Figure 7. E-liquid flavoring agent inhibition of microsomal recombinant CYP2A6

Isoamyl Acetate		
Concentration	Mean	SEM
0.0305	2	1
0.1221	3	2
0.4883	1	1
1.9531	3	1
7.8125	3	2
31.25	4	1
125	6	2
500	10	3

Ethyl Vanillin		
Concentration	Mean	SEM
0.0305	4	1
0.1221	7	1
0.4883	6	2
1.9531	7	2
7.8125	13	2
31.25	34	2
125	69	1
500	93	0

Vanillin Mean SEM Concentration 2 0.0305 1 0.1221 4 1 5 2 0.4883 9 1 1.9531 7.8125 15 3 35 31.25 5 69 5 125 3 500 91

Benzaldehyde

Concentration	Mean	SEM
0.0305	12	2
0.1221	20	3
0.4883	29	5
1.9531	48	8

7.8125	70	6
31.25	87	3
125	96	1
500	98	1

Cinnamaldehyde

Concentration	Mean	SEM
0.0305	17	3
0.1221	22	3
0.4883	39	4
1.9531	64	5
7.8125	80	3
31.25	87	3
125	89	4
500	93	2

E-cigarette liquid flavoring agents identified in e-cigarette liquids tested earlier, as well as isoamyl acetate and benzaldehyde, were screened for inhibition of CYP2A6 at concentrations ranging from 0.03 to 500 μ M. The two smallest aromatic aldehydes, cinnamaldehyde and benzaldehyde, exhibited the strongest dose-dependent inhibition of microsomal recombinant CYP2A6. The large molecular weight aromatic aldehydes vanillin and ethyl vanillin exhibited a less potent inhibition of microsomal recombinant CYP2A6. Isoamyl acetate, an aliphatic ester, exhibited no inhibition of microsomal recombinant CYP2A6 at even the highest concentration. Results were used to calculate IC₅₀ values using GraphPad Software Inc. Prism 8.0.1. Percent inhibition calculated from the PG/VG baseline. Each concentration was screened on three separate days in duplicate (n=3). Mean +/- SEM

uM: micromolar SEM: standard error of the mean v: volume CYP2A6: cytochrome (enzyme) 2A6 IC50: 50% Inhibitory Concentration PG: polyethylene glycol VG: vegetable glycerin AW: apple watermelon SP: strawberry poptart FT: flame thrower p: p-value