

## **SUPPLEMENTARY FILES**

## Supplementary Table 1

Pharmacokinetics parameters derived from the plasma following 5mg and 2mg IV d6- $\alpha$ -tocopherol administration to 6 healthy (Subjects matched).

Parameter	IV d6 $\alpha$ -tocopherol		p-values
	5mg intervention	2mg intervention	
Elimination Rate ( $K_e$ )	0.0214 $\pm$ 0.002		NS
	0.0236 $\pm$ 0.001		
Half-life (h)	33.7 $\pm$ 3.0		NS
	30.0 $\pm$ 1.4		
$C_{max}$ ( $\mu$ M) <sup>1</sup>	1.06 $\pm$ 0.09 emp.		<0.001
	1.06 $\pm$ 0.05 mod.		
$T_{max}$ (h) <sup>2</sup>	0.48 $\pm$ 0.02 emp.		NS
	0.48 $\pm$ 0.04 mod.		
$AUC_{0-72}$ ( $\mu$ M-h) <sup>3</sup>	6.7 $\pm$ 0.3		NS
	7.7 $\pm$ 1.1		
$AUC_{0-8}$ ( $\mu$ M-h)	43.0 $\pm$ 4.8 emp.		<0.001
	43.0 $\pm$ 3.1 mod.		
$AUC_{0-8}$ ( $\mu$ M-h)	17.5 $\pm$ 0.9 emp.		<0.001
	17.5 $\pm$ 2.4 mod.		

Pharmacokinetics parameters derived from the plasma d6- $\alpha$ -T concentrations following 5mg and 2mg IV d6  $\alpha$ -tocopherol administration to 6 healthy (Mean  $\pm$  SEM).

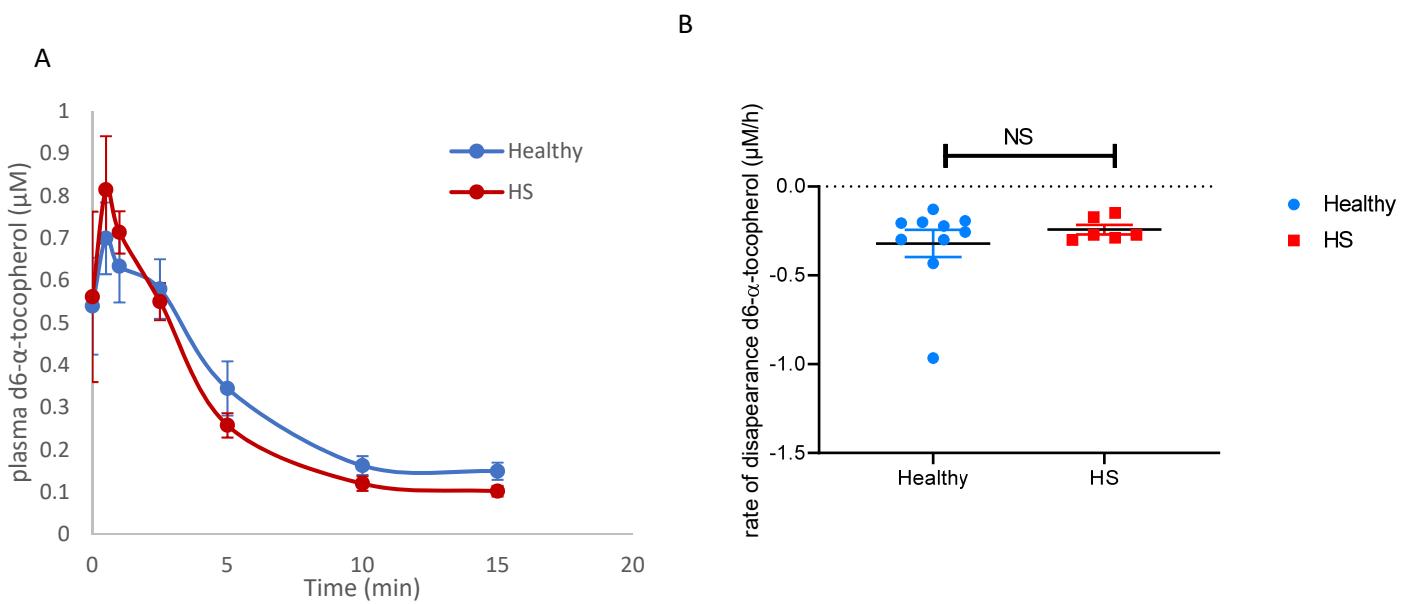
(1)  $C_{max}$ , maximum concentration post-nadir for the IV dose.

(2)  $T_{max}$ , Time of maximum concentration post-nadir for the IV dose.

(3)  $AUC_{0-72}$  is calculated from the plasma concentrations from 0 to 72 h.  $AUC_{0-8}$  is calculated from the plasma concentration from 0 to 8 h corresponding to the time of  $C_{max}$  for d6- $\alpha$ -tocopherol.  $C_{max}$ , maximal concentration;  $T_{max}$ , time of maximum concentration,  $K_e$ ; Constant of elimination; emp., empirical standard errors of the mean (SEM); mod., Mixed-effects model-based standard errors of the mean.

## Supplementary figure 1

Initial disappearance of d6- $\alpha$ -tocopherol in Healthy and HS patient.

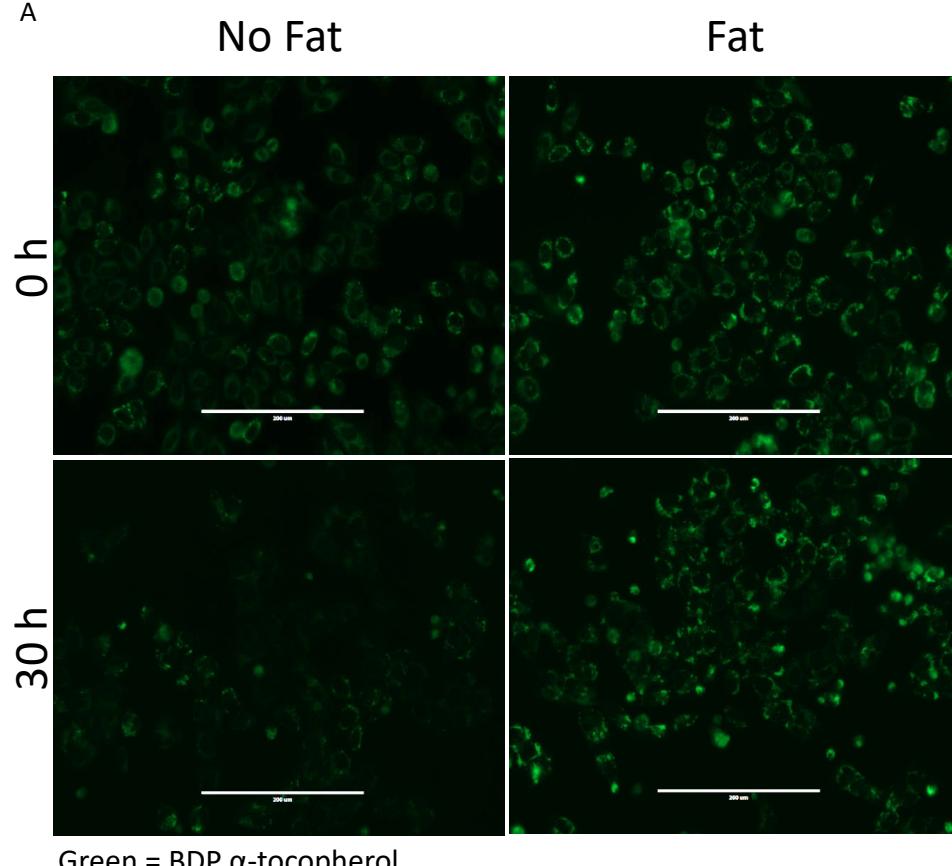


Initial disappearance of d6- $\alpha$ -tocopherol in 10 Healthy and 6 HS patient from 0 to 15min following IV injection of 2mg d6- $\alpha$ -tocopherol (a) or matched to patient status (b). Statistics . Statistics: (a) two-way ANOVA (multiple comparisons), (b) Student's t test (Unpaired, Two-tailed).

## Supplementary Figure 2

Fat traps vitamin E *in vitro*

A

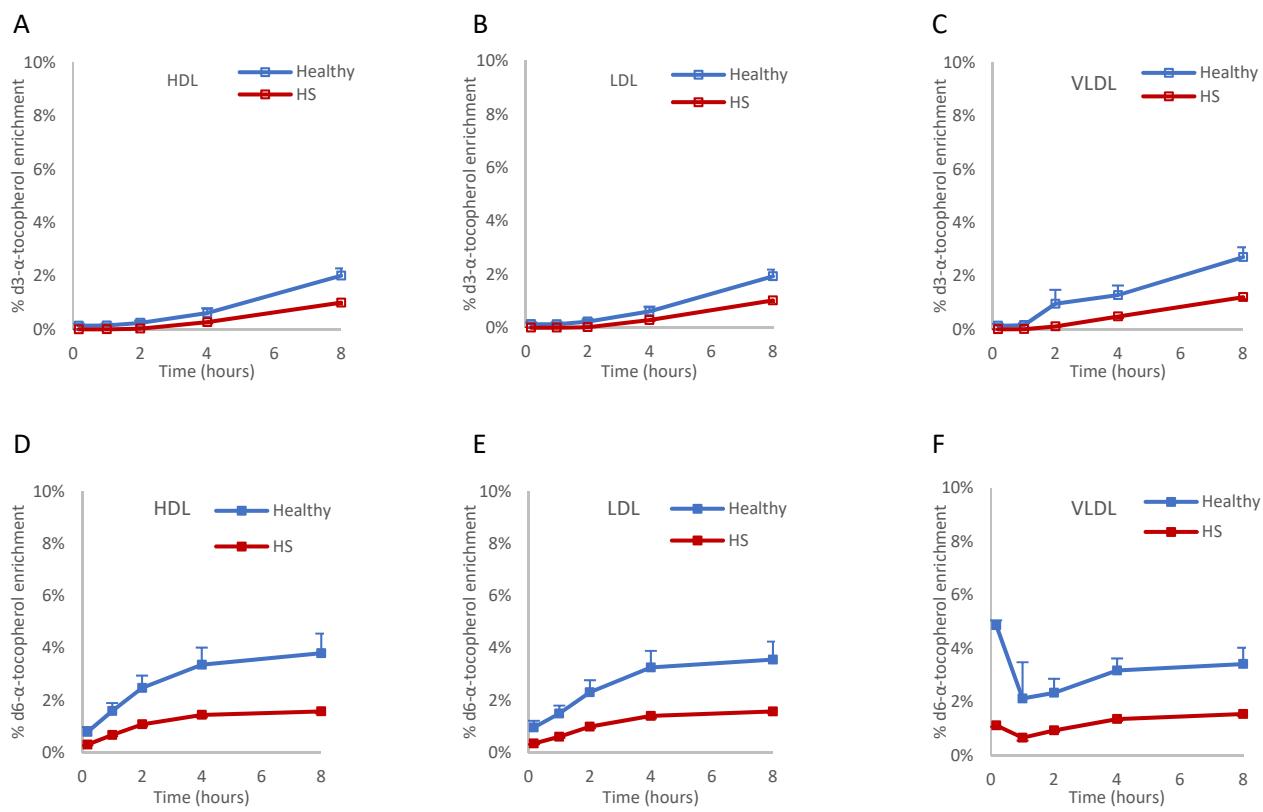


Green = BDP  $\alpha$ -tocopherol

(a) IHH cells loaded with or without 100 $\mu$ M of oleic acid for 24h, followed by incubation with 10 $\mu$ M BDP- $\alpha$ -tocopherol (green) for 2h. Images were taken at this time (0h), or 30h later after incubation in media without BDP- $\alpha$ -tocopherol.

## Supplementary Figure 3

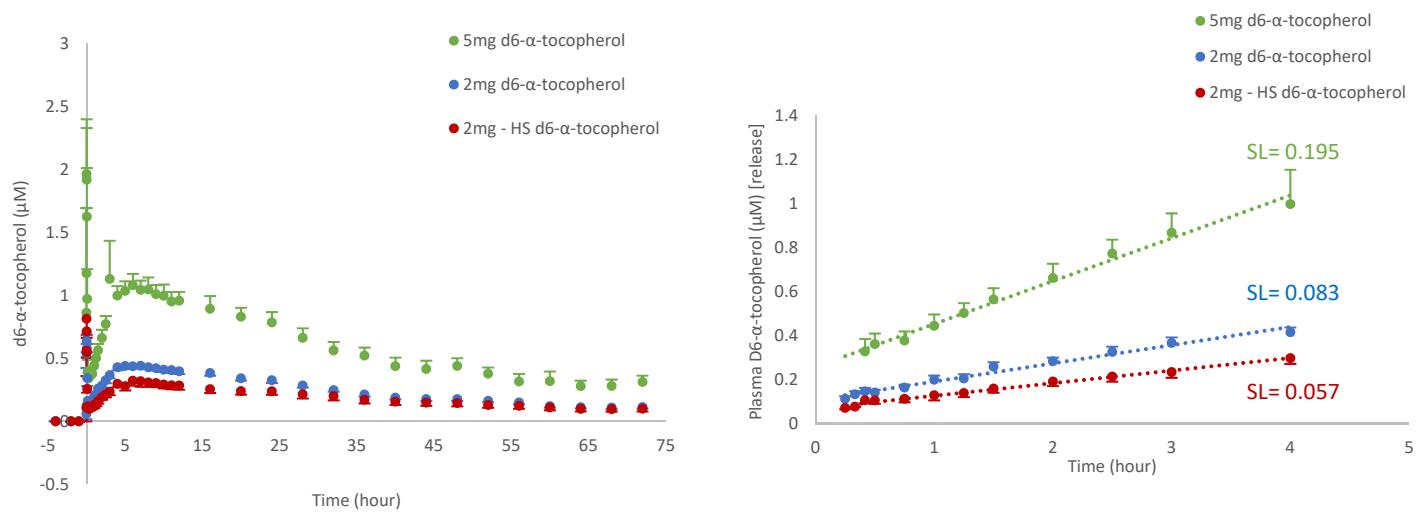
d6-, d3- $\alpha$ -Tocopherol %enrichment into lipoproteins in 10 healthy vs 6 HS women



(a-c) percent d3- $\alpha$ -tocopherol (d-f) and percent d6- $\alpha$ -tocopherol enrichment for VLDL, LDL and HDL from 0 to 8 hours in 10 healthy (□, ■) and 6 HS (□, ■) subjects following 2mg IV and oral administration of deuterated tocopherols. Statistics: two-way ANOVA (multiple comparisons).

## Supplementary figure 4

Plasma concentrations of d6- $\alpha$ -tocopherol healthy and 6 HS women



A) d6- $\alpha$ -tocopherol 5mg (●), 2mg (○) and 2mg (●) in HS subjects was administered intravenously at time zero, plasma samples obtained over 72h. (b) Initial disappearance of d6  $\alpha$ -tocopherol in Healthy (5mg (●)), 2mg (○) and 2mg in HS subjects (●) (0-72h). Data for d6- $\alpha$ -tocopherol 2 and 5mg for healthy subjects are also shown in figure 1G\_Inset. SL, Slope