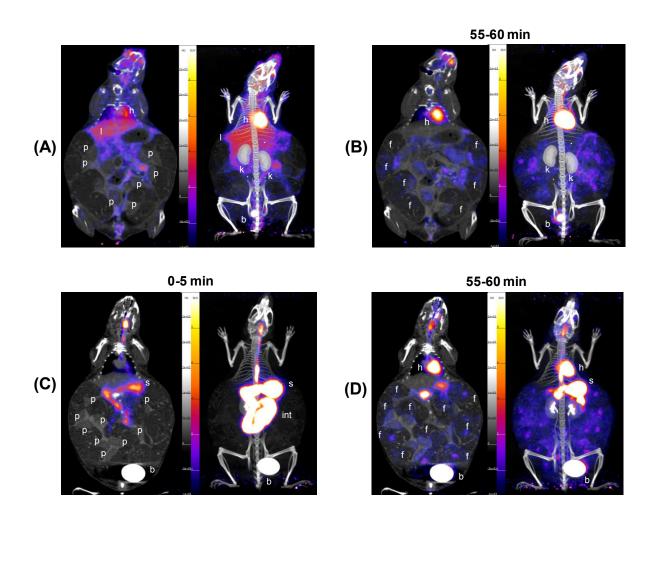


Figure S2. Representative MIP (CT only) of a mouse with regions of interest drawn within each fetus (larger spheres) and placenta (smaller spheres).

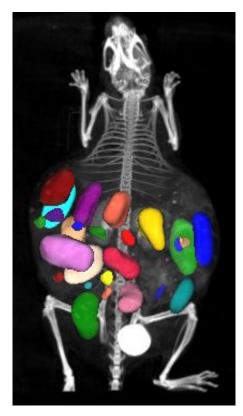
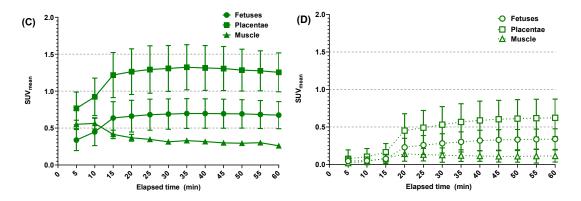


Figure S3. Representative time activity curves for the uptake of **(A)** [18 F]FDG vial tail vein and **(B)** oral gavage. The data in the fetuses (circles), placentae (squares) and muscle (triangle) are presented as an average <u>+</u> standard deviation. Solid lines and filled in shapes are indicative of tail vein studies while dotted line and outlined shapes are oral gavage studies.



TAC analysis of [¹⁸F]FDG as expected illustrated a different trend. With both studies, the uptake was slow and the tracer uptake appeared to plateau around 20-25 min after dosing. At 25 min the SUVmean in the placentae and fetuses was 1.29 ± 0.32 and 0.68 ± 0.21 while at 60 min, the values were 1.25 ± 0.26 and 0.68 ± 0.18 , respectively. As with the [¹⁸F]C8 and [¹⁸F]C8 oral gavage studies, initial uptake of [¹⁸F]FDG was lower in placentae and fetuses as well with just 0.08 ± 0.12 and 0.046 ± 0.05 , respectively. This high standard deviation is probably due to the differences in time from start of fasting to start of the individual mouse study. The longer the fasting, the more desire for the body to metabolize sugar in the form of [¹⁸F]FDG. The placentae and fetuses still followed the same trend as observed in the tail vein injection, although the tracer uptake plateau was around 45 min instead of 25 min as seen in the tail vein injection.

Table S1. Statistical comparison (P values) when comparing uptake of [¹⁸F]C8, [¹⁸F]C6, and [¹⁸F]FDG at 1 hour post **(A)** tail vein injection and **(B)** oral gavage. Red wording is indicative of significance.

(A)		[¹⁸ F]C6 vs [¹⁸ F]C8	[¹⁸ F]FDG vs [¹⁸ F]C8	[¹⁸ F]FDG vs [¹⁸ F]C6
	Blood	0.055	0.021	0.011
	Heart	0.831	0.021	0.011
	Lungs	0.394	0.043	0.011
	Pancreas	0.201	0.564	0.67
	Spleen	0.67	0.021	0.011
	Stomach	0.088	0.021	0.011
	Liver (all)	0.028	0.021	0.014
	Kidneys	0.201	1	0.055
	Sm Intestine	0.033	0.773	0.033
	L Intestine	0.011	0.021	0.201
	Fat	0.088	0.564	0.201
	Skin	0.522	0.564	0.67
	Muscle	0.201	0.149	0.088
	Femur	0.831	0.773	0.831
	Brain	0.011	0.021	0.011
	Uteruses	0.033	0.773	0.011
	Fetuses	0.019	0.266	0.005
	Placentae	0.083	<0.0001	<0.0001

(B)		[¹⁸ F]C6 vs [¹⁸ F]C8	[¹⁸ F]FDG vs [¹⁸ F]C8	[¹⁸ F]FDG vs [¹⁸ F]C6
	Blood	0.014	0.016	1
	Heart	0.014	0.009	0.014
	Lungs	0.014	0.076	0.014
	Pancreas	0.028	0.917	0.050
	Spleen	0.014	0.009	0.014
	Stomach	NA	0.117	NA
	Liver (all)	0.014	0.009	0.086
	Kidneys	0.014	0.117	0.028
	Sm Intestine	0.014	0.465	0.014
	L Intestine	0.050	0.016	0.014
	Fat	0.014	0.251	0.142
	Skin	0.014	0.917	0.014
	Muscle	0.086	0.251	0.624
	Femur	0.014	0.465	0.050
	Brain	0.221	0.009	0.014
	Uteruses	0.014	0.251	0.014
	Fetuses	0.092	0.201	0.014
	Placentae	0.528	0.012	0.006