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Little Cigars vs 3R4F Cigarette: Physical Properties and HPHC Yields

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Abstract

Objective—Our objective was to characterize physical properties and semivolatile harmful and potentially harmful constituent yields in the mainstream smoke (MSS) of 4 popular little cigars compared to the 3R4F reference cigarette.

Methods—We used the ISO and Canadian Intense Regimen protocols to generate MSS for Cheyenne (Full Flavor and Menthol) and Swisher Sweets (Original and Sweet Cherry) little cigars; and the 3R4F. We examined physical properties such as length, tobacco filler mass, pressure drop,

Research leading to the preparation of this paper did not involve use of human subjects.

Conflict of Interest Statement

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Human Subjects Approval

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and ventilation for each product. Nicotine, benzo[a]pyrene, and tobacco-specific nitrosamine (TSNA) yields were determined in the MSS.

Results—Little cigars were longer (~15mm), contained more tobacco filler (100–200 mg), and had a higher pressure drop (~1.3X) compared to the 3R4F. Ventilation holes were found only on the filter paper of the 3R4F. Nicotine transmitted to the MSS was similar for all products under the intense smoking protocol. The highest yields of TSNAs and benzo(a)pyrene were measured for the little cigars.

Conclusions—Little cigars may deliver similar levels of nicotine but higher levels of carcinogens to the MSS compared to cigarettes. Thus, previous reports on the toxicity of tobacco smoke based on cigarettes might not apply to little cigar products.

Keywords

little cigar; 3R4F cigarette; tobacco physical properties; tobacco harmful and potentially harmful constituents (HPHCs)

The toxicity of tobacco smoke is the underlying cause of most tobacco related disease, which is the leading cause of preventable death in the United States (US).^{1–4} Most of the studies that discuss the toxicity, nature, and chemical constituents of tobacco smoke were derived from examining cigarette smoke; $^{2-6}$ however, there is some evidence that the use of cigar products is associated with increased risk of cancer, including lung, larynx, oral cavity and esophageal cancers.⁴ Little cigars are comparable to cigarettes with respect to shape, size, filters and packaging,⁷ but their mainstream smoke (MSS) properties are different.^{8–10} In part, this might be attributed to different tobacco blends, storage time of the tobacco leaves prior to manufacturing, and differences in the manufacturing processes of little cigars versus cigarettes.^{10–16} For instance, when compared to cigarettes, the mainstream smoke of little cigars was more basic, showing higher levels of the volatile nicotine fraction,¹⁰ the socalled "free-base" nicotine that might be more bio-active.¹⁷ Additionally, a previous study in our laboratory showed that the mainstream smoke of little cigars has some unique chemicals and some of the same, but highly concentrated chemicals compared with those measured in mainstream cigarette smoke,⁹ indicating that little cigars may have different toxicity profiles than cigarettes. Moreover, physical properties and smoking behaviors have been found to impact and change the level of the toxic/carcinogenic chemicals in the MSS of cigarettes; 5,10,18-20 but these variables have not been well examined in little cigars yet.

In 2012, the US Food and Drug Administration, Center for Tobacco Products (FDA CTP) established a list of 93 harmful and potentially harmful constituents (HPHCs) in tobacco smoke from cigarettes, pipe, and smokeless tobacco, but not from cigar products.²¹ The tobacco industry promoted cigar products, including little cigars, as alternatives to cigarettes. ^{7,22} A decrease in the consumption of cigarettes (by about 33%) was observed between 2000–2011, which was concurrent with the increase in the consumption of other non-cigarette products including cigar products (by about 123%).²³ This might be to some degree attributed to the fact that these products were not initially placed under FDA's regulatory jurisdiction as part of the 2009 Family Smoking Prevention and Tobacco Control Act.²⁴

In addition to less stringent regulation, increases in little cigar consumption also may be explained by the lower cost of these products, which can be less than half the price of cigarettes in some states with wide disparities in excise taxes levied for little cigars compared to cigarettes.²⁵ Recently, cigar products have fallen under the FDA's regulatory authority.²⁶ Studies are needed to understand the chemical constituents and toxicity of these novel tobacco products to inform regulation and protect public health. This study aims to address some of the data gaps surrounding the differences in the physical properties and yields of MSS semivolatile HPHCs between little cigars and cigarettes. To this end, we evaluated the physical properties and MSS chemistry of 4 popular little cigars: Cheyenne Full Flavor (CF), Cheyenne Menthol (CM), Swisher Sweets Original (SO), and Swisher Sweets Sweet Cherry (SC),^{27,28} and a reference comparator that is representative of a medium-delivery, American-blended cigarette (3R4F, University of Kentucky Reference Cigarette 3R4F).²⁹

METHODS

Selection of Products

Cheyenne [Full Flavor (CF), and Menthol (CM); (Cheyenne International, LLC; Grover, NC)] and Swisher Sweets [Original (SO) and Cherry (SC); (Swisher International, Inc; Jacksonville, FL)] little cigars were purchased in Columbus, OH in February and March 2015. The 3R4F reference cigarette was purchased from the University of Kentucky in 2014.

The selected little cigars in this study have been shown to hold significant shares of the US markets.^{27,28} Reports show that Swisher Sweets and Cheyenne represented about 30% of the little cigars market.³⁰

All products were stored in their original packaging, refrigerated at 4°C, and conditioned (temperature 22 ± 2 °C, and humidity $60 \pm 3\%$, according to ISO 3402:1999(E) protocol)³¹ prior to machine smoking.

Length and Circumference

The total length of each rod was measured from end to end using a digital caliper (SC-6"C, Mitutoyo). The circumference was measured by slitting each rod longitudinally. The paper was then removed, flattened, and the width was measured with a digital caliper (ISO 4387:2000(E), 2971 protocol).³² Each measurement was made on 10 replicates of each product.

Tobacco Mass and Consumed Tobacco

To measure the tobacco mass, each product was weighed individually in triplicate. The paper and filter of each product was removed and weighed separately, and the weight of the filter plus the paper was then subtracted from the total weight of the product. The weight was determined in a temperature and humidity-controlled environment ($22 \pm 2^{\circ}$ C, and $50 \pm 5\%$ relative humidity). To determine the mass of the consumed tobacco, the weight of each little cigar/cigarette was determined before and after smoking by placing the product in a 20 mL

glass vial with its corresponding lid and weighing it. The mass was determined based on 3 replicates.

Filter and Filter Paper

The thickness of the paper wrap (around the filter), along with the cross section and longitudinal section of the filters, were examined using Starrett 0–1 inch Micrometers (MS-100) and Flexbar Opti-Flex Vision System (QCZ-2000) (Figures SI5–9, and Table SI3 in the supporting information). Acetone and sulfuric acid were used to test the type of the filter of each product. Filters made of cellulose acetate should dissolve in acetone, whereas filters made of paper dissolve in sulfuric acid.³³

Pressure Drop

Pressure drop was measured following the Cores-ta N^o 41 method.³⁴ Briefly, ambient air was drawn through the rod at 4 defined flow rates (1–4 L/min) (Figure SI1 in the supporting information) using a Thomas pump, and the resulting pressure drop was measured using a Magnehelic Gage (Dwyer Institute Inc, Michigan City, IN). This method was used to measure the pressure drop of the whole rod (little cigar/cigarette), and the filter. The pressure drop of the tobacco part of the cigar/cigarette was then obtained by subtracting the pressure drop of the filter from the whole rod pressure drop (Figure SI2b). Pressure drop was measured and determined based on 3 replicates.

Total Particulate Matter (TPM): Generation and Collection

Total particulate matter (TPM) from each product was generated using a linear, 5-port smoking machine (Hawktech Model FP2000, Tri-City Machine Works, VA) and collected on standard 44 mm quartz fiber filters (QFFs) per the 2 specified methods, International Organization for Standardization/US Federal Trade Commission ISO/FTC (2-second puff duration, 35 mL puff volume, 60-second interval),³² and Canadian Intense Regimen CIR (2second puff duration, 55 mL puff volume, 30-second interval).³⁵ Before smoking, the product and the QFFs were conditioned at $22 \pm 2^{\circ}$ C, and $50 \pm 5^{\circ}$ relative humidity for at least 12 hours. The time between the conditioning and the use of product and QFFs was 7-9 days. One product was smoked per port and TPM was collected onto an individual QFF. The string cut-off method was used as described in ISO 4387:2000E protocol.³² However, the tobacco in each rod was smoked to a butt length of 41.1 - 43.2 mm, or the length of the externally visible cigarette filter over-wrap plus 8 mm. This is a difference of 5.2 ± 0.07 mm longer butt length compared to the ISO methodology.^{36,37} Filter ventilation holes were (100%) blocked/covered by tape in the case of the CIR protocol. TPM mass was obtained gravimetrically by calculating the weight differences in the QFF before and after the smoking process. The OFFs were weighed using an analytical microbalance in a humidityand temperature-controlled room ($22 \pm 2^{\circ}$ C and $50 \pm 5\%$ relative humidity).

TPM Semivolatile Organic Compound (SVOC) Analysis

Chemical analysis of the collected TPM samples for select semivolatile HPHCs was based on methods previously developed in our laboratories for similar matrices and chemical classes.^{38–40} Briefly, the collected QFFs were spiked with internal standards and extracted

with 50% dichloromethane in acetonitrile for one hour using sonication and an orbital shaker (150 rpm). The extract was then split into 2 aliquots. One aliquot was solvent exchanged into 100 mM ammonium acetate and analyzed by liquid chromatography-tandem mass spectrometry (LC-MS/MS) for the tobacco specific nitrosamines (TSNAs), N[']- nitrosonornicotine (NNN) and 4-(methylnitrosamino)-1-(3-pyridyl)-1-butanone (NNK). The second aliquot was analyzed by automated gas chromatography/mass spectrometry (GC/MS) for the remaining SVOCs including, nicotine, menthol, and benzo[a]pyrene (BaP).

Nicotine and Menthol Content Analysis in the Unburned Tobacco and Paper

To understand the distribution of nicotine and menthol in the selected products, and assess the percent transfer of nicotine and menthol to the mainstream smoke (MSS) of the little cigars compared to the 3R4F cigarette, total nicotine and menthol were measured in the unburned paper wrapper and the tobacco leaves of each product. Nicotine and menthol in the paper and tobacco were determined using methods previously reported by our laboratory.⁴¹ Briefly, each product was separated into rod (tobacco and paper) and filter. Tobacco and paper were then weighed, extracted in a solution containing isopropanol, methyl *tert*-butyl ether, 2N sodium hydroxide, and the internal standard (quinoline) via agitation using an orbital shaker (160 rpm) for 4 hours. The extracts were then analyzed for nicotine and menthol using gas chromatography with flame ionization detection (GC/FID).

RESULTS

Physical Properties of Little Cigars versus 3R4F Cigarette

Filter and wrapper materials—Filters of the 3R4F reference cigarette and the selected little cigars dissolved in acetone, suggesting all filters of the products used in this study are made of cellulose acetate.³³ However, different patterns of MSS adsorption were observed in the filter of the 3R4F compared to little cigars after smoking under ISO conditions (Figures SI6–8 in the Supporting Information). Ventilation holes were found in the filter paper of the 3R4F, whereas no holes were detected in any of the filter papers of the selected little cigars (Figure SI5). Additionally, thickness measurements of the filter overwrap paper of the little cigars showed 3 zones of different thicknesses (Figure SI9 and Table SI3), but the 3R4F filter paper showed uniform thickness (Table SI3).

The level of nicotine in the paper wrapper of the 3R4F cigarette was found to be 1.6–3.8X higher than the nicotine levels of the little cigar paper wrappers of Cheyenne (both flavors) and Swisher Sweets-SO, but was comparable to the nicotine levels in the papers of the little cigar Swisher Sweets-SC (Table 1).

Tobacco mixture/leaves appearance—Different textures of tobacco mixtures were detected between the 2 product classes, as shown in Figure SI4 in the supporting information. The tobacco leaves/mixture of the 3R4F reference cigarette had a more uniform size, color, and texture than the tobacco leaves/mixture of the little cigars. Tobacco leaves/ mixtures of the selected little cigars showed different colors, sizes, and textures among the selected flavors and brands (Figure SI4). The Swisher Sweets-SC little cigar showed the most uniform color, size, and texture compared to other little cigars (Figure SI4). Both the

3R4F and the Swisher Sweet-SC tobacco have yellow to light brown color, whereas other products, including Cheyenne (both flavors) and the Swisher Sweets-SO have multi-colored tobacco leaves including yellow, light brown, dark brown, and dark blue (Figure SI4).

Pressure drop—A positive relationship was observed ($R^2 = 0.88$ for little cigars; and $R^2 = 0.91$ for 3R4F) between the flow rate and the pressure drop across all evaluated products (Figure SI1). The pressure drop of the selected little cigars was higher than that of the 3R4F cigarette (Figures SI1). Pressure drop was also measured for both the tobacco rod and filter separately (Figure SI2b). In the 3R4F, the filter accounted for most of the resistance to draw, as the pressure drop that was measured for the filter only was about 16X higher than that of the tobacco part (Figure SI2b). In the little cigars, the tobacco rod and filter both contributed significantly to the resistance to draw. The filter of the little cigars showed a higher (1.5–2X) pressure drop than that of the tobacco (Figure SI2b).

Tobacco mass, consumed tobacco, and TPM—The average tobacco mass of the 3R4F reference cigarette measured in this study $(753 \pm 10 \text{ mg})$ was like that found in previous reports $(755-760 \text{ mg})^{29,42}$ (Table SI1). The longer butt length used in this study resulted in a smaller (~7%) mass of consumed tobacco.⁴² Table 1 shows the comparison between results for the 3R4F and the selected little cigars. The tobacco mass of the little cigars was about 100–200 mg higher than that measured for the 3R4F reference cigarette. There was no significant difference between the consumed mass of tobacco under both puffing regimens (ISO and CIR). However, the consumed tobacco mass of Cheyenne (both flavors) was higher than that of Swisher Sweets (both flavors). The consumed mass of tobacco when smoking the 3R4F was lower than all the little cigars (Table 1).

As others have found, approximately 2 times more TPM mass was collected under CIR conditions than with the ISO smoking regimen.^{43,44} TPM also was evaluated as a percentage of mass of consumed tobacco and is presented in Figure 1. Figure 1 clearly shows that the percentage of TPM per mass of consumed tobacco under the ISO smoking regimen is significantly (p < .001) lower than that determined under the CIR for all products. The little cigars showed significantly higher percentages of TPM per mass of consumed tobacco under the ISO regimen compared with the 3R4F cigarette (p < .01). There were no significant differences observed across all the little cigars (Figure 1). Under CIR, the Swisher Sweets-SO showed a significantly higher percentage of TPM per mass of consumed tobacco compared to other little cigars and 3R4F (p < .05) (Figure 1).

Length, circumference, and puff number—The circumference and length of the 3R4F measured in this study are comparable with previously published data.^{29,42,43,45} The longer butt length used for this study resulted in fewer puffs taken under both smoking ISO and CIR regimens (Table SI1).^{43,45} Table 1 shows the length, circumference and number of puffs taken for the 4 little cigars versus the 3R4F. The circumference was similar across all little cigars in this study and comparable with the 3R4F. The length of the whole rod of the little cigars was higher (~15 mm) than the length of the 3R4F, and the length of the filter part of the little cigars was about 8 mm greater than that of 3R4F (Table 1 and Figure SI2a). More puffs were taken for the little cigars compared with the 3R4F under the ISO and CIR regimen (Table 1).

Mainstream Smoke Semivolatile HPHC Yields

Mainstream smoke semivolatile HPHC yields measured under ISO and CIR conditions for the 3R4F were normalized per mass of mainstream TPM and are presented along with previously published data in Table SI1. Our results showed a slightly higher (~10%) average yield per mass of TPM than previous reports.^{42,43,45} Normalized semivolatile HPHC yields for the 3R4F obtained under ISO conditions were higher (1.4–2X) than those obtained under CIR conditions. These results agreed with the ISO versus CIR differences (1.5–2X) obtained previously⁴³ (Table SI1).

Table 2 shows the semivolatile HPHC yields per mass of TPM for the selected little cigars and 3R4F. When obtained under the ISO regimen, TSNAs (NNN, NNK) and BaP levels per mass of TPM for Cheyenne (both flavors) were comparable to those for the 3R4F. The highest TSNA levels were measured for Swisher Sweet-SO (p < .05) compared to other little cigars and the 3R4F (Table 2 and Figure 2). In contrast to the TSNAs, the lowest yield per mass of TPM for BaP was found for Swisher Sweets-SO and the highest (p < .05) for Cheyenne-CF when compared to other little cigars and the 3R4F under both smoking regimens. Under the CIR, BaP normalized yields for the 3R4F and the little cigars were not significantly different (Figure 2). Under the ISO smoking regimen, nicotine per mass of TPM obtained for 3R4F was significantly (p < .05) higher than both brands of little cigars (Table 2 and Figure 2). Additionally, the percentage of nicotine compared to the mass of TPM for 3R4F was significantly higher than that measured for the selected little cigars when using ISO (p < .001) and CIR (p < .01) (Figure SI3). The percentage of nicotine to the mass of TPM collected under ISO conditions was significantly higher (p < .05) than the percentage found when using CIR across all products (Figure SI3).

When smoking under ISO conditions, menthol was detected only in Cheyenne-CM (41.7 \pm 2.5 µg/mg TPM) (Table 2). But when smoking under CIR, menthol was detected in Cheyenne-CF (0.3 \pm 0.01 µg/mg TPM); and Swisher Sweet-SO (0.3 \pm 0.03 µg/mg TPM); along with the Cheyenne-CM (34 \pm 2.5 µg/mg TPM) (Table 2).

In general, semivolatile HPHCs measured in MSS generated under the ISO regimen for the selected products had the same pattern when compared with the measurements obtained under CIR (Figure 2). Also, semivolatile HPHC yields normalized by mass of TPM were not significantly different between the 2 puffing regimens for a given product (Figure 2).

Yields of nicotine, NNN, NNK, and BaP were also normalized per mass of consumed tobacco and are presented in Figure 3. Under the CIR regimen, like the normalization to mass of TPM results, the highest (p < .05) TSNA normalized yields were obtained for Swisher Sweets-SO. The highest nicotine (p < .05) result was obtained for the 3R4F cigarette; and the highest yield of BaP per mass of consumed tobacco was obtained for the Cheyenne-CF (p < .05) (Figure 3). In general, the yields per mass of consumed tobacco obtained under CIR were significantly (p < .05) higher than those obtained under the ISO smoking regimen (Figure 3).

To understand the amount of semivolatile HPHCs emitted per mass of nicotine that is delivered to the mainstream smoke, compounds measured under both smoking regimens

were normalized to MSS nicotine yields. Results, presented in Table SI2, showed that all normalized yields are higher for the selected little cigars compared to 3R4F cigarette. Among the selected little cigars, the highest yields of TSNAs per mass of MSS nicotine were obtained for Swisher Sweets-SO, and the highest BaP result was obtained for the Cheyenne-CF under both smoking regimens (ISO and CIR) (Table SI2).

Nicotine and Menthol Content

Nicotine content of the tobacco filler of the 3R4F cigarette was significantly higher (p < .01) than the nicotine in the tobacco filler of all the selected little cigars (Table 1). In the MSS, nicotine yield per rod (tobacco filler plus paper) was higher for the little cigars under ISO, but lower under the more intense CIR regimen compared to the 3R4F yield. An exception was the nicotine yield of the mentholated little cigar, which yielded significantly higher amounts of nicotine ($\sim 1.5X$) than the non-mentholated product of the same brand for both regimens (Table 1). The fraction of nicotine that is transmitted from the tobacco filler to the MSS was significantly (p < .01) higher for the little cigars compared to the 3R4F cigarette under ISO conditions, but levels were comparable under CIR conditions (Table 1). Across both product types, the transmission of nicotine from the tobacco filler to the MSS under CIR was significantly higher (p < .01) than that under the ISO regimen (Table 1).

Menthol was detected in the tobacco filler and paper wrapper of all the products, including the 3R4F cigarette, except for the flavored little cigar Swisher Sweets-SC (Table 1).

DISCUSSION

Physical Properties of Little Cigars and 3R4F Cigarette

Physical properties, cigarette design, and smoking behavior may influence the levels of HPHCs in the MSS.^{5,10,18–20} Previous studies have examined the effect of different physical properties and smoking regimens on the levels and toxicity of the MSS HPHCs across different brands of cigarettes.^{3,10,19,33,42,43,45,46} To understand how these variables in cigarettes are different from little cigars, we compared the physical properties of the most popular little cigars^{27,28} with the 3R4F. Our results showed that each of the selected little cigars are longer (~15 mm), and therefore, hold more tobacco mass (100–200 mg), than the 3R4F. Additionally, a higher MSS TPM mass was obtained for the selected little cigars versus the 3R4F (Table 1). Different levels of MSS TPM mass and its associated semivolatile HPHCs might have different toxicity.^{3,43}

Additionally, the selected little cigars showed different tobacco textures and colors implicating different blends that were used for manufacturing these products. The 3R4F that was used in this study is a US blend of tobacco (35.41% of flue cured "bright" tobacco, 21.62% burley, 12.07% oriental, 1.35% Maryland, and 29.55% reconstituted tobacco²⁹). The homogeneous mixture of 3R4F tobacco contrasted with the different colors, textures, and sizes of tobacco leaves of the selected little cigars (Figure SI4). These different mixtures of tobacco, suggesting different types of tobacco leaves and different manufacturing processes, may affect the levels of the MSS HPHCs.^{14,15,39–41} Thus, studying the role of different

tobacco blends on the levels and toxicity of MSS HPHCs of little cigars could help in regulating these products.

Other product design variables that could contribute to the transmission of chemicals from the tobacco filler to the MSS include filter ventilation.^{13,47,48} Our results showed that the 3R4F cigarette filter paper had ventilation holes, whereas no holes were observed in any of the selected little cigars (Figure SI5). Ventilation holes have been shown to reduce MSS yields from machine smoking by dilution.⁴⁹ Also, the filter paper manufacturing and design of the 3R4F cigarette were different from the selected little cigars (Table SI3 and Figure SI9). These observations might explain the differences in the MSS adsorption pattern in the filter (Figures SI6–SI8) after smoking and the higher pressure drop of little cigars versus the 3R4F cigarette (Figures SI1 and SI2b).

These results show some distinct differences between the physical properties of a small group of selected little cigars and those of a representative reference cigarette. Rigorous studies of the physical properties of more little cigar products are needed to understand their impact on the MSS HPHC yields. This information is important to the development and implementation of future regulations surrounding little cigars, given FDA CTP's new authority over these products.

Semivolatile HPHCs of Little Cigars and 3R4F Cigarette

The little cigars had higher yields of NNN, NNK and BaP per mass of TPM than the 3R4F, but the 3R4F cigarette showed the highest nicotine yields (Table 2 and Figure 2). Nicotine in the unburned tobacco of the 3R4F was also higher than in all the selected little cigars (Table 1). Studies have found that tobacco leaves that pass through the fermentation process (an additional aging process applied after curing tobacco) undergo chemical transformation by which the nitrogenous compounds in the tobacco leaves (eg, nicotine) interact with the ambient nitrogen oxides to form TSNAs (ie, NNN and NNK), and thus, nicotine levels decrease.^{50,54} The tobacco used in little cigars undergoes additional fermentation, whereas that used in cigarettes does not.^{15,50–53} This additional processing of the little cigar tobacco might explain the lower nicotine levels in the unburned tobacco. Decreased nicotine could influence user behaviors and the addictive nature of little cigars. This processing also may affect TSNA levels in the MSS, although the transmission efficiency of TSNAs was not determined in this study. In fact, when compared amongst the little cigars, MSS TSNA yields that were normalized to the TPM (Figure 2), and to the consumed tobacco (Figure 3), showed high variabilities. For example, Swisher Sweets-SO had the highest levels of TSNAs followed by Swisher Sweets-SC, and Cheyenne (both flavors) had MSS TSNA yields comparable to the 3R4F cigarette. This might implicate other parameters such as the type of curing and the tobacco type/blend,^{54,55} as well as the storage of the tobacco leaves (before manufacturing the product).⁴⁵ in the formation of TSNAs in the MSS. Nevertheless, these results imply that tobacco processing may play a role in the HPHCs yields in mainstream smoke from little cigars. These processes may be important to consider in regulatory efforts to reduce the level of toxicant delivery from little cigars.

The delivery of some toxicants may be related. Previous studies on US and non-US brand cigarettes have shown an inverse relationship between MSS BaP and TSNA levels,⁵⁵ a

finding attributed to the decomposition of the PAHs, including BaP, by the nitrogen oxides that result in the formation of TSNAs.^{54–56} Our data show a similar trend in that the TSNA levels are mirror images of the BaP levels for the 4 little cigar brands tested (Figures 2 and 3).

For all products, our results showed that the transmission of nicotine from the unburned tobacco to the mainstream smoke when smoking under the CIR is higher than when smoking under the ISO regimen (Table 1), ie, more intense puffing and blocked ventilation holes result in a larger fraction of the tobacco's nicotine content being transferred into the MSS. In addition, the more intense CIR regimen eliminated differences in nicotine transmission between the little cigars and the 3R4F. Under both regimens and per rod, delivery of nicotine for the little cigars was equivalent to or greater than that of the 3R4F cigarette (Table 1), results that concur with those Chen and Pankow¹⁰ reported. This finding makes little cigars attractive from a consumer's viewpoint, in that little cigars deliver just as much nicotine, if not more, per rod than cigarettes, but cost half as much. However, this relationship may be offset by the fact that little cigars also deliver a greater mass of TPM than the 3R4F (Table 1), which is associated with harshness.⁵⁷ Further studies are needed to investigate the links between the MSS chemistry of little cigars and their growing appeal in the US.²³

Menthol has been detected in non-mentholated cigarettes.⁵⁸ Our results also show the presence of menthol in the 3 non-mentholated little cigars studied. As with cigarettes, this may be done by the manufacturer to reduce the harshness of the "unflavored" little cigar smoke. Studies of cigarettes have found that menthol may be added intentionally to non-menthol cigarettes to make them more appealing, but the addition also may result from contamination/carry-over at the manufacturing facilities, or occur naturally in the tobacco leaves.^{58,62,63}

When compared with previously published data for cigarettes, the menthol level in the Cheyenne Menthol little cigar was about 12X the level of menthol in the cigarettes labeled as mentholated cigarettes.⁵⁸ We also found that the per rod nicotine delivery of the mentholated Cheyenne little cigar was substantially higher (1.5X) than the non-mentholated little cigar of the same brand. This may impact the product appeal as well as user behaviors and exposures. Menthol reduces the tobacco smoke's harshness⁵⁹ and was found to inhibit nicotine metabolism which enhances nicotine delivery and increases exposure.^{60,61} As these data are only from one little cigar, more work is needed to understand mentholation levels and nicotine delivery across a variety of mentholated little cigars.

The rise in little cigar use, particularly among youth, dictates the need to better understand the potential health implications of these tobacco products, yet, little is currently known. Cigar smoking in general has been linked to multiple serious health consequences,⁴ and regular cigar smoking, including little cigars, has been shown to account for 9000 premature deaths annually.⁶⁴ Yet, much of the public believes that cigars are safer than cigarettes.⁶⁵ Reports have indicated that cigar smoke in general is as toxic as cigarette smoke.⁴ Our data indicate that under more intense machine smoking regimens, which still may underestimate human smoking behaviors,⁶⁶ yields of toxic HPHCs from little cigars in particular are 1–4X

higher than those found in a representative reference cigarette. In addition, the delivery of nicotine at levels like those found in the reference cigarette indicate the addictive potential and appeal of little cigars. Our studies focused only on laboratory machine smoking HPHC yields across a small number of products. More studies, particularly those based on human puffing topographies and exposures, across a wider range of little cigars are needed to understand the health implications of the increasingly popular little cigars more clearly, and support the FDA's regulations of these newly deemed products.

IMPLICATIONS FOR TOBACCO REGULATION

The FDA now has broad authority to regulate or restrict the manufacturing of little cigars to protect public health.²⁶ This could include banning or restricting the use of flavors, as with cigarettes, or setting specific product standards that reduce the harm caused by use of little cigars. The FDA relies on objective, science-based evidence to inform its rulemaking. Yet, studies examining the physical properties and mainstream smoke HPHC yields of little cigars are scarce. This report shows that physical properties and MSS semivolatile HPHC yields of the little cigars tested are different from those of the 3R4F reference cigarette. These results indicate that important variables to be considered in the regulation of little cigar products include the tobacco blend, manufacturing processes (curing and fermentation), storage of tobacco leaves before manufacturing, little cigar design, and menthol content and delivery. Although these variables have been well studied in cigarettes, it is important that they be studied in cigar products to understand how the toxicity of little cigars may differ from that of cigarettes.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Figure 1. The Total Particulate Matter (TPM) Expressed as a Fraction of the Mass of Tobacco Consumed during Machine Smoking According to ISO and CIR Regimens for the Selected Little Cigars Note.

(CF= Cheyenne Full Flavor; CM = Cheyenne Menthol; SO = Swisher Sweets Original; SC = Swisher Sweets Cherry) and reference cigarette (3R4F); [*]: Statistically significant difference between ISO and CIR (p < .001); [**]: Swisher Sweet Original is statistically significantly higher than other little cigars, as well as the 3R4F (p < .05) under the CIR.

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Figure 2. Levels of Semivolatile HPHCs in the Mainstream Smoke of the Selected Little Cigars Note.

(CF = Cheyenne Full Flavor; CM = Cheyenne Menthol; SO = Swisher Sweets Original; SC = Swisher Sweets Cherry) and the reference cigarette (3R4F), expressed as a fraction of the mass of mainstream TPM (mg), generated using machine smoking under the ISO and CIR regimens; [*] indicates that the result is statistically significantly (p < .05) higher than other products.

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Figure 3. Levels of Semivolatile HPHCs in the Mainstream Smoke of Selected Little Cigars Note.

(CF = Cheyenne Full Flavor; CM = Cheyenne Menthol; SO = Swisher Sweets Original; SC = Swisher Sweets Cherry) and the reference cigarette (3R4F), expressed as a fraction of the tobacco consumed (mg) during machine smoking under the ISO and CIR regimens; [*] indicates that the result is statistically significantly (p < .05) higher than other products.

Hyaical Fundrat Hyaical Fundrat Tobeo Mass (mg) S8< ± 30	Physical Properties		Cheyenne Full Flavor	Cheyenne Menthol	Swisher Sweets Original	Swisher Sweets Sweet Cherry	3R4F
Indecordance (mg) S8.5.4)			Physical Pro	perties			
Konstand ISO 78 ± 32 957 ± 32 61 ± 85 700 ± 10 579 ± 32 The function of the control of the contro of the control of the cont	Tobacco Mass (mg)		988 ± 50	991 ± 50	805 ± 57	892 ± 20	753 ± 10
contant notice (int) CIR $72 = 57$ $88 = 10$ $688 = 63$ $688 = 17$ $568 = 53$ $568 = 53$ $688 = 17$ $568 = 53$ $568 $	Ē	ISO	787 ± 32	957 ± 3.2	641 ± 8.5	700 ± 10	579 ± 5
FM register FO [89 ± 0.6 ± 1.3] $[89 \pm 0.6 \pm 1.3]$ $[89 \pm 0.6 \pm 1.3]$ $[17 \pm 0.7]$ $[66 \pm 0.8]$ $[37 \pm 0.7]$ $[37 \pm 0.7]$ $[39 \pm 1.3]$ $[31 \pm 0.4]$ $[31 \pm $	Consumed Lobacco (mg)	CIR	772 ± 57	882 ± 10	638 ± 6.3	688 ±17	568 ± 9
		ISO	18.9 ± 0.9	22.4 ± 1.1	17.8 ± 0.7	16.6 ± 0.8	7.3 ± 0.3
It Mumber (v) It S0 115 ± 1.3 125 ± 1.0 88 ± 0.3 90 ± 0.05 70 ± 0.05 <	1 PM (mg)	CIR	40.6 ± 1.3	45.6 ± 2.2	42.9 ± 1.5	42.9 ± 1.5	31.9 ± 1.3
rut number (b) CIR $[63 \pm 1]0$ $[53 \pm 3]2$ $[13 \pm 0.4]$ $[12 \pm 0.4]$ $[90 \pm 0.0]$ Circumference (nm) 24 ± 0.1 2		ISO	11.5 ± 1.3	12.5 ± 1.0	8.8 ± 0.3	9.0 ± 0.05	7.0 ± 0.0
Citcumterene (min) 24 ± 0.1 $24 $	Puff Number (N)	CIR	16.3 ± 1.9	15.3 ± 3.3	10.3 ± 0.5	11.2 ± 0.4	9.0 ± 0.0
Length (mm) 99 ± 0.2 88 ± 0.2 99 ± 0.3 99 ± 0.2 839 ± 0.2 83 ± 0.2 81 ± 0.2 81	Circumference (mm)		24 ± 0.1	24 ± 0.1	24 ± 0.1	24 ± 0.1	24.5 ± 0.1
Hiter Length ^d (mm) 353 ± 0.4 353 ± 0.2 349 ± 0.2 349 ± 0.2 507 ± 0.0 500 ± 0.03	Length (mm)		99 ± 0.2	98.8 ± 0.2	99 ± 0.3	99 ± 0.2	83.9 ± 0.2
Nicotine Total nicotine in paper wrapper (mgg) Nicotine Nicotine </td <td>Filter Length^a (mm)</td> <th></th> <td>35.3 ± 0.4</td> <td>35.2 ± 0.2</td> <td>34.9 ± 0.3</td> <td>34.9 ± 0.2</td> <td>26.7 ± 0.0</td>	Filter Length ^a (mm)		35.3 ± 0.4	35.2 ± 0.2	34.9 ± 0.3	34.9 ± 0.2	26.7 ± 0.0
Total nicotine in paper wrapper (mg/g) 09 ± 0.1 14 ± 0.1 0.6 ± 0.1 2.2 ± 0.1 2.3 ± 0.1 Total nicotine per mass of unburned tobacco (mg/g) 8.8 ± 0.8 10.2 ± 0.6 11.7 ± 1.6 10.5 ± 0.9 16.1 ± 0.4 Total nicotine per mass of unburned tobacco (mg/g) ISO 0.8 ± 0.06 0.88 ± 0.04 0.81 ± 0.03 0.60 ± 0.03 Mainstream smoke nicotine (mg/rod) ISO 0.84 ± 0.02 1.18 ± 0.06 0.88 ± 0.04 0.81 ± 0.03 0.60 ± 0.03 Transmission of micotine (mg/rod) ISO 0.84 ± 0.02 1.18 ± 0.06 0.88 ± 0.04 0.81 ± 0.03 0.60 ± 0.03 Transmission of nicotine to smoke % ISO 1.33 ± 0.06 2.01 ± 0.07 1.52 ± 0.09 1.50 ± 0.03 1.79 ± 0.03 Transmission of nicotine to smoke % ISO 12.1 ± 1.0 11.8 ± 1.7 10.9 ± 1.1 6.4 ± 0.3 Transmission of nicotine to smoke % ISO 19.7 ± 2.5 22.4 ± 1.5 20.5 ± 3.1 20.6 ± 1.9 19.5 ± 0.03 Total mentiof per mass of unburned tobacco and paper wrapper (mg/g) 10.7 ± 2.5 22.4 ± 1.5 20.5 ± 3.1 10.5 ± 1.9 19.5 ± 0.03 10.5 ± 1.5 10.5 ± 1.5 10.5 ± 1.5 $10.5\pm1.$			Nicotin	0			
Total nicotine per mass of unburned tobacco (mg/g) 8.8 ± 0.8 10.2 ± 0.6 11.7 ± 1.6 10.5 ± 0.9 16.1 ± 0.4 Total nicotine per mass of unburned tobacco (mg/g) ISO 0.84 ± 0.02 1.18 ± 0.06 0.88 ± 0.04 0.81 ± 0.03 0.60 ± 0.03 Mainstream smoke nicotine (mg/rod) ISO 0.84 ± 0.02 1.18 ± 0.06 0.88 ± 0.04 0.81 ± 0.03 0.60 ± 0.03 Transmission of nicotine to smoke % ISO 1.33 ± 0.06 2.01 ± 0.07 1.52 ± 0.09 1.50 ± 0.03 1.79 ± 0.06 Transmission of nicotine to smoke % ISO 12.2 ± 1.3 12.1 ± 1.0 11.8 ± 1.7 10.9 ± 1.1 6.4 ± 0.3 Transmission of nicotine to smoke % ISO 12.7 ± 2.5 22.4 ± 1.5 20.5 ± 3.1 10.9 ± 1.1 6.4 ± 0.3 Transmission of nicotine to smoke % ISO 10.7 ± 2.5 22.4 ± 1.5 20.5 ± 3.1 10.9 ± 1.1 10.9 ± 1.1 Transmission of nicotine to smoke % ISO 10.9 ± 3.1 10.9 ± 1.1 10.9 ± 1.1 10.9 ± 1.1 10.9 ± 1.1 Transmission of nicotine to smoke % ISO 10.5 ± 3.1 10.9 ± 1.1 10.9 ± 1.1 $10.9 $	Total nicotine in paper wrapper (mg/g)		0.9 ± 0.1	1.4 ± 0.1	0.6 ± 0.1	2.2 ± 0.1	2.3 ± 0.1
Image: Main stream structure in (mg/rod) ISO 0.84 ± 0.02 1.18 ± 0.06 0.88 ± 0.04 0.81 ± 0.03 0.60 ± 0.03 Main stream structure (mg/rod) CIR 1.33 ± 0.06 2.01 ± 0.07 1.52 ± 0.09 1.50 ± 0.03 1.79 ± 0.06 Transmission of nicotine to struct $\sqrt[6]{6}$ ISO 1.22 ± 1.3 1.21 ± 1.0 11.8 ± 1.7 10.9 ± 1.1 64 ± 0.3 Transmission of nicotine to struct $\sqrt[6]{6}$ CIR 19.7 ± 2.5 22.4 ± 1.5 20.5 ± 3.1 20.6 ± 1.9 19.5 ± 0.8 Transmission of nicotine to struct $\sqrt[6]{6}$ Math 10.7 ± 2.5 22.4 ± 1.5 20.5 ± 3.1 20.6 ± 1.9 19.5 ± 0.8 Menthole to struct to the tot struct (mg/g) 0.08 70.9 70.9 0.05 ND^b 0.05	Total nicotine per mass of unburned tobacco (mg/g)		8.8 ± 0.8	10.2 ± 0.6	11.7 ± 1.6	10.5 ± 0.9	16.1 ± 0.4
Answer convertigation CIR 1.33 ± 0.06 2.01 ± 0.07 1.52 ± 0.09 1.50 ± 0.03 1.79 ± 0.06 Transmission of nicotine to smoke % ISO 12.2 ± 1.3 12.1 ± 1.0 11.8 ± 1.7 10.9 ± 1.1 6.4 ± 0.3 Transmission of nicotine to smoke % CIR 19.7 ± 2.5 22.4 ± 1.5 20.5 ± 3.1 20.6 ± 1.9 19.5 ± 0.8 Total menthol per mass of unburned tobacco and paper wrapper (mg/g) 0.08 70.9 0.05 ND^b 0.05	Maineteonu emolo niodin (modod)	ISO	0.84 ± 0.02	1.18 ± 0.06	0.88 ± 0.04	0.81 ± 0.03	0.60 ± 0.03
Transmission of nicotine to smoke % ISO 12.2 ± 1.3 12.1 ± 1.0 11.8 ± 1.7 10.9 ± 1.1 6.4 ± 0.3 Transmission of nicotine to smoke % CIR 19.7 ± 2.5 22.4 ± 1.5 20.5 ± 3.1 20.6 ± 1.9 19.5 ± 0.8 Menthol Menthol 70.9 0.05 ± 3.1 20.6 ± 1.9 19.5 ± 0.8 Menthol 22.4 ± 1.5 20.5 ± 3.1 20.6 ± 1.9 19.5 ± 0.8 Menthol 20.5 ± 3.1 20.6 ± 1.9 19.5 ± 0.8 Menthol 20.5 ± 3.1 20.6 ± 1.9 19.5 ± 0.8 Menthol $10.0 \pm 0.6 \pm 0.8$ $70.9 \pm 0.6 \pm 0.8$ Total menthol tor mass of unburned tobacco and paper wrapper (mg/g) 0.08 70.9 0.05 0.05 0.05 0.05	Maustream smoke moune (mgrou)	CIR	1.33 ± 0.06	2.01 ± 0.07	1.52 ± 0.09	1.50 ± 0.03	1.79 ± 0.06
$\begin{tabular}{ c c c c c c c c c c c c c c c c c c c$	Transmission of nicotine to smoke %	ISO	12.2 ± 1.3	12.1 ± 1.0	11.8 ± 1.7	10.9 ± 1.1	6.4 ± 0.3
$Menthol Total menthol per mass of unburned tobacco and paper wrapper (mg/g) 0.08 70.9 0.05 ND^b 0.05 0.05 ND^b 0.0$		CIR	19.7 ± 2.5	22.4 ± 1.5	20.5 ± 3.1	20.6 ± 1.9	19.5 ± 0.8
Total menthol per mass of unburned tobacco and paper wrapper (mg/g) 0.08 70.9 0.05 0.05			Mentho	ľ			
	Total menthol per mass of unburned tobacco and paper w	rapper (mg/g)	0.08	70.9	0.05	q ND p	0.05

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Table 1

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 a^{a} Filter material is cellulose acetate for all tested little cigars and 3R4F;

b = ND: Not detected

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Table 2

Mainstream Smoke Semivolatile HPHCs Yields Expressed as a Fraction of the Total Particulate Matter Generated (Average \pm Uncertainty^{$\frac{1}{2}$}) for the Selected Little Cigars and the 3R4F Reference Cigarette (N = 4)

Chemical Component (Yield/mg TPM)	Cheyenne Full Flavor	Cheyenne Menthol	Swisher Sweets Original	Swisher Sweets Sweet Cherry	3R4F
		OSI			
NNN (ng/mg TPM)	13.6 ± 0.7	14.8 ± 0.8	49.4 ± 2.5	22.8 ± 1.2	14.3 ± 1.1
NNK (ng/mg TPM)	14.2 ± 0.9	11.3 ± 0.6	30.0 ± 1.4	17.5 ± 1.2	11.9 ± 1.0
BaP (ng/mg TPM)	0.9 ± 0.06	0.8 ± 0.06	0.6 ± 0.05	0.7 ± 0.04	0.8 ± 0.1
Nicotine (µg/mg TPM)	44.5 ± 2.4	52.4 ± 2.4	49.2 ± 2.9	48.6 ± 3.0	80.7 ± 5.1
Menthol (µg/mg TPM)	ND^{g}	41.7 ± 2.5	ND^{a}	ND^{a}	ND^{a}
		CIR			
NNN (ng/mg TPM)	10.7 ± 0.6	12.4 ± 0.6	36.2 ± 1.4	18.5 ± 0.6	9.2 ± 0.5
NNK (ng/mg TPM)	10.8 ± 0.9	9.6 ± 0.7	23.2 ± 0.9	15.6 ± 0.6	8.7 ± 0.5
BaP (ng/mg TPM)	0.8 ± 0.05	0.6 ± 0.04	0.4 ± 0.05	0.5 ± 0.04	0.4 ± 0.02
Nicotine (µg/mg TPM)	32.8 ± 1.8	44.1 ± 2.6	35.5 ± 2.6	39.2 ± 1.2	55.9 ± 2.9
Menthol (µg/mg TPM)	0.3 ± 0.01	34.0 ± 2.5	0.3 ± 0.03	ND^{a}	ND^{a}
Note.					
c					

 a^{a} = ND: Not detected

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fCombined Standard Uncertainty = (X value/Y value) *[SQRT (x standard deviation/X value)² + (y standard deviation/Y value)²]