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# Cigarette smokers' concurrent use of smokeless tobacco: dual use patterns and nicotine exposure

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#### **Abstract**

Background—The concurrent use of cigarettes with other tobacco products, such as smokeless tobacco (SLT), is increasingly common. Extant work with cigarette smokers who also use SLT is based heavily on retrospective reports and between-group comparisons. The purpose of this study was to assess prospectively the patterns of dual users' product use and nicotine exposure on days when cigarettes were smoked exclusively (single use) versus concurrently with SLT (dual use).

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Contributors MDB conceptualised and supervised all aspects of the project, including manuscript preparation and review. SGF developed and oversaw use of the EMA software, and also assisted with manuscript preparation and review. NJF and JEOH collected the data, conducted statistical analyses and assisted with manuscript preparation and review. SK assisted with data collection, and reviewed and edited the drafts of the manuscript. GD and IH reviewed and edited the drafts of the manuscript. Author note All work was performed at West Virginia University.

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**Design**—Forty-six dual cigarette-SLT users recorded their product use in real time via ecological momentary assessment for a 2-week longitudinal design. They responded to questions about situational factors (eg, location, mood) using this same diary, and collected saliva samples each night for later cotinine measurement. At the end of this 2-week period, users reported on their reasons for and beliefs about SLT use.

**Results**—Cotinine levels were significantly higher on dual versus single use days (mean ±SEM=374.48±41.08 ng/mL vs 300.17±28.13 ng/mL, respectively; p<0.01), and the number of cigarettes logged was higher on dual versus single use days (11.13±0.98 vs 9.13±1.11, respectively; p<0.01). Product use was distinguished by situational factors, with the strongest predictor being location of use. Moreover, the most common reason for initiating (56.52%) and continuing (67.39%) SLT use was to circumvent indoor smoking restrictions.

**Conclusions**—Results support the idea of product supplementation rather than replacement among this convenience sample of dual users. For smokers whose primary motivation for SLT use involves situations where they would otherwise be tobacco free, the potential benefits of clean indoor air laws may be diminished.

## INTRODUCTION

The tobacco use landscape in the USA has been shifting over recent decades, with cigarette smoking now at an all-time low (~14%)<sup>1</sup> and use of other tobacco products either increasing or remaining stable.<sup>23</sup> Also more popular than in previous years is the concurrent use of cigarettes with other tobacco products, including loose or pouched smokeless tobacco (SLT). <sup>45</sup> Indeed, nearly 40% of current cigarette smokers have reported use of at least one other tobacco product,<sup>6</sup> with nearly 8% reporting that the other tobacco product is a form of SLT.<sup>5</sup> The use of SLT alone and in combination with cigarettes is most prevalent among young Caucasian males,<sup>7–10</sup> and this disparity is most pronounced among those who reside in rural regions (eg, West Virginia, Kentucky, Mississippi).<sup>1112</sup>

Cigarette smokers' increasing use of other tobacco products like SLT is not surprising given the myriad of challenges to smoking they have faced over these same decades, and their exposure to tobacco industry marketing that encourages the use of cigarette alternatives. The number of states with comprehensive indoor (ie, workplace, restaurants) smoke-free laws increased from 0 to 27 in the period of 2000–2015. Today, smoking bans also exist for some outdoor spaces (eg, college campuses, playgrounds), <sup>1415</sup> multiunit housing facilities, <sup>16</sup> and personal vehicles with children (for review, see Hyland et al<sup>14</sup>). In anticipation of this changing tide, both cigarette and SLT manufacturers began the development and/or marketing of SLT products for use in smoking-restricted locations. <sup>1718</sup> Advertisements for SLT products such as those pouched and spitless in nature informed smokers that they could be used 'anywhere' and 'anytime'. 19 Many smokers now report use of SLT in places where they cannot smoke. 92021 Another tactic employed by tobacco manufacturers was to tout SLT as a harm reduction product. 1822 Emphasis was put on snus, a moist snuff product with relatively low levels of tobacco-specific carcinogens.<sup>23</sup> Interestingly, whereas a minority of smokers (eg, ~11%–24%) believe that SLT products are less harmful than are cigarettes, <sup>2425</sup> a notable portion of smokers have confirmed their use of SLT as a method for quitting cigarettes.<sup>2627</sup>

Importantly, much of what is known about dual use of cigarettes and SLT is based on retrospective reports and between-group comparisons. In one such study,  $^{28}$  cigarette smokers who used SLT daily reported a lower number of cigarettes per day (CPD; n=13) than smokers who used SLT non-daily or never (n=20). Other work suggests no differences in the average number of CPD between smokers with varying levels of SLT use (daily, non-daily, not at all) (n=17-19; n=17-18) (ref 29 30, respectively). For this latter work, cotinine levels were higher for daily cigarette smokers who used SLT daily versus non-daily or never, but were comparable between those who used SLT non-daily versus never.  $^{30}$  Of course, different groups of cigarette smokers may use SLT for different reasons (eg, circumvent smoking restrictions; quit cigarettes), and consequently have different patterns of use and nicotine exposure.

The present study was designed to prospectively assess patterns of dual cigarette-SLT use via ecological momentary assessment (EMA) methods. Primary aims were to compare smokers' cigarette use and nicotine exposure (salivary cotinine levels) on days when SLT was also used (dual use days) versus on days when SLT was not used (single use days). Secondary aims were to evaluate whether contextual factors (eg, smoking-restricted vs non-restricted locations) differentiated type of product used (cigarettes vs SLT), and to describe the reasons for and beliefs about SLT use among a sample of dual users who were not currently interested in quitting smoking.

## **METHODS**

#### Inclusion/exclusion criteria

Dual cigarette-SLT users were recruited from March 2015 to May 2017 in various counties throughout West Virginia (eg, Monongalia, Marion, McDowell, Raleigh) via fliers, online postings and word of mouth. Inclusion criteria included being 18-60 years of age; smoking 5 CPD for 1 year; and using SLT 2 times per day for 4 days per week for 6 months. These cut-offs for CPD and SLT were chosen to ensure sufficient power to detect differences between single versus dual use days for primary outcomes. To verify tobacco use status, participants also were required to provide an exhaled air carbon monoxide (CO) level of 7 ppm (CoVita; Haddonfield, NJ) and a urinary cotinine >3 (NicAlert; Nymox Pharmaceutical; Hasbrouck Heights, NJ). Exclusion criteria included diagnosis of schizophrenia or bipolar disorder; current pregnancy (verified by urinalysis) or breast feeding; use of marijuana >5 days in the past month; any other illicit drug use in the past 3 months; use of alcohol >15 days in the past month; regular use of other tobacco products; or active engagement in tobacco cessation. A power analysis indicated that ~35 participants were needed to detect small-medium differences between single versus dual use days for primary outcomes (logged CPD and cotinine levels), with a desired power of 0.80 and a type I error rate of 0.05.

#### **Procedures**

Using a longitudinal study design, dual cigarette-SLT users provided responses to assessments via EMA device every day for 2 consecutive weeks. They also visited the laboratory on four occasions for screening and training (day 1), compliance checks (days 3,

9 and 15) and/or completion of questionnaires (day 15). Participants were paid \$50 on day 3, \$100 on day 9 and \$150 on day 15 for a total of \$300 for study completion.

**Baseline (day 1)**—Following informed consent, participants completed questionnaires that assessed demographics, medical and drug use history, as well as dependence on cigarettes (Fagerström Test for Cigarette Dependence; FTCD)<sup>31</sup> and SLT (Severson Smokeless Tobacco Dependency Scale).<sup>32</sup> FTCD scores range from 0 to 10: 0–2 (very low dependence), 3—4 (low dependence), 5 (medium dependence), 6–7 (high dependence) and 8–10 (very high dependence). Severson Smokeless Tobacco Dependency Scale scores range from 0 to 19, with higher scores indicating greater levels of dependence. Participants also provided urine and breath samples for verification of tobacco use status, and females were tested for pregnancy. Those deemed eligible were then trained on all study procedures outlined below, and left the laboratory with relevant supplies and instruction materials.

Assessment period (days 1–15)—For 14 consecutive days, participants used their own brand of cigarette and SLT products ad libitum. They also engaged daily with an EMA monitoring device by logging all cigarettes and/or SLT uses immediately before the product was used. For a randomly selected portion of these logged products, participants were further prompted to complete questions that addressed mood, withdrawal symptoms and situational factors. Items related to mood (eg, sad, happy, enthusiastic, bored, calm/relaxed) and withdrawal (eg, craving, difficulty concentrating, irritable) were measured using a visual analogue scale that ranged from 0 (not at all) to 100 (extremely). Situational factors were measured via multiple choice or yes/no items. Situational factors included location (eg, home, other's home, vehicle, workplace, outside, bar, restaurant, other), cigarette availability (ie, easily, with difficulty, no), smoking norms (ie, allowed, discouraged, forbidden), social context (eg, with others, others smoking, and so on) and activities (eg, working, inactive/ leisure, eating/drinking, between activities, other). The number of prompts randomly selected for these additional questions was based on participants' self-reported number of CPD at baseline in order to standardise the number of prompts across participants. Participants also answered these same questions in response to random prompts that occurred independent of product use. At the end of the day, participants again completed questionnaires (eg, withdrawal symptoms, mood), as well as tallied and logged any products used but not recorded in real time during the day. Similar sampling procedures have been outlined in extensive detail elsewhere. 3334 Participants also were required to collect the filters from all cigarettes smoked for storage in containers prelabelled for each day of the week, as well as to collect a saliva sample each night using kits provided to them.

**Study visits (days 3, 9, 15)**—At each visit, participants returned their spent cigarette filters, saliva samples and EMA device. The number of cigarette filters returned was counted and compared with the number of cigarettes logged via monitoring device. The devices were checked to evaluate compliance, with a minimum threshold of 80% responses to random prompts. 3435 If compliance was <80%, participants were counselled by staff and provided with additional device training.

During the final study visit (day 15), participants were asked to choose their reason(s) for initiating use of SLT and again for their current use of SLT from the following options: (1) to

improve health; (2) to assist with quitting smoking; (3) to use in places where I can't smoke; and (4) other. When 'other' was chosen, participants were asked to describe their other reason(s). In addition, participants were asked to report their beliefs about SLT<sup>21</sup>: (1) snuff/dip/chew products are (less/more/same/don't know) harmful than cigarettes; (2) snus products are (less/more/same/don't know) harmful than cigarettes; (3) snuff/dip/chew products (help/do not help/don't know) smokers quit cigarettes; (4) snus products (help/do not help/don't know) smokers quit cigarettes; (5) I use smokeless tobacco when I cannot smoke: (true/false/not sure).

#### **Materials**

**EMA device**—The mobile devices used were BLU Dash 5.0 (BLU Products, Doral, FL) with an Android operating system. Each device was preloaded with software customised specifically for this study (https://www.utas.edu.au/health/research/groups/school-of-medicine/behavioural-and-situational-research-group-bsrg/hbart). Participants were not able to access features of the device other than the software used for data collection.

**Saliva samples**—SalivaBio Oral Swabs (Salimetrics, State College, PA) were used for the collection of passive drool samples. Abstinence from food and drink was required for 1 hour prior to sample collection. During collection, participants were instructed to rinse their mouth with water, wait for 10 min and then place the cotton swab in their mouth for 2 min. The swab was then placed into a plastic vial and stored in the participants' freezer until their next scheduled in-person visit. These instructions were also provided via the monitoring device, which allowed for recording the completion of this task. Once returned to the lab, saliva samples were stored at  $-80^{\circ}$ C until assayed. Cotinine levels were determined by liquid chromatography with tandem mass spectrometry using extraction and processing methods described by Cappendijk and colleagues. The limit of quantification was 1 ng/mL.

#### Data analysis

**Primary outcomes**—EMA records were used to categorise study days as 'dual use' (days when both cigarettes and SET were logged) or 'single use' (days when only cigarettes were logged, there was only 1 day in which only SLT was logged). CPD and cotinine levels were averaged across days for each participant and compared between dual and single use days via dependent samples t-tests (all p<0.05).

**Secondary outcomes**—A secondary aim was to determine whether cigarettes and SLT were being used under different circumstances (eg, location, mood). A subset of participants provided a large enough sample (n=13 for 733 assessments combined) to calculate the area under the curve for the receiver operating characteristic curve (ALTC-ROC). An ALTC-ROC value was generated for each participant, then weighted by the inverse of the SE and averaged across participants to create one ALTC-ROC value. AUC-ROC values range from 0.5 (no discrimination between products) to 1.0 (complete discrimination) and describe the probability of accurately identifying which product was used based on the situational factor. These values were calculated for the following items: location (home; work; bar/restaurant/other; vehicle; or outside); smoking restrictions (forbidden; discouraged; or allowed);

craving; and affect. Affect was derived from a factor analysis of 14 mood items followed by a Varimax rotation. Data were determined to be suitable for factor analysis based on a significant Bartlett test ( $X^2(91) = 4521.3$ , p<0.001), and an overall Kaiser-Meyer-Olkin value of 0.91. All factor loadings were 0.4 and larger (range = 0.4–0.9). Affect consisted of seven of the 14 items: 'irritable', 'angry/frustrated', 'calm/relaxed', 'happy', 'miserable', 'bored' and 'enthusiastic'. Following calculation of ALTC/ROC values, weighted t-tests were performed to determine whether the obtained values were significantly larger than 0.5 (ie, chance). Analyses were performed using R statistical software (http://www.r-project.org/) in the *ROCR*, *pROC*, *psych*, *weights* and *GRArotation* libraries and outcomes were considered statistically significant when p<0.05. Another secondary aim was to describe the reasons for and beliefs about SLT use among our sample. For these questions, the proportion of participants who endorsed each response option is provided.

## **RESULTS**

### Missing data

Excluded from all analyses were individuals who failed to respond to prompts or log their product use (n=7), or to return saliva samples (n = 3) entirely. For the remaining completers, data were missing on 3.4% of study days for logs of product use and on 2.8% of study days for cotinine levels. For situational and mood items, an inconsistency between logging and the program algorithm resulted in missing data for a notable number of participants (n=33). The remaining 13 participants provided a total of 733 combined assessments for cigarette and SLT logs.

## **Participants**

Of 69 participants who consented to participate, 56 (81.2%) were enrolled and 46 (68.7%) completed the study. Seventy per cent of non-completers failed to respond to random prompts or to log their used products via EMA device. Non-completers did not differ from completers on any baseline demographic characteristic. Table 1 outlines these characteristics for those who completed the study.

#### **Primary outcomes**

**EMA-based logs of SLT use—**SLT use was logged on 7 days/week for 35.6%, 5–6 days/week for 53.3% and 3–4 days/week for 11.1% of participants. Daily SLT users logged an average of 3.38 (SD = 1.61) dips/day, while non-daily users reported using an average of 2.42 (SD = 1.44) dips/day. Of the 25 participants who reported daily use at baseline, 72.0% logged SLT on 13–14 study days and 28.0% logged SLT on 10–12 study days. The mode was 2 dips/day for daily users and 1 dip/day for non-daily users. One-third of participants logged an average of 2 or fewer dips/day, 51.1% logged 2–4 dips/day and 15.6% logged 4 or more dips/day. The mean dips/week was 17.22 (SD = 11.03).

**Dual versus single use days**—Both cigarettes and SLT products (ie, dual use) were logged on 85.0% of study days. As for single use, only cigarettes were logged on 13.7% of study days and only SLT was logged on 0.2% of study days. Logged cigarettes differed significantly between single (M = 9.13, SEM = 1.11) and dual use days (M = 11.13, SEM = 1.11)

SEM=0.98), t(25)=-3.25, p=0.00 (see figure 1). Cotinine levels also differed significantly between these days: 300.17ng/mL (SEM=28.13) for single use versus 374.49 ng/mL (SEM=41.09) for dual use, t(23)=-2.95, p=0.01 (see figure 2). These same analyses were run after removal of outliers (eg, participants with only one single use day). For this analysis, logged cigarettes did not differ significantly between single (M = 11.26, SEM = 1.27) and dual use days (M = 12.49, SEM = 1.31), t(16) = -1.69, p=0.11. Cotinine levels remained significantly different between days: 322.75 ng/mL (SEM=31.36) for single use versus 401.53 ng/mL (SEM=52.08) for dual use, t(16) = -2.43, p=0.03.

## Secondary outcomes

**Stimulus control of product use**—Figure 3 shows AUC-ROC values for situational factors, all of which were statistically significant (all p's<0.01). Cigarette use was logged most often at home (53.8%), outside (18.1%), in a vehicle (11.0%) and at work (7.7%). SLT use was logged most often at home (59.2%), at work (11.5%), outside (10.6%) and in a vehicle (9.9%). Cigarette use was logged 92.0% of the time in smoking-permitted locations and 8.0% of the time in smoking-discouraged/forbidden locations, respectively. SLT use was logged 82.4% vs 17.6% of the time in these respective locations.

**SLT reasons/beliefs**—Nearly all participants (97.8%) reported that they use SLT on at least some occasions when they cannot smoke cigarettes. Using SLT for this same purpose was the most commonly reported reason for initiating SLT use (56.5%), as well as for continuing to use SLT today (67.4%). Other reasons reported were to aide smoking cessation (10.9% initiation; 10.9% continuation), to improve health (4.4% initiation; 0.0% continuation), other (eg, 13.0% for both initiation and continuation) and more than one of these reasons (15.2% initiation; 8.8% continuation). For the option of 'other', reasons given mentioned social use, convenience and relaxation.

When asked whether traditional SLT use is as harmful as cigarette smoking, 63.0% reported 'same', 17.4% reported 'less', 13.0% reported 'more' and 6.5% reported 'don't know'. When this same question was asked for the moist product snus, 45.7% reported 'same', 19.6% reported 'less', 10.9% reported 'more' and 23.9% reported 'don't know'. Participants also were asked whether they believe that SLT helps with quitting cigarettes, with answers being 'does help' (39.1% for traditional; 23.9% for snus), 'does not help' (47.8% for traditional; 50.0% for snus) and 'don't know' (13.0% for traditional; 26.1% for snus).

## DISCUSSION

This study is the first to use a prospective longitudinal design to characterise patterns of product use and nicotine exposure among a convenience sample of dual cigarette-SLT users. For this sample, the average number of cigarettes smoked per day was higher on days when cigarettes were used concurrently with SLT than versus exclusively. Consequently, cotinine levels were higher on days when SLT was also used. Moreover, the type of product used was associated with situational factors, with the strongest situational predictor being location of use. Together, results do not support the idea of product replacement for this sample of dual users.

The dual users sampled here reported their use of SLT primarily for circumvention of indoor smoking restrictions (similar to McClave-Regan and Berkowitz).<sup>21</sup> Indeed, those enrolled were not actively trying to quit cigarettes, and very few reported that they initiated and/or continued SLT use as a cessation or harm reduction method. Additionally, about half reported that they do not believe that traditional SLT or snus assists with quitting (consistent with McClave-Regan and Berkowitz).<sup>21</sup> For smokers whose sole motivation for SLT use involves situations where they would otherwise be tobacco free, the potential benefits of clean indoor air laws may be diminished. Relative to exclusive cigarette or SLT use, dual use has been associated with more negative outcomes: increased risk of nicotine dependence, 530 decreased likelihood of successful quit attempts<sup>2630</sup> and higher rates of serious medical conditions.<sup>3738</sup> Even for smokers who use SLT specifically for harm reduction or cessation, the benefits remain unknown. Some work shows that use of SLT as a cessation aid is more likely among dual cigarette-SLT users than cigarette-only smokers.<sup>39</sup> There also exists little evidence that, in the USA, dual cigarette-SLT users switch to exclusive SLT use. 4041 In extant randomised controlled trials, poor long-term smoking cessation rates have been observed for snus relative to controls, <sup>4243</sup> Importantly, smokers may find SLT products less appealing than cigarettes because of unacceptable sensory characteristics, minimal withdrawal relief and/or poor nicotine delivery capabilities. 44-46

Notably, the large majority of these dual users stated that both traditional SLT and snus are as or more harmful than are cigarettes. This finding is consistent with other work with either cigarette smokers or SLT users. <sup>21254748</sup> Arguments have been made for the evaluation of dual use patterns within the context of SLT harm perceptions. <sup>254849</sup> That is, perhaps smokers would be more likely to replace rather than supplement cigarettes with SLT if they were told, and believed, that doing so would significantly reduce their risks of tobacco-related disease. <sup>49</sup> Still others suggest that such a promotion of SLT products would fail to result in health benefits at the population level. <sup>50</sup>

Given the observational nature of our study, we are unable to answer many of these questions that surround the dual use debate. Another study limitation includes the generalisability of results given our convenience sample. Those enrolled were overwhelmingly Caucasian males, and recruited from relatively rural geographic locations; however, this demographic reflects what has been reported repeatedly in the literature. Results also may not generalise to those with a different pattern of dual use (eg, daily SLT users who are non-daily smokers). For the assessment of SLT consumption, measures are crude relative to those for cigarettes. The size/weight and duration of a single bout of SLT use (eg, one 'pinch' of chew) were not considered, factors which likely affect nicotine exposure.

Additional work is needed to replicate these findings, and should consider examining patterns of use as a function of SLT use reasons and beliefs. That is, dual use patterns may differ by participants' motivation for using SLT, as well as their beliefs about such products in terms of harm reduction. Characterising such differences in the patterns of dual use is important for understanding user toxicant exposure and subsequent health risks. Indeed, regulators would benefit from a better understanding of the context of SLT use among smokers as they make decisions about how these products are marketed. These same ideas

might be applied to smokers' use of other products, such as electronic cigarettes, which are used at a much higher rate than SLT products.<sup>56</sup> Major LJS cigarette manufacturers are now entering the electronic cigarette market,<sup>51</sup> as they have done in the past for the SLT market. They also may be employing the same marketing strategy as that used for SLT, with some advertisements positioning electronic cigarettes as an alternative product for use in smoking-restricted situations.<sup>52</sup>

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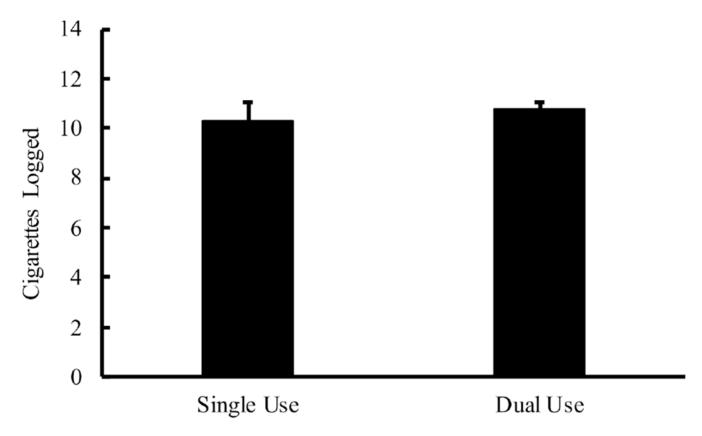
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## What this paper adds

Previous work about dual use of cigarettes and smokeless tobacco (SLT) is
primarily based on retrospective reports and between-group comparisons and
little is known about patterns of use in this population.

- This is the first study to assess prospectively the patterns of product use and nicotine exposure for dual users on days when cigarettes were smoked exclusively (single use) versus concurrently with SLT (dual use).
- Relative to single use days, dual use days revealed a larger number of cigarettes smoked and higher levels of cotinine.
- The patterns of dual use among these samples do not support the idea of product replacement.



**Figure 1.** Mean (SEM) number of cigarettes logged between single use (cigarette only) and dual use (cigarette+smokeless tobacco (SLT)) days, which were significantly different at p<0.05.

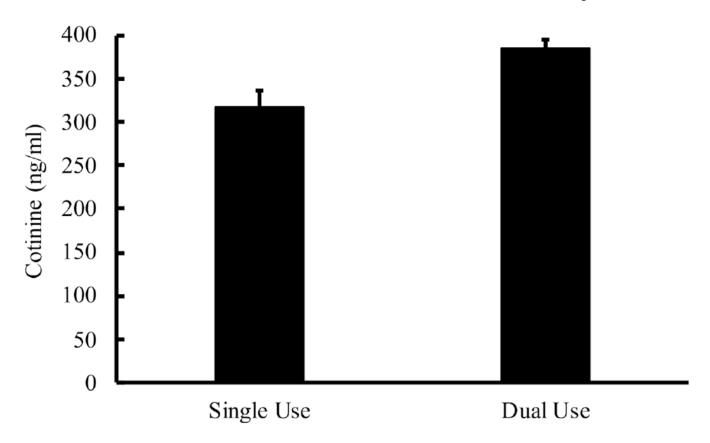
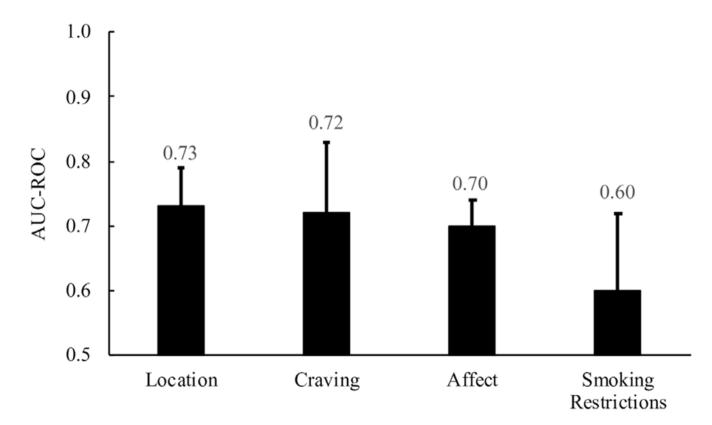


Figure 2.

Mean (SEM) cotinine levels between single use (cigarette only) and dual use (cigarette +smokeless tobacco (SLT)) days, which were significantly different at p<0.05.



**Figure 3.** AUC-ROC (SEM) values across situational domains(all p's<0.01). AUC-ROC, area under the curve for the receiver operating characteristic curve.

Table 1

## Participants' baseline characteristics

	M (SD) or %
Male	93.5%
Non-Hispanic, Caucasian	97.8%
Age (years)	30.39 (8.99)
Cigarettes per day	19.50 (8.63)
Years smoking	10.74 (6.47)
Expired air CO (ppm)	25.52 (14.63)
FTCD score*	6.02 (2.50)
SLT products	
Snuff/dip/chew	80.4%
Snus	6.5%
Multiple	13.0%
Wintergreen/mint	65.2%
SLT uses/day	4.27 (2.21)
SLT days/week	5.74 (1.54)
Years SLT use	9.32 (7.94)
SSTDS score †	8.30 (3.75)

<sup>\*</sup>Fagerström Test for Cigarette Dependence  $(0-10)^{31}$ 

SLT, smokeless tobacco.

 $<sup>^{\</sup>dagger}$ Severson Smokeless Tobacco Dependency Scale (0–19) $^{32}$