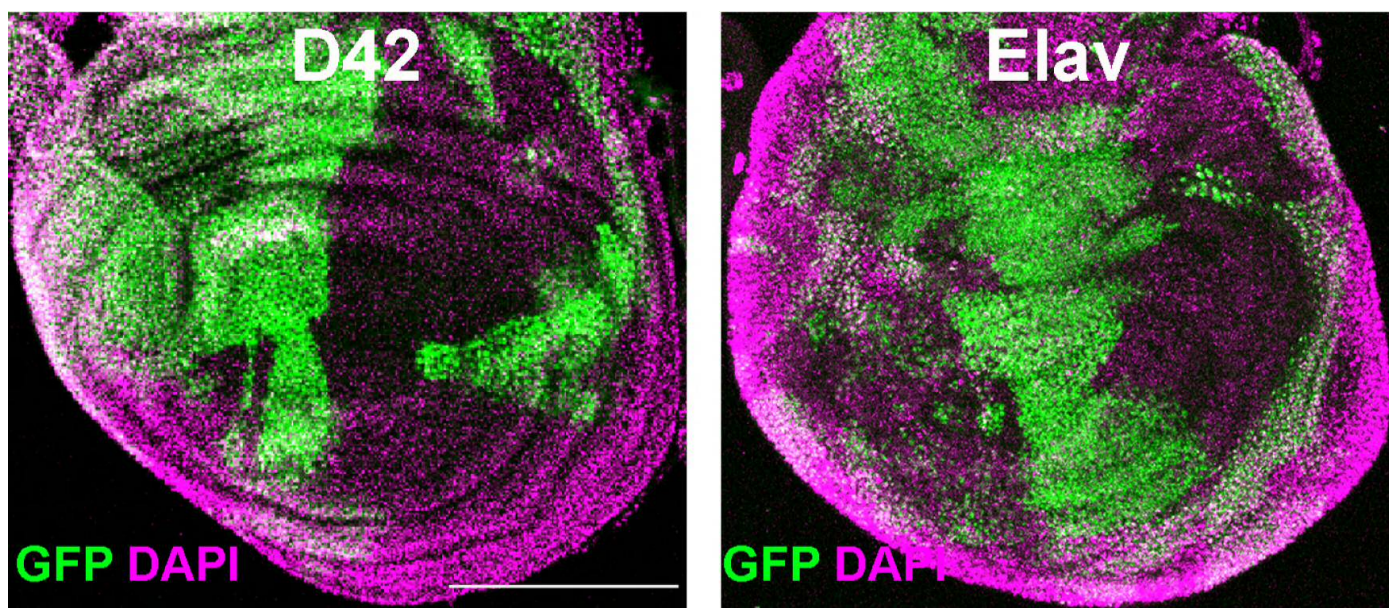
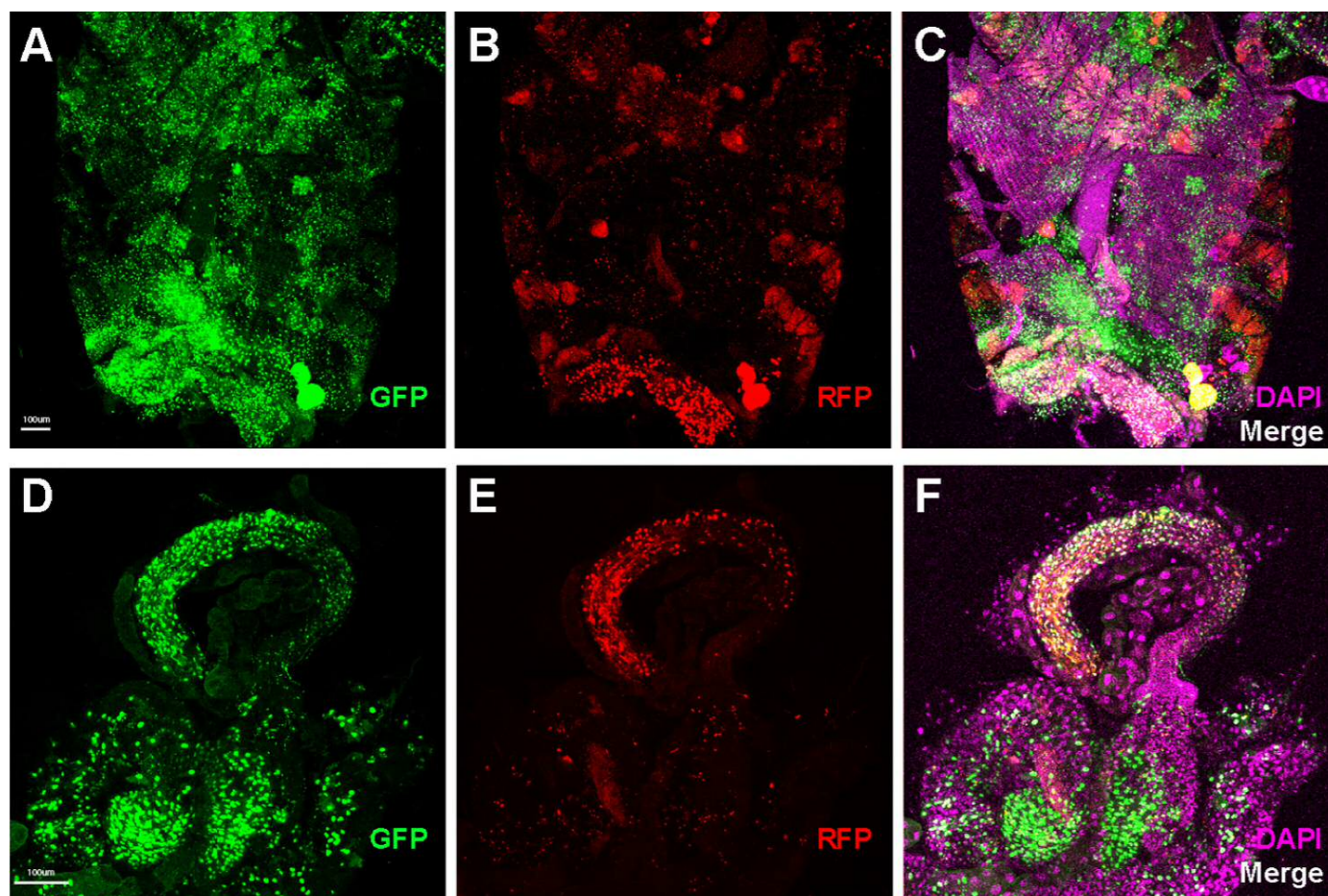


Fig. S1.- ^1H nuclear magnetic resonance (NMR) spectra of *Drosophila melanogaster* metabolites. NMR spectra of flies expressing PI3K (blue), A β 42 (red) and both A β 42 and PI3K (black) 15 days after induction of protein expression. The displayed spectra show the mean intensities of three replicates.



Suppl Fig. S2

Fig. S2.- Transient activation of Gal4 drivers in the wing disc. Using the G-TRACE technique, the initial activation of enhancers controlling the Gal4 expression can be monitored by GFP. Once activated, the GFP reporter maintains its expression in a Gal4 independent manner allowing its visualization even when the enhancer controlling the Gal4 expression is switched off. The images shown correspond to late third instar larva, a time at which the *D42* and *elav-Gal4* drivers are no longer expressed in the wing disc.



Supp. Fig. S3

Fig. S3.- Transient activation of *elav-Gal4* in the adult. Images show a squashed adult female abdomen (A-C) and thorax (D-F) 15-days post switching-on the driver. Genotype: *elav-Gal4/+* ; *UAS-RedStinger*, *UAS-FLP*, *Ubip63FRT-stop-FRT-Stinger/+* ; *tub-Gal80^{TS}/+*. The historical (green) versus the current (red) expression domains of the driver are revealed. All these cells contributed to the metabolic signature of Aβ42.

Table S1.**O2PLS-DA and OPLS-DA model statistics for parameter prediction from metabolite data**

Modeled groups	Age (days)	A[†]	N[‡]	R^{2§}	Q^{2*}
control <i>vs.</i> A β 42 <i>vs.</i> PI3K	7, 15	2+0	18	0.40	0.41
A β 42 <i>vs.</i> control and PI3K	7, 15	1+1	18	0.42	0.76
	7	1+0	9	0.46	0.81
	15	1+0	9	0.39	0.70
PI3K <i>vs.</i> control and A β 42	7, 15	1+1	18	0.42	0.64
A β 42+PI3K <i>vs.</i> control and PI3K	7, 15	1+1	18	0.52	-0.11

[†]A describes the number of model components where the first number accounts for the predictive component(s) correlating with the predicted variable, and the second the orthogonal component(s).

[‡]N describes the number of observations included in the model.

[§]R² describes how much of the total metabolite variation that is explained by the model.

*Q² represents the predictability of the total model and is related to the statistical validity of the model. Q² was calculated using cross-validation with all measurements for one condition left out at a time.