

Aldehyde Detection in Electronic Cigarette Aerosols

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Supporting Information

ABSTRACT: Acetaldehyde, acrolein, and formaldehyde are the principal toxic aldehydes present in cigarette smoke and contribute to the risk of cardiovascular disease and noncancerous pulmonary disease. The rapid growth of the use of electronic cigarettes (e-cigarettes) has raised concerns over emissions of these harmful aldehydes. This work determines emissions of these aldehydes in both free and bound (aldehyde–hemiacetal) forms and other carbonyls from the use of e-cigarettes. A novel silicon microreactor with a coating phase of 4-(2-aminooxyethyl)-morpholin-4-ium chloride (AMAH) was used to trap carbonyl compounds in the aerosols of e-cigarettes via oximation reactions. AMAH–aldehyde adducts were measured using gas chromatography–mass spectrometry. ¹H nuclear magnetic resonance spectroscopy was used to analyze hemiacetals in the aerosols. These aldehydes were detected in the aerosols of all e-cigarettes. Newer-generation e-cigarette devices generated more aldehydes than the first-generation e-cigarettes because of higher battery power output. Formaldehyde–hemiacetal was detected in the aerosols generated from some e-liquids using the newer e-cigarette devices at a battery power output of 11.7 W and above. The emission of these aldehydes from all e-cigarettes, especially higher levels of aldehydes from the newer-generation e-cigarette devices, indicates the risk of using e-cigarettes.

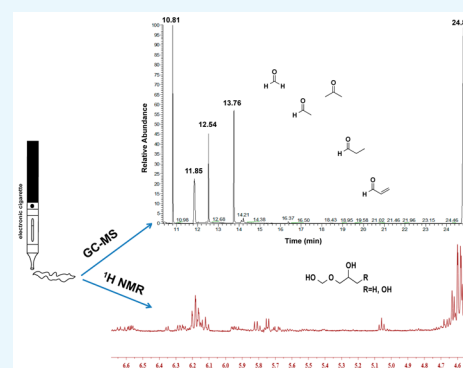


Table 1. Characteristics of e-Cigarette Cartridges and e-Liquids Used in This Study

product code	brand name	type	nicotine content (label)	manufacturer
EC01	blu Classic Tobacco	cartridge	16 mg	Imperial Tobacco
EC02	blu Magnificent Menthol	cartridge	16 mg	Imperial Tobacco
EC03	blu Vivid Vanilla	cartridge	16 mg	Imperial Tobacco
EC04	blu Cherry Crush	cartridge	16 mg	Imperial Tobacco
EL01	eVo Black Diamond	e-liquid	6 mg/mL	NicoPure Lab USA
EL02	Smooththol	e-liquid	6 mg/mL	NicQuid
EL03	Perfected Vapes/Clearwater	e-liquid	6 mg/mL	Delaware Vapor USA
EL04	Halo Café Mocha	e-liquid	6 mg/mL	Halo USA
EL05	Halo Menthol Ice	e-liquid	6 mg/mL	Halo USA
EL06	Halo South Classic	e-liquid	6 mg/mL	Halo USA

reactivity, it is difficult to quantify volatile aldehydes (and other carbonyls), and current methods that depend solely on capture by 2,4-dinitrophenylhydrazine (DNPH)-impregnated silica gel cartridges are generally criticized to be inaccurate at the low levels encountered in cigarette smoke and e-cigarette aerosols.²⁸ Moreover, a recent study reported that formaldehyde may form reversible formaldehyde–hemiacetal that may contribute to a proportion of the total aldehyde.²⁶ Unfortunately, no study has measured both free aldehydes and aldehyde–hemiacetals in the aerosols of e-cigarettes.

To address the challenge of accurately measuring aldehydes in aerosols of e-cigarettes, we used a newly developed technology that combines an enhanced carbonyl trapping agent along with a microfabricated silicon microreactor to capture aldehydes via oximation reactions. This microreactor approach has demonstrated superior lower limit of detection in an efficient and quantitative trap of trace carbonyl compounds in air and exhaled breath.^{29,30} In addition, we investigated the contribution of aldehyde-specific hemiacetals to the total aldehyde presence in the aerosols, using nuclear magnetic resonance (NMR) spectroscopy. We report herein the measurements of toxic aldehydes and their respective hemiacetals in the aerosols generated from both the first-generation e-cigarettes with cartridges and a second-generation e-cigarette with a “tank-type” atomizer for vaporization of a number of popular e-liquids.

2. MATERIALS AND METHODS

2.1. Materials. All reagents and solvents, including formaldehyde, 1,3,5-trioxane, cyclohexanone, acetone (acetone-*d*₆), and poly-4-vinylpyridine (PVP), were purchased from Sigma-Aldrich. 4-(2-Aminoxyethyl)-morpholin-4-ium chloride (AMAH) was synthesized according to a published procedure.³⁰ A popular first-generation e-cigarette blu with a fixed battery output voltage of 3.7 V along with four cartridges (coil resistance 3.0 Ω) was purchased online from Amazon. A newer-generation e-cigarette device consisting of an atomizer EVOD2 made by KangerTech (coil resistance 1.5 Ω) and an iTaste VV V3.0 battery with a digital voltage output of 3.3–5 V was purchased from a local e-cigarette store. Three popular e-liquids, EL01–EL03, were purchased from a local e-cigarette store. Each of these e-liquid products contained 10 mL of solution with a nicotine level of 6 mg/mL. Another three flavored e-liquids, EL04–EL06, were purchased from their manufacturer. These e-liquids contained 7 mL of solution with a nicotine level of 6 mg/mL. Table 1 lists the characteristics of these e-cigarette cartridges and e-liquids.

2.2. Microreactors. The silicon microreactors were fabricated from silicon wafers using standard microelectrome-

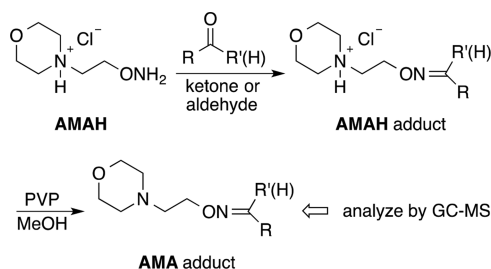
chanical system fabrication techniques. The microreactors have been used for trapping carbonyl compounds in environmental air and exhaled breath.^{29,30} Detailed fabrication and characterization procedures of the microreactors for the capture of carbonyl compounds were published elsewhere.^{29,30} Each microreactor contains thousands of micropillars. The surface of the micropillars was functionalized with AMAH by infusing a solution of AMAH (32 μL, 1 × 10⁻⁶ mol) in methanol into a microreactor followed by evaporation of the solvent in a vacuum oven. Fused silica capillary tubes with 350 μm o.d. and 250 μm i.d. were connected to the inlet and outlet ports of the microreactor with a silica-based bonding agent, respectively. The use of AMAH-coated microreactors to capture carbonyl compounds followed by conversion of AMAH adducts to neutral analytes for gas chromatography–mass spectrometry (GC–MS) analysis have been published elsewhere.³⁰

2.3. Generation of Aerosols from e-Cigarettes.

Aerosols were generated from 10 puffs (puff duration of 4 s, puff volume of 91 mL/puff, and puff frequency of 2 puffs/min) and were collected in Tedlar bags using a software-controlled (FlexiWare) cigarette-smoking robot (SCIREQ, Montreal, Canada). The puff duration, puff volume, and puff frequency in this study are within the ranges used by e-cigarette users.^{31,32} To study the effect of puffing topography on the emission of aldehydes in aerosols, polypropylene syringes with a 60 mL capacity were also used to collect aerosols of e-cigarettes by manually varying puff duration and puff volume. Whereas the first-generation e-cigarette has a fixed battery power output of 4.6 W (voltage 3.7 V), the battery power output of the second-generation e-cigarette was tested at 9.1 W (3.7 V), 11.7 W (4.2 V), 14.7 W (4.7 V), and 16.6 W (5.0 V) for vaporization of e-liquids. Aerosol samples collected in Tedlar bags were evacuated through the microreactors by a vacuum pump at a flow rate of 3.5 mL/min for the capture of carbonyl compounds. For aerosols collected from the EL04, EL05, and EL06 e-liquids at the battery power output of 14.7 and 16.6 W, the aerosol samples were diluted 50 times with N₂ and then were drawn through the microreactors because of much higher levels of generated aldehydes. After the evacuation process, the microreactor was eluted with 150 μL of MeOH followed by addition of an internal reference, AMAH–cyclohexanone (1 × 10⁻⁷ mol). Cyclohexanone was chosen because it is symmetrical (i.e., no geometrical isomers for the AMAH–cyclohexanone adduct) and was not detected in e-cigarette aerosols. Calibration curves of the internal standard against all detected aldehydes at different concentrations were established for quantitative measurements. PVP (5 mg) was added to the eluted solutions to neutralize the positively charged AMAH adducts to AMA (4-(2-aminoxyethyl)morpholine) adducts

(Scheme 1).³⁰ The suspensions were vortex-mixed for 30 s and allowed to stand for 30 min for the sedimentation of PVP particles, after which a 2 μ L aliquot was used for GC–MS analysis.

Scheme 1. Microreactor Oximation of AMAH with Carbonyl Compounds and Neutralization of Adducts with PVP before GC–MS Analyses



2.4. GC–MS Analysis of Carbonyl Adducts. A Thermo Scientific GC–MS instrument equipped with an AI 1310 automatic sampler, a TRACE 1310 GC with a split/splitless injector, and an ITQ 1100 series ion trap MS was used for analysis. The GC had an Agilent J&W DB-17ms column (60 m \times 0.25 mm \times 0.25 μ m film thickness). The flow rate of the carrier gas helium was 1.5 mL/min. The column temperature was 50 $^{\circ}$ C for 1 min, then increased by 10 $^{\circ}$ C/min up to 160 $^{\circ}$ C, and then to 200 $^{\circ}$ C by 2 $^{\circ}$ C/min. After that, the temperature was increased by 12 $^{\circ}$ C/min up to 280 $^{\circ}$ C and was held at 280 $^{\circ}$ C for 5 min. The total running time was 41 min. The samples were split-injected with a split flow of 15 mL/min and a split ratio of 10.

2.5. NMR Analysis of Hemiacetals. Ten puffs (puff volume of 35 mL, puff duration of 4 s, and puff frequency of 1 puff/min) of the aerosols generated by the first-generation e-cigarette and the second-generation e-cigarette were collected using a 60 mL polypropylene syringe with a very short rubber tube to connect the e-cigarette. After collection, the rubber tube and e-cigarette were immediately removed, and then the syringe was fitted with a stainless steel needle to transfer the aerosolized liquid into an NMR test tube in an ice bath. During the transfer, most of the aerosol condensed and was collected as liquid. Four hundred microliters of deuterated dimethyl sulfoxide- d_6 (DMSO) was added into the NMR tube followed by the addition of a known amount of benzene (1.72×10^{-6}

mol) as an internal standard. Then, 1 H NMR spectra (referenced to tetramethylsilane (TMS)) were immediately taken at 400 MHz. To verify the formation of formaldehyde–hemiacetal in e-liquids, formaldehyde was also generated by heating 1,3,5-trioxane and 8 N sulfuric acid at 95 $^{\circ}$ C accordingly³³ and then introduced as a gas into e-liquids. Formaldehyde–hemiacetal was quantified by relative integration against the known amount of benzene added as an internal standard.

2.6. Statistical Data Analysis. All measured amounts of carbonyl volatile organic compounds were analyzed using the Wilcoxon test to statistically determine the differences in the changes in puff voltage, puff volume, and puff duration of e-cigarettes. A *p*-value smaller than 0.05 was defined as statistically significant. The Wilcoxon tests were performed using Minitab version 16.0.

3. RESULTS AND DISCUSSION

3.1. Measurement of Aldehydes in Aerosols of e-Cigarettes. To measure aldehydes and other carbonyl compounds in aerosols of e-cigarettes, we used a sensitive microreactor-capture approach with an AMAH coating as previously reported (Scheme 1).³⁰ The use of microreactors with aminoxy coatings for chemoselective capture of trace carbonyl compounds via oximation reactions has been established.^{29,30} Table 2 lists the measured amounts of the four most abundant carbonyl compounds in the aerosols of the blu e-cigarettes at the battery power output of 4.6 W and the newer-generation e-cigarette at the battery power output of 9.1 W at the same battery output voltage of 3.7 V. All tested e-cigarettes produced acetaldehyde, acrolein, and formaldehyde. The amounts of acetaldehyde and formaldehyde were much higher than that of acrolein in aerosols of all e-cigarettes. The new-generation e-cigarette with the tank-type atomizer (EVOD2) and iTaste VV V3.0 battery for vaporization of the six e-liquids produced much higher levels of aldehydes and acetone than did the blu e-cigarettes because of the higher battery power output. Formaldehyde and acrolein in 10-puff aerosols generated from the six e-liquids by the newer e-cigarette device ranged from 8.2 to 40.4 μ g and 1.6 to 5.8 μ g, respectively, which were lower than these aldehydes in mainstream smoke of conventional cigarettes measured using Health Canada intense puffing regime shown in Table 2 (formaldehyde 74 μ g/cigarette and acrolein 120.4 μ g/

Table 2. Carbonyl Compounds Formed from e-Cigarettes and e-Liquids^a

product code	acetaldehyde μ g/10 puffs	acrolein μ g/10 puffs	formaldehyde μ g/10 puffs	acetone μ g/10 puffs
EC01	0.57 \pm 0.03	0.05 \pm 0.01	0.55 \pm 0.03	4.97 \pm 0.31
EC02	0.49 \pm 0.03	0.24 \pm 0.01	0.62 \pm 0.05	5.90 \pm 0.36
EC03	0.52 \pm 0.04	0.07 \pm 0.01	0.43 \pm 0.05	6.21 \pm 0.34
EC04	0.15 \pm 0.02	0.02 \pm 0.002	0.18 \pm 0.02	1.29 \pm 0.16
EL01	63.1 \pm 3.5	1.6 \pm 0.2	26.8 \pm 2.6	9.3 \pm 0.7
EL02	23.3 \pm 2.4	1.9 \pm 0.5	8.2 \pm 0.2	4.2 \pm 0.9
EL03	44.8 \pm 3.3	2.0 \pm 0.04	40.4 \pm 0.4	12.5 \pm 0.3
EL04	13.3 \pm 0.02	5.8 \pm 0.8	15.2 \pm 0.02	1.3 \pm 0.2
EL05	13.9 \pm 5.3	2.1 \pm 0.5	20.1 \pm 3.7	2.9 \pm 0.2
EL06	15.2 \pm 4.8	3.1 \pm 0.4	21.8 \pm 5.1	3.0 \pm 0.5
tobacco cigarette ^b	1240.3 \pm 17.7	120.4 \pm 14.7	74.0 \pm 23.7	641.9 \pm 71.2

^aAerosol samples were generated at a battery power output of 4.6 W for the blu e-cigarettes EC01–EC04 and a battery power output of 9.1 W for e-liquids EL01–EL06 using an EVOD2 atomizer (puff volume of 91 mL and puff duration of 4 s). Each experiment was performed in triplicate, and the data are expressed as the average (\pm SD) of the measured values. ^bData from Counts et al. (Health Canada Intense puffing regime).³⁴

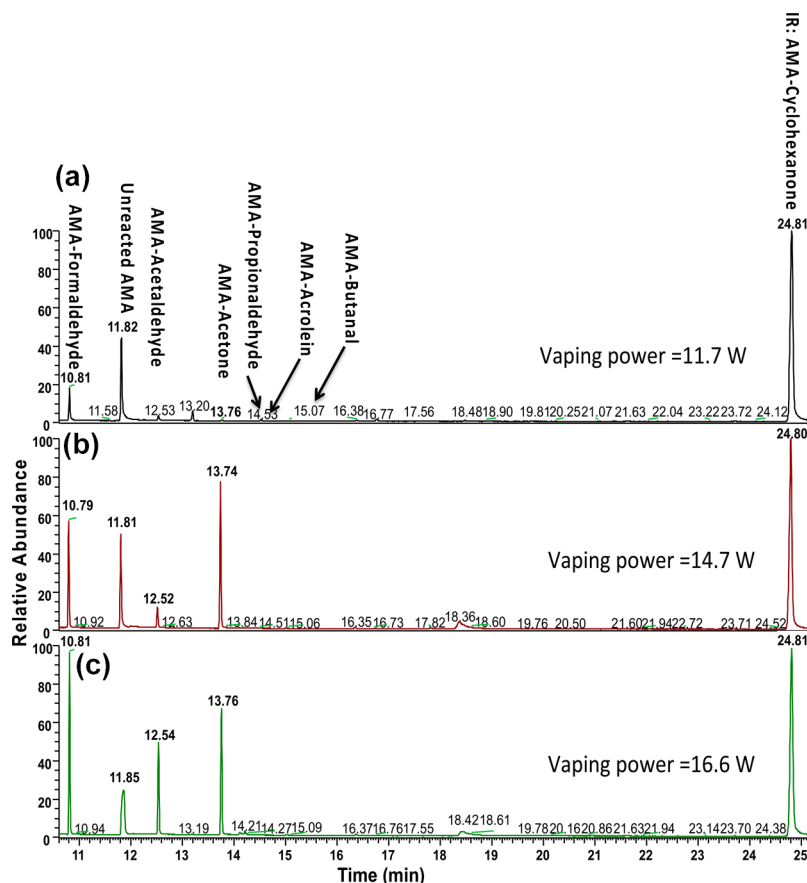


Figure 1. Representative GC–MS chromatograms of aldehydes. The newer e-cigarette device (iTaste) was used to vaporize e-liquid EL05 at a battery power output of (a) 11.7 W (4.2 V), (b) 14.7 W (4.7 V), and (c) 16.6 W (5.0 V).

cigarette).³⁴ The amounts of acetaldehyde and acetone in 10-puff aerosols of these e-liquids were much lower than those in mainstream smoke of conventional cigarettes.³⁴ Acetone in tobacco cigarette smoke has a much less toxic effect than that of the three aldehydes.^{2,3} The levels of propionaldehyde and butyraldehyde in conventional cigarette smoke are much lower than the levels of formaldehyde and acetaldehyde, and the toxic effects of these two compounds were not documented.²

Previous studies of carbonyl compounds in aerosols of e-cigarettes have found that generation of aldehydes including acrolein, formaldehyde, and acetaldehyde was related to the battery power output of the e-cigarette device and thermal decomposition of humectants and flavoring chemicals in e-liquids.^{8,35–37} Several studies have investigated the mechanisms of thermal decomposition of propylene glycol and glycerin, which are the main constituents of e-liquids.^{37–39} Thermal decomposition of propylene glycol generates more acetone, more acetaldehyde, and less formaldehyde,^{37,39} whereas thermal decomposition of glycerin results in more acrolein and formaldehyde.^{37,38} A recent study have also found that thermal decomposition of some flavoring chemicals in e-liquids can exponentially increase the generation of aldehydes.⁴⁰ Numerous studies of animal models have demonstrated the high sensitivity of pulmonary and CV systems to acrolein and formaldehyde at a relatively low level of inhalation.^{19–24} Therefore, newer-generation e-cigarettes with emissions of higher levels of formaldehyde and acrolein will need further animal studies to examine the health risk.

Because the puffing topography of e-cigarettes affects the formation of toxic aldehydes, we used a simple polypropylene syringe to imitate the characteristics of e-cigarette users by varying puff volumes and puff durations in the first-generation blu e-cigarettes with two different flavor cartridges, that is, Classic Tobacco and Magnificent Menthol. When puff volume was varied from 20 to 60 mL/puff at a fixed puff duration of 2 s, the amounts of acetaldehyde, acrolein, and formaldehyde were proportionally increased with the puff volume for both Classic Tobacco and Magnificent Menthol (Figures S1 and S2). For Magnificent Menthol, the amounts of acetaldehyde, acrolein, and formaldehyde were higher than those for Classic Tobacco at the same puff conditions (Figure S1). Varying puff duration from 1 to 6 s at a fixed puff volume of 35 mL/puff had a more complicated effect as the amount of acetaldehyde, acrolein, and formaldehyde increased when the duration was increased from 1 to 4 s but then declined as the puff duration was increased to 6 s for both Classic Tobacco and Magnificent Menthol (Figures S3 and S4). The maximum amounts of formaldehyde, acetaldehyde, and acrolein were generated at a puff duration of approximately 4 s/puff. Coincidentally, several studies of puffing topography of e-cigarette users reported average puff durations between 3.5 and 4.3 s.^{31,32}

Because increasing the battery power output of newer e-cigarette devices increases the atomizer coil temperature that results in emissions of more aldehydes in aerosols of e-liquids,^{8,26,36} we investigated the effect of the battery power output of the newer e-cigarette device on the generation of carbonyl compounds in aerosols of e-liquids. EL04 and EL05

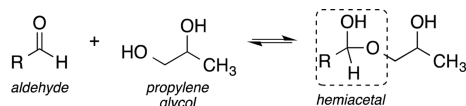
Table 3. Effect of Varying Battery Power Output on the Generation of Aldehydes in the Aerosols from e-Liquid EL05^a

voltage (W)	acetaldehyde (μg)	acrolein (μg)	formaldehyde (μg)	acetone (μg)	propionaldehyde (μg)	butyraldehyde (μg)
11.7	22.71 \pm 3.35	1.22 \pm 0.82	129.55 \pm 9.66	11.46 \pm 0.50	0.57 \pm 0.41	0.49 \pm 0.32
14.7	134.30 \pm 7.8	3.18 \pm 0.71	386.77 \pm 11.00	984.92 \pm 50.10	3.37 \pm 1.52	4.95 \pm 8.55
16.6	532.10 \pm 60.2	16.21 \pm 0.30	819.81 \pm 76.80	808.72 \pm 72.6	17.92 \pm 0.90	13.60 \pm 0.53

^aA total of 10 puffs of aerosol was collected at the puff volume of 91 mL and puff duration of 4 s. Each experiment was performed in triplicate, and the data are expressed as the average (\pm SD) of the measured values.

generated much more carbonyl compounds at a power output of 11.7 W (voltage 4.2 V) and above. Figure 1 shows GC–MS chromatograms of AMAH and its adducts from the aerosol samples collected from EL05 at the battery power output of 11.7 W (4.2 V), 14.7 W (4.7 V), and 16.6 W (5.0 V). Acetaldehyde, acrolein, formaldehyde, propionaldehyde, and butyraldehyde were detected. Increasing the power from 11.7 to 16.6 W resulted in dramatic increases in the levels of these aldehydes (Table 3). Similar results of dramatic increases in acetaldehyde, acrolein, and formaldehyde with increasing vaping power output to 9 and 10 W have been reported.³⁶ High battery power output results in overheating of the coil and leads to excessive aldehyde generation by thermal decomposition of humectants (“dry puff” condition).³⁶

3.2. Measurement of Hemiacetals in Aerosols of e-Cigarettes. Hemiacetals are formed when alcohols, such as propylene glycol in e-liquids, add reversibly to the carbonyl functional group of aldehydes, as shown in Scheme 2. The

Scheme 2. Reversible Formation of a Hemiacetal through the Reaction of an Aldehyde and an Alcohol

reaction between formaldehyde and propylene glycol or glycerin of e-cigarette liquids during vaporization is, therefore, thought to form measurable formaldehyde–hemiacetal, as detected using ¹H NMR spectroscopy.²⁶

A recent report has suggested that emission of formaldehyde in e-cigarette aerosols is higher than a direct measurement of formaldehyde because a portion of formaldehyde is sequestered in the form of a hemiacetal, which prompted more health concerns over using e-cigarettes.²⁶ Unfortunately, this work on the measurement of formaldehyde–hemiacetal did not measure free or unreacted formaldehyde or any other aldehydes in the aerosols of the e-cigarettes. The work was generally criticized for using unrealistic battery output power (voltage of 5 V) for the measurement of formaldehyde–hemiacetal.⁴¹ The reported formaldehyde–hemiacetal in aerosols was generated from e-liquid EL04. Thus, there is a need to measure total aldehydes in the aerosols of other e-cigarettes, especially for newer-generation e-cigarettes with variable battery power output and highly variable e-liquid compositions that could influence hemiacetal formation (e.g., humectant and flavoring chemicals as well as nicotine concentration). To quantify the fraction of aldehydes that reacted with propylene glycol and/or glycerin to form hemiacetals in the aerosols of e-cigarettes, we collected aerosolized e-liquids in NMR tubes containing DMSO-*d*₆ using the newer e-cigarette device. We initially attempted to detect the formation of a formaldehyde-derived hemiacetal in aerosols of all e-cigarettes in Table 1. No formaldehyde–hemiacetal

signal was detected in any of the aerosols generated from the first-generation blu e-cigarettes with all tested puff volume and puff duration scenarios (data not shown). No hemiacetal was detectable in the aerosols of EL01, EL02, and EL03 at any battery power output from 9.1 to 16.6 W as shown in Figure 2a.

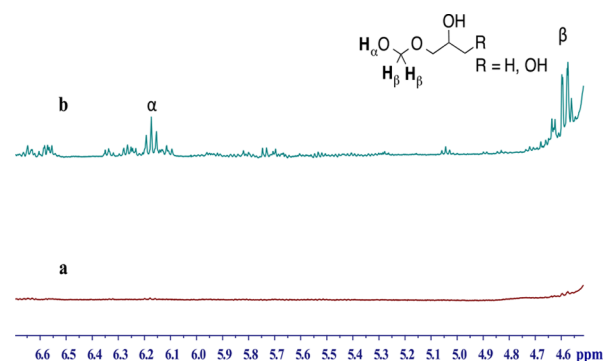


Figure 2. ¹H NMR spectra (DMSO-*d*₆) for detection of hemiacetals: (a) e-liquid EL01 vaporization at the battery power output of 16.6 W (no hemiacetal detected) and (b) e-liquid EL01 spiked with formaldehyde.

In a positive control experiment, formaldehyde gas was introduced into EL01 e-liquid, and under this condition, a triplet signal at δ 6.18 ppm and a doublet signal at δ 4.61 ppm were observed, confirming the formation of a formaldehyde–hemiacetal in this e-liquid (Figure 2b). Thus, the lack of detectable formaldehyde–hemiacetal in aerosols of the first-generation e-cigarette and e-liquids EL01, EL02, and EL03 was likely related to the generally low amount of formaldehyde present in these aerosols, even at higher battery power output.

Formaldehyde hemiacetal in aerosols of EL05 was detected at the battery power output from 11.7 to 16.6 W using NMR. No other aldehyde hemiacetal such as acetaldehyde hemiacetal and acrolein hemiacetal was detected. Figure 3 shows that there was an increase in peak intensity of the hemiacetal as the e-cigarette battery power output was increased from 11.7 to 16.6 W. The limit of detection of formaldehyde–hemiacetal using NMR was determined to be 7.1 $\mu\text{g}/10$ puffs. The formaldehyde–hemiacetal level was below the limit of detection at a battery power output of 9.1 W. The amounts of formaldehyde–hemiacetal in aerosol increased as the power was increased, and the calculated amounts (mean \pm SD) of hemiacetal were based on the internal standard (Figure 4). At a battery power output of 11.7 W, 78.6 \pm 23.8 $\mu\text{g}/10$ puffs of formaldehyde hemiacetal was measured, whereas at 16.6 W, 250.4 \pm 56.1 $\mu\text{g}/10$ puffs of the hemiacetal was measured. We were able to estimate the amount of the bound formaldehyde to be 22.2 \pm 6.7 and 70.7 \pm 15.8 μg from the measured formaldehyde–hemiacetal at 11.7 and 16.6 W, respectively. These amounts of formaldehyde could be released from the reversible reaction of formaldehyde–hemiacetal. Given the puff volume of 35 mL of

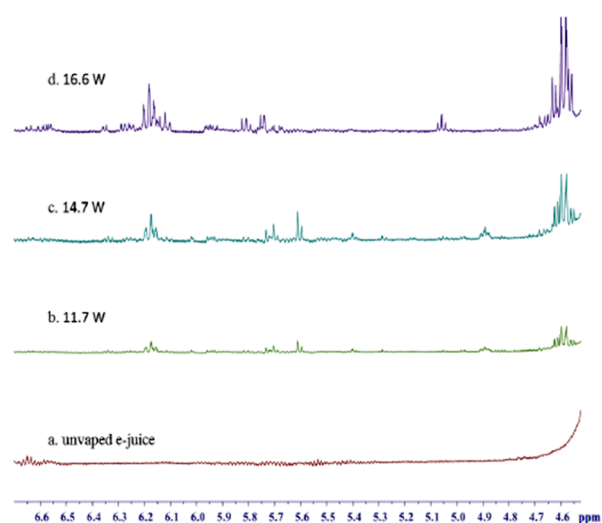


Figure 3. ^1H NMR spectra (DMSO- d_6) of hemiacetals: (a) unvaporized e-liquid EL05; the tank-type e-cigarette (iTaste) was used to vaporize the e-liquid EL05 at a battery power output of (b) 11.7 W (4.2 V), (c) 14.7 W (4.7 V), and (d) 16.6 W (5.0 V).

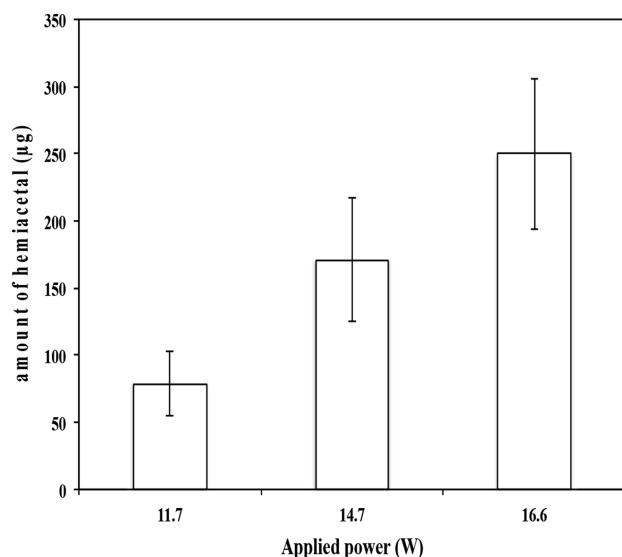


Figure 4. Relationship between the amounts of formaldehyde-hemiacetal in 10 puffs of aerosols collected from e-liquid EL05 at the battery power output of 11.7, 14.7, and 16.6 W.

this work, we could estimate that the formaldehyde in formaldehyde-hemiacetal could be approximately 44.6% of free formaldehyde at the power output of 11.7 W and approximately 22.4% of free formaldehyde at the power output of 16.6 W. Higher formaldehyde levels of 380 (puff volume of 50 mL for 10 puffs) was reported from formaldehyde-hemiacetal at the e-cigarette voltage output of 5 V for vaporization of EL04 e-liquid.²⁶

The effects of tobacco products on CVD risk are distinct from those on other organ systems. Because of their low capacity to detoxify xenobiotics,⁴² CV tissues appear to be relatively more sensitive to smoking. As a result, effects of smoking on CV tissues appear at exposure levels far lower than those that cause other diseases, such as cancer.⁴³ Moreover, in contrast to cancer, substantial CVD risk is associated even with very light smoking, such that 80% of the risk of smoking >20 cigarettes per day is associated with <3 cigarettes per day.⁴⁴

These published data suggest that even though there may be a decrease in the risk of other diseases with e-cigarette use, substantial harm to the CV system could occur because of high sensitivity of this target system to aldehydes. The emission of aldehydes including acrolein, acetaldehyde, and formaldehyde in aerosols of e-cigarettes as measured in this study supports the need to maintain an appropriate level of concern over active and passive exposures to harmful emissions of e-cigarettes and calls for more research on the newer-generation e-cigarettes and their potentially harmful effects.

■ ASSOCIATED CONTENT

§ Supporting Information

The Supporting Information is available free of charge on the ACS Publications website at DOI: 10.1021/acsomega.6b00489.

The effects of puff volume and puff duration on the amounts of aldehydes in aerosols of the first-generation e-cigarette (PDF)

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Author Contributions

The manuscript was written through contributions of all authors.

Notes

The authors declare no competing financial interest.

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