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EPA prevents insulin resistance and glucose intolerance in mice fed a high fat-high sucrose diet

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Introduction:

Insulin resistance (IR) is a key feature of obesity. IR favors the progression of metabolic syndrome (MetS), increases the risk of type 2 diabetes and cardiovascular diseases. IR results from metabolic dysfunctions caused by ectopic fat depots in the liver and skeletal muscle when adipose tissue storage capacity is exceeded. Nutritional strategies using n-3 PUFA were proposed to prevent IR and MetS associated to obesity by unclear mechanisms.

Objectives:

The aim of the present work was to compare the effect of ALA, EPA and DHA on insulin resistance and metabolic dysfunctions during a high fat (HF)-high sucrose (HS) challenge.

Materials & Methods :

C57BL6 and Ob/Ob mice were fed a HF (45% energy)-HS (17% energy) diet without or with 1% energy replaced by ALA, EPA or DHA for 16 and 6 weeks respectively. IR was evaluated by intraperitoneal (ip) injection of insulin followed by monitoring of glycaemia over 2 hours. Glucose tolerance was similarly assessed using glucose injection. Fatty acid composition of tissues was determined by gas chromatography.

Results:

Figure 1. Body weight of mice during HF/HS diet

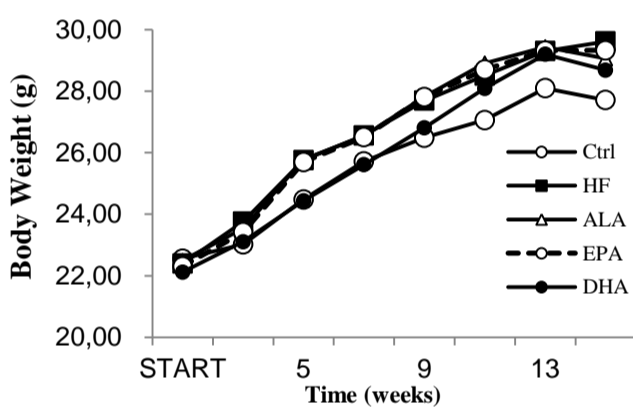


Table 1. plasma concentration of cholesterol (Chol) and non esterified fatty acids (NEFA)

	CTRL	HF	ALA	EPA	DHA
Chol (g/L)	0,72±0,03 ^c	0,92±0,04 ^a	0,83±0,08 ^{ab}	0,73±0,04 ^c	0,74±0,02 ^{bc}
NEFA (g/L)	0,30±0,01 ^{ab}	0,27±0,01 ^{bc}	0,37±0,03 ^a	0,22±0,01 ^c	0,31±0,01 ^{ab}

Figure 2. ip-insulin tolerance test.

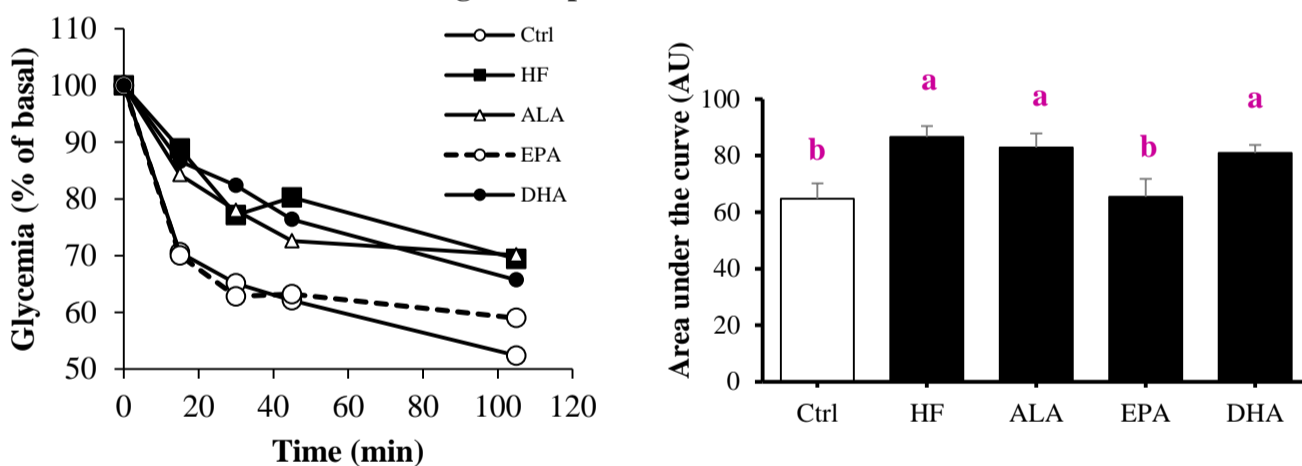


Figure 3. ip-Glucose tolerance test.

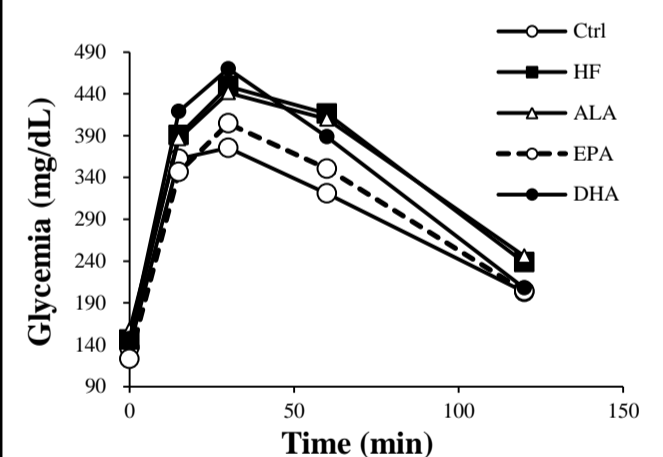


Table 2. n-3 PUFA incorporation into plasma

	(% of total fatty acids)				
	CTRL	HF	ALA	EPA	DHA
C18: 3 n-3	0,45 ± 0,05 ^a	0,31 ± 0,05 ^c	0,63 ± 0,03 ^b	0,31 ± 0,04 ^c	0,32 ± 0,04 ^c
C20: 5 n-3	0,37 ± 0,02 ^a	0,39 ± 0,05 ^a	0,77 ± 0,06 ^a	4,84 ± 0,26 ^b	2,84 ± 0,36 ^c
C22:5 n-3	0,24 ± 0,01 ^a	0,26 ± 0,02 ^a	0,35 ± 0,01 ^b	0,82 ± 0,04 ^c	0,23 ± 0,01 ^a
C22: 6 n-3	5,19 ± 0,17 ^a	6,00 ± 0,27 ^a	6,95 ± 0,19 ^b	8,07 ± 0,31 ^c	11,65 ± 0,51 ^d

Figure 4. n-3PUFA enrichment in the adipose tissue

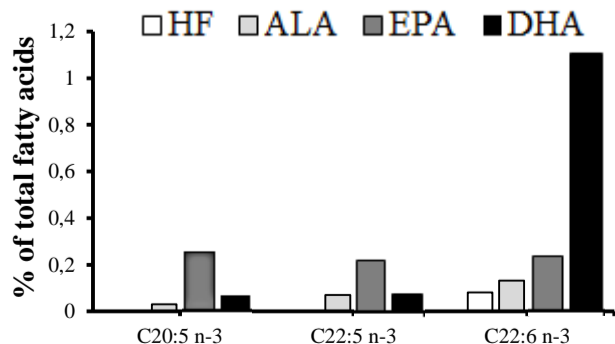
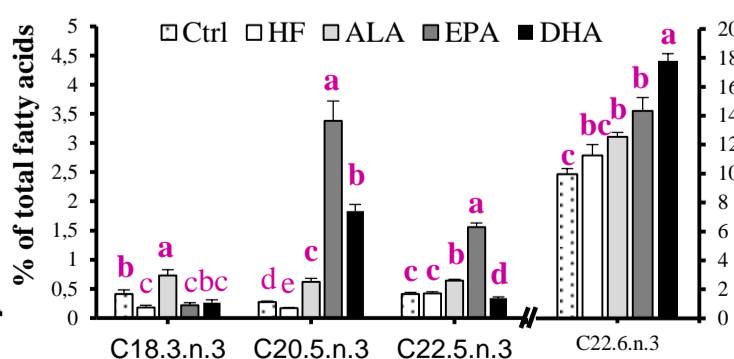


Figure 5. n-3PUFA enrichment in the liver



Conclusion :

Long chain omega 3 fatty acids have distinct effects on IR and glucose tolerance. By contrast with ALA and DHA, EPA has interesting protective effect against several key parameters of MetS.