

UC Office of the President

Recent Work

Title

Nicotine, Cotinine, and Tobacco-Specific Nitrosamines Measured in Children's Silicone Wristbands in Relation to Secondhand Smoke and E-cigarette Vapor Exposure

Permalink

<https://escholarship.org/uc/item/3qq4h2kh>

Journal

Nicotine & Tobacco Research, 23(3)

ISSN

1462-2203

Authors

Quintana, Penelope JE
Lopez-Galvez, Nicolas
Dodder, Nathan G
[et al.](#)

Publication Date

2021-02-16

DOI

10.1093/ntr/ntaa140

Peer reviewed

Nicotine, cotinine and tobacco-specific nitrosamines (TSNAs) measured in children's silicone wristbands in relation to secondhand smoke and e-cigarette vapor exposure

Journal:	<i>Nicotine & Tobacco Research</i>
Manuscript ID	NTR-2020-347.R1
Manuscript Type:	Original Investigation
Date Submitted by the Author:	n/a
Complete List of Authors:	<p>Quintana, Penelope; San Diego State University, School of Public Health Lopez-Galvez, Nicolas; San Diego State University Research Foundation Dodder, Nathan; San Diego State University Research Foundation Hoh, Eunha ; San Diego State University, School of Public Health Matt, Georg; San Diego State University, Department of Psychology Zaskarian, Joy; San Diego State University Research Foundation, (retired) Chu, Linda; San Diego State University, School of Public Health Vyas, Mansi; San Diego State University, School of Public Health Akins, Brittany; San Diego State University, School of Public Health Padilla, Samuel; San Diego State University Research Foundation Anderson, Kim; Oregon State University College of Agricultural Sciences, Environmental and Molecular Toxicology Hovell, Melbourne; San Diego State University, School of Public Health</p>
Keywords:	Exposure assessment, Silicone wristbands, Electronic cigarettes, Nicotine or derivatives, Special populations

SCHOLARONE™
 Manuscripts

1
2
3
4
5 Title:

6 Nicotine, cotinine and tobacco-specific nitrosamines (TSNAs) measured in children's silicone
7 wristbands in relation to secondhand smoke and e-cigarette vapor exposure.
8
9

10
11 Authors:

12 Quintana, Penelope JE, PhD, MPH*

13 Lopez-Galvez, Nicolas, MPH, MA

14 Dodder, Nathan G., PhD

15 Hoh, Eunha, PhD

16 Matt, Georg E., PhD

17 Zakarian, Joy M., MPH

18 Vyas, Mansi, MPH

19 Chu, Linda, BS

20 Akins, Brittany, BS

21 **Padilla, Samuel, BS**

22 Anderson, Kim A., PhD

23 Hovell, Melbourne F., PhD, MPH
24
25
26
27
28
29
30
31
3233 **corresponding author*
34
3536 **Corresponding author:**

37 Penelope JE (Jenny) Quintana, PhD, MPH

38 Professor and Graduate Advisor for MPH and MS

39 School of Public Health

40 San Diego State University

41 5500 Campanile Dr.

42 San Diego, CA 92182-4162

43 619-594-1688/ fax 619-594-6112

44 jquintan@sdsu.edu
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60

Abstract (250 words)

Introduction: Simple silicone wristbands hold promise for exposure assessment in children. We previously reported strong correlations between nicotine in wristbands worn by children and urinary cotinine. Here, we investigated differences in wristband chemical concentrations among children exposed to secondhand smoke from conventional cigarettes (CC) or secondhand vapor from electronic cigarettes (EC), and children living with non-users of either product (NS).

Methods: Children (n=53) wore three wristbands and a passive nicotine air sampler for 7 days and one wristband for 2 days, and gave a urine sample on day 7. Caregivers reported daily exposures during the 7-day period. We determined nicotine, cotinine, and tobacco-specific nitrosamines (TSNA) concentrations in wristbands, nicotine in air samplers, and urinary cotinine through isotope-dilution liquid chromatography with triple-quadrupole mass spectrometry (LC-MS/MS).

Results: Nicotine and cotinine levels in wristbands (WB) in children differentiated between groups of children recruited into NS, EC exposed, and CC exposed groups in a similar manner to urinary cotinine (UC). Wristband levels were significantly higher in the CC group (WB nicotine median 233.8 ng/g silicone, UC median 3.6 ng/ml, n=15) than the EC group (WB nicotine median: 28.9 ng/g, UC 0.5 ng/ml, n=19), and both CC and EC group levels were higher than the NS group (WB nicotine median: 3.7 ng/g, UC 0.1 ng/ml, n=19). TSNA, including the known carcinogen NNK, were detected in 39% of wristbands.

Conclusion: Silicone wristbands show promise for sensitive detection of exposure to tobacco-related contaminants from traditional and electronic cigarettes and have potential for tobacco control efforts.

IMPLICATIONS

Silicone wristbands worn by children can absorb nicotine, cotinine and tobacco-specific nitrosamines, and amounts of these compounds are closely related to the child's urinary cotinine. Levels of tobacco-specific compounds in the silicone wristbands can distinguish patterns of children's exposure to secondhand smoke and e-cigarette vapor. Silicone wristbands are simple to use and acceptable to children, and therefore may be useful for tobacco control activities such as parental awareness and behavior change, and effects of smoke-free policy implementation.

For Peer Review

INTRODUCTION

Simple techniques for measuring exposure to secondhand and thirdhand tobacco smoke (SHS, THS) and second and thirdhand vapor from electronic cigarettes (SHeV, THeV) can assist tobacco control efforts. Methods for assessing these exposures in children are especially needed

1. Current methods, while accurate, often rely on biological specimens such as urine and saliva, which can be difficult to obtain, and require special handling of specimens
2. Silicone wristbands, which are already worn by children by choice, have been shown to absorb toxic chemicals from the wearer's environment when worn for several days to weeks³⁻¹¹.

Our group was the first to apply silicone wristbands to measurement of tobacco product exposure in children, and we demonstrated that nicotine levels in silicone wristbands worn by children (n = 31) for 2 and 7 days were both highly correlated with cotinine, a metabolite of nicotine, in urine from the same child¹². Here, we further investigated the ability of silicone wristbands to record differences in nicotine exposures among children recruited into different exposure groups: children exposed to SHS or those exposed to SHeV compared to children living with non-users of either conventional or electronic cigarettes. We also compare silicone wristband performance to air monitors carried by the children as well as to urinary cotinine. We expand analysis of tobacco products measured in silicone wristbands to cotinine and tobacco-specific nitrosamines (TSNAs) a group of chemicals specific to tobacco that includes known carcinogens¹³.

METHODS

Recruitment

Approvals were obtained from the Human Research Protection Program at San Diego State University and informed consent and assent was obtained. Participants were recruited from a past home air quality study who had agreed to be re-contacted (n = 27), referrals from other study participants (n = 5), tabling, i.e. **research assistants** recruited at tables placed in public locations

1
2
3 such as shopping centers (n = 8) and advertisements on Craigslist (n = 1), Facebook (n = 11),
4 and Instagram (n = 1). Children were all nonusers of conventional cigarettes (CC) or electronic
5 cigarettes (EC). We recruited children who lived with at least one adult who smoked a minimum
6 of 7 CC/week inside the home (n = 19), children who lived with at least one adult who used EC at
7 least 4 days/week inside the home and used e-liquids with nicotine (n = 19), and children not
8 exposed to nicotine products, who lived with adult nonsmokers and nonusers of EC who had a
9 complete ban on smoking and EC use inside their home (n = 15). Participants in the CC and EC
10 groups were more likely to be classified as African-American/Black or Biracial/mixed race (82%)
11 than Latinos (33%, $p < 0.05$).
12
13
14
15
16
17
18
19
20
21
22

23 Sample Collection

24
25 All samples and interview data were collected during two visits to participants' homes between
26 March 2017 and December 2018.
27
28
29

30 *Face-to-face home interviews.* Parents/caregivers were interviewed by trained research
31 assistants to collect data on education level, family income, and child's age, gender,
32 race/ethnicity; household characteristics including number of residents and rooms, and years
33 living in the current home; child's exposure to CC and EC over the sampling week; actual
34 household residents' usage frequency and amount of CC and EC products over the sampling
35 week; the location (inside home, inside the car, or outside) where CC and EC were used when
36 child was present.
37
38
39
40
41
42
43
44

45 *Daily monitoring.* Caregivers were asked to report their daily use of CC/EC and the child's
46 exposure to CC/EC during brief (5-minute) daily telephone calls with a research assistant, and to
47 verify wearing of the air monitor and wristband through a daily texted photo.
48
49
50
51

52 *Urine samples.* Single spot urine samples were collected from child participants using procedures
53 from our previous studies^{12,14,15}. Caregivers were asked to have the child collect their sample on
54
55
56
57
58
59
60

1
2
3 the morning of the second home visit, but sometimes samples were collected at other times due
4 to child and caregiver schedules.
5
6

7
8 *Silicone wristbands.* Cleaned prepared bulk wristbands in Teflon bags were obtained from K.
9 Anderson.⁸ Wristbands were individually stored and transported in clear glass jars with
10 polytetrafluoroethylene (PTFE)-faced PE-lined caps, or 2.0 mil thick Teflon PFA bags (2-day
11 wristband). Each child participant received three wristbands at the initial home visit. Two were
12 placed on the **child's** arm at the home visit, to be worn for 7 days, and one was given to the
13 caregiver **for child to wear** for the last 2 days. A reminder was given by phone to caregivers on
14 the morning of day 5 to place wristband on child. The actual wearing time for the 7-day wristbands
15 was from 5.8 days to 8.7 days, with a median of 7.0 days and 2.4 days for the 2-day wristbands.
16
17 The research assistants also collected a wristband field blank for each participant, handling and
18 transporting it in the same way as the worn wristbands, except it was immediately replaced in the
19 container. Caregivers texted a picture of the child wearing the wristband and air monitor (below)
20 once a day to verify wearing. Participants were asked to wear wristbands at all times during the
21 study. Most (36/53) children received 'small' size wristbands (median weight 3.8 g), 8/53 received
22 'extra small' size wristbands (median weight 3.7 g) and 9/53 received 'large' size wristbands
23 (median weight 4.3 g). Nicotine was present in 8/36 of the analyzed field blanks, and the average
24 blank value was 1.14 ng nicotine/wristband. The majority (6/8) of the field blanks with detectable
25 nicotine levels were from CC group, with one from EC group and the lowest one (0.6 ng
26 nicotine/wristband) in NS group. We reported **results as** ng nicotine/ wristband in our previous
27 paper. Here we report nicotine, cotinine, and TSNAs in ng nicotine/g silicone in wristband as in
28 **other studies**^{6,16}. For 31 wristbands **used for nicotine analysis, the** actual wristband weight was
29 not recorded, so we used the median weight of wristbands in that size category. Both 2-day and
30 7-day wristband nicotine and cotinine levels adjusted for silicone wristband weight (ng/g silicone)
31 were highly correlated with levels expressed as ng/wristband (all $\rho > 0.99$, $p < 0.01$). Sample
32
33
34
35
36
37
38
39
40
41
42
43
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60

1
2
3 wristbands and field blanks were transported cooled in individual borosilicate amber glass vials,
4 with Thermoset lids lined with PTFE, and stored at -20°C until analysis.
5
6

7
8 *Air nicotine.* Air nicotine concentrations were measured using passive air monitors ¹⁷. Child
9 participants were asked to wear the passive diffusion monitor badge (protected by a stainless
10 steel mesh 'tea ball') pinned to their outer clothing or backpack strap over one week, and to wear
11 the badge at all times except when bathing, swimming, sleeping, or when it interfered with
12 activities such as engaging in vigorous sports. Many participants reported not wearing the
13 monitors for short times (for example, forgetting in car when at church). The times were not
14 adjusted for non-wearing of the badge. Deployment time was used to calculate minutes monitor
15 was exposed and ranged from 5.9 days to 12 days (median 7.0 days). An air monitor field blank
16 was collected for each participant. Of these, 10% were analyzed, and if the average blank level
17 was < 30% of the level in the sample, the average blank level was subtracted from the sample
18 level. If the average blank level was > 30% of the level in the sample, the sample was censored.
19
20 Results are reported as ng nicotine/m³ of air.
21
22
23
24
25
26
27
28
29
30
31
32

33 **Laboratory Analysis**

34
35 Detailed laboratory methods are available in online supporting materials.
36
37

38
39 *Wristband Nicotine.* The QuEChERS extraction procedure modified for nicotine analyses in dust
40 and surface wipes was utilized to extract nicotine from the silicone wristbands ¹⁸. Nicotine
41 quantification was conducted by LC-MS/MS (Agilent 1200 Series LC system coupled to an Agilent
42 6460 Triple Quadrupole system) operated in positive electrospray ionization (ESI+) mode. The
43 estimated method detection limit (MDL) ¹⁹ was 0.19 ng/wristband. Detailed information on the
44 sample preparation, extraction and quantification of nicotine from wristbands has been described
45
46
47
48
49
50

51 ¹².

1
2
3 *Wristband Cotinine.* Cotinine was extracted from the same wristband as nicotine using the
4 nicotine wristband extraction method, above, with a final spiked cotinine-d3 concentration of 5
5 ng/mL, and quantified with the same instrumental method as for urinary cotinine, below ^{14,15}. The
6 estimated MDL was 0.10 ng/wristband. We added this analyte to the second half of samples (n =
7 22) after considering literature suggesting that sweat could be a route of exposure for the
8 wristbands.
9
10
11
12
13
14
15

16 *Wristband TSNAs.* The extraction procedure is described in detail in the supporting materials. A
17 separate wristband was analyzed for TSNAs. One half of the wristband was weighed, then spiked
18 with the four internal standards (NNK-d4, NAB-d4, NAT-d4, NNN-d4). The final concentration of
19 each of the deuterated TSNAs was 12.5 ng/mL. The TSNAs were quantified by LC-MS/MS,
20 operated in positive electrospray ionization (ESI+) mode as in ¹⁵. The estimated MDL was 0.10
21 ng/wristband. A total of 51 wristbands were analyzed for TSNAs.
22
23
24
25
26
27
28

29 *Urinary Cotinine.* Urinary cotinine was determined through isotope dilution LC-MS/MS ^{12,14,15}. The
30 estimated MDL was 0.033 ng/mL urine, and cotinine levels were reported in ng/mL urine ²⁰.
31
32
33

34 *Air Nicotine.* The extraction procedure is described in detail in the supporting materials, and used
35 a procedure similar to nicotine in wristbands. The final concentration of nicotine-d₄ was 5 ng/mL.
36 The instrumental method and calibration procedure described for wristband nicotine was used for
37 quantification.¹² The estimated MDL was 0.13 ng/badge.
38
39
40
41
42

43 **Statistical Analysis**

44
45 Descriptive statistics were produced using SPSS v26. The Kruskal Wallis test followed by pairwise
46 Mann-Whitney U-test were utilized to determine differences of nicotine and cotinine
47 concentrations among participants' groups. Spearman rank-order correlations (*rho*) were used to
48 determine associations. The Type I error rate was set at 5% (two-tailed). Statistical tests were
49 conducted with SPSS v25 and 26.
50
51
52
53
54
55
56
57
58
59
60

RESULTS

Demographic characteristics

All children recruited for this study were nonsmokers and nonusers of EC. The majority of participants were multiracial (39.6%) or Latino/a (22.6%) (Table 1). The majority of children were females (60%), and a median 9 years of age (range 3 – 14). Participants in this study lived a median 3 years in their home with a median number of occupants of 5 (range 2 – 16). African American and multiracial participants had significantly higher urinary cotinine than Latino participants.

Detection levels for nicotine and cotinine in wristbands, cotinine in urine and air nicotine samplers

Nicotine was detected in 100% of all silicone wristbands, with a range over three orders of magnitude (from 2.5 ng of nicotine to over 3000 ng per wristband) (supporting online material Table S2) and cotinine in almost all wristbands (91% and 95%, 7-day and 2-day, respectively). Urinary cotinine had a similar detection frequency with only one non-detect, (<2%). In contrast, only 36% of air samples had detectable levels of nicotine (Table S1).

Differences in wristband, air, and urine measures between exposure group

Table 2 gives the medians by exposure group for the children for 7-day wristband nicotine, 7-day wristband cotinine, urinary cotinine and air nicotine. In addition to children's exposure groups based on recruitment criteria (Table 2, classification I), we present groups classified based on daily interviews during the 7 day period that the child was wearing the wristband, which differed in some cases from the report at recruitment: these groups are reported exposure of the child in the same indoor room (classification II), as well as product use by residents, whether or not the child was present (classification III).

1
2
3 Nicotine and cotinine levels in wristbands distinguished between CC, EC, and NS recruitment
4 groups (Table 2, **classification I**), as did urinary cotinine and air nicotine. Nicotine and cotinine
5 levels were significantly higher in 7-day wristbands (WB), as well as children's urinary cotinine
6 recruited in the CC group (WB nicotine median: 233.8 ng/g silicone; urinary cotinine: 3.6 ng/ml)
7 than the EC group (WB nicotine median: 28.9 ng/g; urinary cotinine: 0.5 ng/ml), and both CC and
8 EC group levels were higher than in the NS group (WB nicotine median: 3.7 ng/g; urinary cotinine:
9 0.1 ng/ml). Wristband cotinine concentrations also distinguished between the CC, EC and NS
10 recruitment groups (Table, 2, **classification I**). Table S3 (supporting online materials), presents
11 data for 2-day wristband nicotine and cotinine, with similar results.
12
13
14
15
16
17
18
19
20
21

22 Classifications II and III were based on caregiver report over the 7 days of the study. For all
23 measures, the **wristbands worn by children in the CC group were** significantly more contaminated
24 compared to EC and NS groups (Table 2, **classifications II and III**). Examining the residents' use
25 of CC use group further (detailed examination of III, Table 2), it is clear that the highest WB
26 nicotine? levels, as well as air and urinary cotinine levels, were measured when residents smoked
27 CC inside: WB nicotine for CC inside group median 428.4 ng/g silicone vs. CC outside only
28 median 36.7 ng/g, $p < 0.01$, Table 2. For WB nicotine, cotinine and urinary cotinine, the EC group
29 was also significantly higher than NS groups for classification III (Table 2). We further examined
30 the NS groups in II and III to examine wristband sensitivity to low levels of exposure. In
31 classification II, 24 of the children were classified as 'NS' by their caregivers, and of these, 14
32 were classified as NS in both II and III (Table 2). The 10 children (24 total – 14) from homes where
33 residents used CC or EC but the caregiver reported the child was not exposed (NS in classification
34 II only) had significantly higher, WB nicotine, WB cotinine and urinary cotinine than the 14 children
35 from homes with no EC/CC use and a total ban (classification as NS in both II and III). For WB
36 nicotine, **the** median was 46.5 ng/g silicone) vs. 3.9 ng/g, for WB cotinine 22.3 ng/g vs. 0.3 ng/g,
37 and for urinary cotinine 0.3 ng/ml (0.2-3.5) vs. 0.1 ng/ml (0.0-0.2), all $p < 0.01$.
38
39
40
41
42
43
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60

Correlation of wristband nicotine and cotinine with quantitative measures of tobacco-product use

To determine whether levels of nicotine and cotinine in silicone wristbands were sensitive to the levels of tobacco product use, we determined correlations with reported measures of indoor CC and EC use (Table 3). For both the child's exposure as reported by the caregiver (exposure classification II) as well as by the resident's reported use, classification III), the amount of nicotine (ng/g silicone) in the wristband was significantly correlated with the number of cigarettes smoked indoors over the 7 day period ($\rho = 0.706, 0.725$, respectively, $p < 0.01$). The amount of vaping reported was also significantly correlated with nicotine levels in wristbands, both for reported exposure to EC in minutes ($\rho = 0.442, 0.557$, respectively, $p < 0.01$) and in reported mLs of EC-fluid used per week ($\rho = 0.557, 0.581$, respectively, $p < 0.01$), even though the amount of nicotine in the product was unknown.

Measurement of tobacco-specific nitrosamines in silicone wristbands

We detected TSNA in 39% of silicone wristbands (online supporting material Table S1), mostly in children exposed to CC (Table 4). For recruitment group (I) as well as exposure (II) and use (III) group, the CC group had a higher level of total TSNA on the wristbands (median 0.25 ng/g silicone vs 0.05 ng/g silicone in III NS group, Table 4) mainly due to the well-studied lung carcinogen NNK (64% detection in product use CC group, 46% detection in CC plus EC group, 7% in EC group and 0% in NS group). Details for each individual TSNA (NNK, NAT, NAB, NNN) are given in the supporting online material (Table S4). NAB was the only TSNA to be detected in the NS groups I, II or III (14%, 9%, and 14% detection in NS groups, respectively, Table 4).

Correlations between analytes

The 2-day and the 7-day wristband nicotine levels were highly correlated ($\rho = 0.94$), with a median 28% difference, and both 2-day and 7-day wristband nicotine levels were highly correlated

1
2
3 with urinary cotinine on day 7 (both $\rho > 0.90$) (supporting online material Table S5). Cotinine in
4 wristbands was also highly correlated with wristband nicotine (both $\rho > 0.90$) and urinary
5 cotinine ($\rho = 0.87$). The median ratio of cotinine to nicotine measured in the same wristband
6
7 decreased in more highly exposed groups, though none of the decreases were statistically
8
9 significant e.g., from a median 0.74 in NS recruitment group vs. 0.50 in EC group and 0.20 in CC
10
11 group) (supporting online materials Table S6).
12
13
14

15 16 **DISCUSSION**

17
18
19 Silicone wristbands recorded nicotine and cotinine over a range spanning three orders of
20 magnitude, with 100% detection rate for nicotine. We demonstrated significant differences in
21 silicone wristband nicotine and cotinine between groups of children recruited into non-smoker,
22 EC exposed, and CC exposed groups in a manner similar to urinary cotinine, demonstrating the
23 validity of the silicone wristband sampler in measuring exposure to tobacco products. Silicone
24 wristbands had a sensitive detection limit for nicotine and cotinine: In children reported as non-
25 exposed by the caregiver, nicotine in wristbands more closely tracked caregiver's use patterns
26 regardless of children's presence, rather than child's exposure as reported by the caregiver, as
27 did urinary cotinine. This increase might be due to child exposures to drifting SHS or SHeV
28 unnoticed by the caregiver²¹, and/or potential exposure to thirdhand smoke residue²²⁻²⁴, and
29 demonstrates the sensitivity of the silicone wristband sampler. The highest levels for wristbands
30 were clearly associated with exposure to CC indoors, demonstrating the ability of the sampler to
31 detect high exposures in a similar matter to urinary cotinine. Within groups, we also detected
32 significant correlations with the number of CC or amount of EC used, indicating silicone
33 wristbands can detect an exposure-response relationship. Silicone wristbands performed in a
34 similar manner to urinary cotinine, and the cost of analysis is similar to air nicotine and cotinine,
35 but wristbands are simpler to deploy and collect. Although wristbands were deployed and
36
37
38
39
40
41
42
43
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60 retrieved in-person in this study, other studies have shipped wristbands internationally at ambient

1
2
3 temperatures^{4,25}, and semi-volatile organic compounds (SVOCs) concentrations were found to
4 be stable in wristbands at room temperatures (nicotine is an SVOC)²⁶. The next step is to validate
5 remote deployment and collection of wristbands for tobacco-related compounds collected under
6 standard mailing conditions to assist in community-based exposure studies.
7
8
9
10

11 We demonstrated that children exposed to secondhand EC vapor by their caregivers have higher
12 levels of nicotine on their wristbands than do children of non-smokers/users. E-cigarette use in
13 the home has been reported to result in measurable air nicotine (geomean 130 ng/m³ nicotine in
14 homes of indoor e-cigarette users compared to 20 ng/m³ in homes of non-smokers/non-users)²⁷
15 and 200 ng/m³ nicotine in homes of indoor e-cigarette users in another study²⁸. In our study, less
16 than 50% of the air monitors carried by children in the EC-only caregiver use registered above
17 our nicotine in air detection limit (median 0.0, 75th percentile 22.0 ng/m³, maximum 180.5), but the
18 air monitor in our study traveled with the child, as opposed to a static home measurement, so
19 these integrated air levels were likely lower. To our knowledge, the elevated level of nicotine in
20 wristbands is the first report of personal measurement of nicotine from secondhand e-cigarette
21 exposure in children.
22
23
24
25
26
27
28
29
30
31
32
33
34
35

36 In our data, the 7-day wearing time and the 2-day wearing time produced comparable results.
37 The 7-day measurement was higher in nicotine, but only by 28%, rather than the 350% expected
38 if the levels were linear over the time exposed. This may be due to saturation of the wristband or
39 degradation of nicotine over time, and this requires further study. The shorter time of wearing may
40 be preferable, as a few 7-day wristbands were accidentally lost.
41
42
43
44
45
46

47 The routes of exposure assessed by the silicone wristbands, whether inhalation, dermal, or
48 ingestion, or a combination, has been debated. Early deployments of the silicone wristbands
49 focused on the ability of the silicone to sample air exposures, which would presumably partition
50 into the wristband based on chemical characteristics²⁶, and PAHs in active air samplers
51 correlated with PAHs in the paired wristbands³. Some data have emerged implicating other routes
52
53
54
55
56
57
58
59
60

1
2
3 of exposure, such as dermal and ingestion exposures. Aerts et al.²⁹ detected pesticide residues
4 in wristbands not present in paired air samples, suggesting that silicone wristbands directly worn
5 on the skin may also capture ingested or dermal contaminants. Wang et al.³⁰ found that wristband
6 analytes were better correlated with dermal wipes plus air measurements than with air alone,
7 indicating that inhalation and dermal routes of exposure were measured by the silicone wristband
8 sampler. The route of exposure assessed by the wristband in children exposed to tobacco
9 products should be further investigated.
10
11
12
13
14
15
16
17

18 Nicotine and cotinine in the wristbands may arise from contact of the wristbands with sweat, as
19 both of these compounds are excreted in sweat, as well as related nicotine metabolites (e.g. OH-
20 cotinine)³¹⁻³³. Evidence that sweat may contribute to observed nicotine and cotinine on the
21 wristband is supported by the strong correlation ($\rho > 0.9$) between urinary cotinine and
22 wristband nicotine. This correlation is much stronger than reported from other silicone wristband
23 studies that compared urinary and wristband levels for other toxicants^{3,5}. Between 87 and 203 ng
24 of nicotine were collected on a commercial sweat patch worn for 72 hours by 5 adults exposed to
25 SHS³⁴. In our participant wristbands, a median 571 ng (range 130 - 2629 ng) of nicotine were
26 collected on wristbands worn for 2 days by children exposed to SHS. In the wristbands, the ratio
27 of cotinine to nicotine in the same wristband decreased (though non-significantly) in more highly
28 exposed subjects. We could not determine the ratio between nicotine and cotinine in sweat from
29 the studies cited for comparison, however. It is possible that sweat is a major contributor to
30 observed levels at lower doses, but air nicotine increasingly contributes to nicotine in wristbands
31 at higher air levels of nicotine.
32
33
34
35
36
37
38
39
40
41
42
43
44
45
46
47

48 We detected TSNAs in silicone wristbands worn for 7 days, mostly in children exposed to CC,
49 including the well-studied lung carcinogen NNK¹³. It is unknown whether a wearing period would
50 increase detection, or if a shorter wearing time would record similar levels of detection. Also, it is
51 possible that we would have detected metabolites of TSNAs excreted in sweat, as well as parent
52
53
54
55
56
57
58
59
60

1
2
3 TSNAs, for reasons discussed above. It is possible that TSNA exposure could arise through
4
5 thirdhand smoke residue in dust or on surfaces ²², as even in the NS group a TSNA (NAB) was
6
7 detected.
8
9

10 There are limitations to this study. One is the complex behavior around children's exposure to
11
12 tobacco-related products. Classification of exposure based on recruitment self-report did not
13
14 always match up with the classification based on smoking and vaping behavior during the 7-day
15
16 study period. For example, some in the 'exposed to EC indoors' recruitment group were only
17
18 exposed outdoors during the study period (n = 2) or exposed to CC outdoors as well (n = 5). Due
19
20 to budget limitations, we could not collect dust or wipe samples to assess the extent of thirdhand
21
22 smoke contamination in the child's home, which may be an unmeasured factor contributing to
23
24 wristband nicotine or TSNA levels¹⁵. Also, if we could have measured children's sweat directly,
25
26 we could compare levels and ratios of nicotine and cotinine in sweat with wristband levels.
27
28

29 **CONCLUSION**

30
31
32 The simple silicone wristband demonstrates a wide range over three orders of magnitude and
33
34 with 100% detection for wristband nicotine. Silicone wristband nicotine and cotinine levels can
35
36 discriminate between groups of children exposed to SHS (CC), SHeV (EC) and children not living
37
38 with a user or smoker, with sample sizes between 15 and 20 children for each group. The
39
40 wristbands detected low and high exposures within groups and discriminated between exposure
41
42 groups for children in a manner similar to urinary cotinine. Our data also indicate that children
43
44 living with e-cigarette users are significantly more exposed to nicotine from e-cigarettes than
45
46 children living with non-smokers.
47
48

49 We demonstrate that silicone wristbands can be used to detect multiple classes of chemicals
50
51 related to tobacco smoke (nicotine, cotinine, TSNAs). We detected carcinogenic TSNAs in
52
53 silicone wristbands worn for 7 days, mostly in children exposed to CC smokers. Significant
54
55 questions remain whether the silicone wristband captures pollutants on the skin or in the sweat
56
57
58
59
60

1
2
3 of wearer or pollutants in the air. The silicone wristband is a simple-to-deploy method for
4
5 assessing exposure to tobacco-related products that shows promise for tobacco control efforts.
6
7
8
9
10
11
12
13
14
15
16
17
18
19
20
21
22
23
24
25
26
27
28
29
30
31
32
33
34
35
36
37
38
39
40
41
42
43
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60

For Peer Review

Acknowledgements

The authors express their gratitude to Christine Batikian, MPH, Viridiana Mendoza, and Madeleine Warman for assisting with data collection.

This research was supported by funds from the California Tobacco Related Disease Research Grants Program Office of the University of California, Grant Number 25IP-0023 (P. Quintana, Principal Investigator).

For Peer Review

Conflict of Interest

KAA discloses a financial interest in MyExposome that is marketing products related to the research being reported. The terms of this arrangement have been reviewed and approved by Oregon State University in accordance with its policy on research conflict of interest. The other authors declare no conflict of interest.

For Peer Review

REFERENCES

1. Hubal EAC, Sheldon LS, Burke JM, et al. Children's exposure assessment: a review of factors influencing Children's exposure, and the data available to characterize and assess that exposure. *Environmental Health Perspectives*. 2000;108(6):475-486.
2. Avila-Tang E, Al-Delaimy WK, Ashley DL, et al. Assessing secondhand smoke using biological markers. *Tob Control*. 2013;22(3):164-171.
3. Dixon HM, Scott RP, Holmes D, et al. Silicone wristbands compared with traditional polycyclic aromatic hydrocarbon exposure assessment methods. *Anal Bioanal Chem*. 2018;410(13):3059-3071.
4. Donald CE, Scott RP, Blaustein KL, et al. Silicone wristbands detect individuals' pesticide exposures in West Africa. *Royal Society Open Science*. 2016;3(8).
5. Hammel SC, Hoffman K, Webster TF, Anderson KA, Stapleton HM. Measuring Personal Exposure to Organophosphate Flame Retardants Using Silicone Wristbands and Hand Wipes. *Environ Sci Technol*. 2016;50(8):4483-4491.
6. Hammel SC, Phillips AL, Hoffman K, Stapleton HM. Evaluating the Use of Silicone Wristbands To Measure Personal Exposure to Brominated Flame Retardants. *Environ Sci Technol*. 2018.
7. Harley KG, Parra KL, Camacho J, et al. Determinants of pesticide concentrations in silicone wristbands worn by Latina adolescent girls in a California farmworker community: The COSECHA youth participatory action study. *Sci Total Environ*. 2019;652:1022-1029.
8. O'Connell SG, Kind LD, Anderson KA. Silicone Wristbands as Personal Passive Samplers. *Environmental Science & Technology*. 2014;48(6):3327-3335.
9. Manzano CA, Dodder NG, Hoh E, Morales R. Patterns of Personal Exposure to Urban Pollutants Using Personal Passive Samplers and GC x GC/ToF-MS. *Environ Sci Technol*. 2019;53(2):614-624.
10. Lipscomb ST, McClelland MM, MacDonald M, Cardenas A, Anderson KA, Kile ML. Cross-sectional study of social behaviors in preschool children and exposure to flame retardants. *Environ Health*. 2017;16(1):23.
11. Paulik LB, Hobbie KA, Rohlman D, et al. Environmental and individual PAH exposures near rural natural gas extraction. *Environ Pollut*. 2018;241:397-405.
12. Quintana PJE, Hoh E, Dodder NG, et al. Nicotine levels in silicone wristband samplers worn by children exposed to secondhand smoke and electronic cigarette vapor are highly correlated with child's urinary cotinine. *J Expo Sci Environ Epidemiol*. 2019;29(6):733-741.
13. Hecht SS. Tobacco smoke carcinogens and lung cancer. *J Natl Cancer Inst*. 1999;91:1194-1210.
14. Matt GE, Quintana PJE, Hoh E, et al. A Casino goes smoke free: a longitudinal study of secondhand and thirdhand smoke pollution and exposure. *Tob Control*. 2018.
15. Matt GE, Quintana PJE, Zakarian JM, et al. When smokers quit: exposure to nicotine and carcinogens persists from thirdhand smoke pollution. *Tob Control*. 2016;26(5):548-556.
16. Reddam A, Tait G, Herkert N, Hammel SC, Stapleton HM, Volz DC. Longer commutes are associated with increased human exposure to tris(1,3-dichloro-2-propyl) phosphate. *Environ Int*. 2020;136:105499.
17. Hammond SK, Leaderer BP. A diffusion monitor to measure exposure to passive smoking. *Environmental Science & Technology*. 1987;21(5):494-497.
18. Paya P, Anastassiades M, Mack D, et al. Analysis of pesticide residues using the Quick Easy Cheap Effective Rugged and Safe (QuEChERS) pesticide multiresidue method in combination with gas and liquid chromatography and tandem mass spectrometric detection. *Anal Bioanal Chem*. 2007;389(6):1697-1714.

19. US Federal Government. Definition and procedure for the determination of the method detection limit. Rev. 1.11. In. Vol 40 CFR Appendix B to Part 136: US Federal Government,; 2011.
20. Matt GE, Wahlgren DR, Hovell MF, et al. Measuring environmental tobacco smoke exposure in infants and young children through urine cotinine and memory-based parental reports: empirical findings and discussion. *Tob Control*. 1999;8(3):282-289.
21. Gambino J, Moss A, Lowary M, et al. Tobacco Smoke Exposure Reduction Strategies—Do They Work? *Academic Pediatrics*. 2020.
22. Jacob P, 3rd, Benowitz NL, Destailats H, et al. Thirdhand Smoke: New Evidence, Challenges, and Future Directions. *Chem Res Toxicol*. 2017;30(1):270-294.
23. Matt GE, Quintana PJ, Hovell MF, et al. Households contaminated by environmental tobacco smoke: sources of infant exposures. *Tob Control*. 2004;13(1):29-37.
24. Matt GE, Quintana PJ, Zakarian JM, et al. When smokers move out and non-smokers move in: residential thirdhand smoke pollution and exposure. *Tob Control*. 2011;20(1):e1.
25. Bergmann AJ, North PE, Vasquez L, Bello H, Ruiz MDG, Anderson KA. Multi-class chemical exposure in rural Peru using silicone wristbands. *Journal of Exposure Science and Environmental Epidemiology*. 2017;27(6):560-568.
26. Anderson KA, Points GL, Donald CE, et al. Preparation and performance features of wristband samplers and considerations for chemical exposure assessment. *Journal of Exposure Science and Environmental Epidemiology*. 2017;27(6):551-559.
27. Ballbè M, Martínez-Sánchez JM, Sureda X, et al. Cigarettes vs. e-cigarettes: Passive exposure at home measured by means of airborne marker and biomarkers. *Environ Res*. 2014;135:76-80.
28. van Drooge BL, Marco E, Perez N, Grimalt JO. Influence of electronic cigarette vaping on the composition of indoor organic pollutants, particles, and exhaled breath of bystanders. *Environmental Science and Pollution Research*. 2019;26(5):4654-4666.
29. Aerts R, Joly L, Sztternfeld P, et al. Silicone Wristband Passive Samplers Yield Highly Individualized Pesticide Residue Exposure Profiles. *Environ Sci Technol*. 2018;52(1):298-307.
30. Wang S, Romanak KA, Stubbings WA, et al. Silicone wristbands integrate dermal and inhalation exposures to semi-volatile organic compounds (SVOCs). *Environ Int*. 2019;132:105104.
31. Kidwell DA, Holland JC, Athanaselis S. Testing for drugs of abuse in saliva and sweat. *Journal of Chromatography B: Biomedical Sciences and Applications*. 1998;713(1):111-135.
32. Concheiro M, Shakleya DM, Huestis MA. Simultaneous analysis of buprenorphine, methadone, cocaine, opiates and nicotine metabolites in sweat by liquid chromatography tandem mass spectrometry. *Anal Bioanal Chem*. 2011;400(1):69-78.
33. Koster RA, Alffenaar J-WC, Greijdanus B, VanDerNagel JEL, Uges DRA. Application of Sweat Patch Screening for 16 Drugs and Metabolites Using a Fast and Highly Selective LC-MS/MS Method. *Ther Drug Monit*. 2014;36(1).
34. Kintz P, Henrich A, Cirimele V, Ludes B. Nicotine monitoring in sweat with a sweat patch. *Journal of Chromatography B: Biomedical Sciences and Applications*. 1998;705(2):357-361.

Table 1. Demographic and household characteristics in relation to urinary cotinine (n = 53)

Demographics	n (%)	Urinary Cotinine (ng/ml) (median, p25-p75)	p-value
Gender			
Male	21 (40)	0.3 (0.1-2.6)	0.960
Female	32 (60)	0.5 (0.1-2.8)	
Ethnicity			
Latino/Hispanic	12 (23)	0.1 (0.1-2.7)	0.014
African American/Black	7(13)	2.8 (1.2-9.9)^a	
Caucasian/White	11 (21)	0.3 (0.2-0.0)	
Asian/ Pacific Islander	2 (4)	0.8 (0.5)	
Bi or Multiracial	21 (40)	1.3 (0.1-4.5)^a	
Yearly income			
\$20,000 and less	11 (21)	1.2 (0.1-4.9)	0.672
\$20,001 to \$50,000	19 (36)	0.9 (0.1-3.6)	
\$50,001 and above	23 (43)	0.3 (0.1-1.4)	
Parents/caregivers education			
High school and below	7 (13)	2.0 (0.3-6.7)	0.395
Some college	26 (49)	0.5 (0.1-3.5)	
Received higher education diploma	20 (38)	0.3 (0.1-1.4)	
Children's age groups			
3 to < 6 years of age	12 (23)	0.8 (0.2-2.6)	0.355
6 to < 11 years of age	28 (53)	0.8 (0.2-3.6)	
11 to 14 years of age	13 (25)	0.0 (0.0-3.1)	
Household characteristics			
Number of occupants			
<= 3	10 (19)	0.3 (0.2-6.2)	0.825
4 - 5	25 (47)	0.5 (0.1-1.7)	
>= 6	18 (34)	0.7 (0.2-2.9)	
Number of rooms			
<= 6	20 (38)	1.0 (0.2-3.5)	0.080
7	15 (28)	0.2 (0.0-1.3)	
>= 8	18 (34)	0.7 (0.3-3.0)	
Years living in residence			
<= 2	19 (37)	1.2 (0.3-4.7)	0.139
2 - <= 5	17 (33)	0.3 (0.1-1.5)	
>= 6	15 (29)	0.5 (0.1-1.9)	

p25-p75: 25th and 75th percentile. ^aAfrican-American/Black and Multiracial groups had significantly higher levels than Latinos.
Note: bolded values are significant (p< 0.05).

Table 2. Child's exposure group related to nicotine and cotinine concentrations in wristbands, nicotine in air and urinary cotinine.

Exposure group by classification scheme (#)	n*	7-day Wristband Nicotine Concentration (ng/g), n = 52 Median (p25–p75)	7-day Wristband Cotinine Concentration (ng/g), n = 22 Median (p25–p75), n	Air Nicotine Concentration (ng/m ³), n = 53 Median (p25–p75)	Urinary Cotinine Concentration (ng/ml), n = 53 Median (p25–p75)
Recruitment (I)					
NS-Non-exposed at recruitment	15	3.7 (1.6–4.6)	0.34 (0.0-1.0), 5	0.0 (0.0–0.0)	0.1 (0.0–0.1)
EC	19	28.9 (15.5–55.5)^a	7.4 (3.4-12.7)^a, 10	0.0 (0.0–14.3)	0.5 (0.3–1.2)^a
CC	19	233.8 (74.7–429.1)^{a,b}	36.8 (15.4-56.1)^{a,b}, 7	90.7 (0.0–291.3)^{a,b}	3.6 (1.4–9.9)^{a,b}
Child's reported exposure (II)					
NS-Non-exposed by caregiver report	24	4.5 (3.1–28.2)	1.5 (0.3-29.4), 11	0.0 (0.0–0.0)	0.1 (0.1–0.3)
EC	14	27.6 (15.1–58.0)^a	6.3 (4.0-10.9), 7	0.0 (0.0–41.4)	0.5 (0.3–1.2)^a
EC+CC	4	176.2 (55.7–561.3)^a	36.9 (– –) ^e , 1	171.5 (21.2–381.9)^{a,b}	2.4 (1.9–8.1)^a
CC	9	242.4 (105.9–470.8)^{a,b}	50.7 (– –), 3	210.2 (68.0–317.1)^{a,b}	4.2 (3.2–7.6)^{a,b}
Residents' tobacco product use (III)					
NS-No reported use by residents	14	3.9 (1.6–5.4)	0.3 (0.0-1.0) 5	0.0 (0.0–0.0)	0.1 (0.0–0.2)
EC	14	27.0 (14.9–53.1)^a	8.6 (4.9-13.4) ^a, 9	0.0 (0.0–0.0)	0.3 (0.2–0.6)^a
EC+CC	13	73.4 (28.4–214.9)^a	22.4 (4.8-49.4) ^a, 4	0.0 (0.0–55.2)	1.4 (0.7–2.8)^a
CC	12	243.4 (102.2–510.5)^{a,b}	43.8 (18.8-66.8) ^a, 4	187.1 (45.9–294.9)^{a,b}	4.2 (3.0–10.4)^{a,b}
Detailed residents' tobacco products use (III)					
NS-No reported use by residents	14	3.9 (1.6–5.4)	0.3 (0.0-1.0), 5	0.0 (0.0–0.0)	0.1 (0.0–0.2)
EC, inside or only outside home, no CC ^c	14	27.0 (14.9–53.1)^a	8.6 (5.0-13.4) ^a, 9	0.0 (0.0–0.0)	0.3 (0.2–0.6)^a
CC outside only, regardless of EC usage ^d	10	36.7 (23.1–91.2)^a	15.4 (– –) ^a , 3	0.0 (0.0–30.5)	1.1 (0.2–1.5)^a
CC and EC inside home	5	230.3 (65.4–311.6)^a	29.4 (– –), 1	47.0 (0.0–276.3)^a	2.8 (1.4–8.3)^a
CC inside home, no EC	10	428.4 (173.4–584.6)^{a,b}	43.8 (18.8-66.8) ^a, 4	279.5 (77.5–320.7)^{a,b}	4.8 (3.5–10.8)^{a,b}

NS: no exposure or no use of tobacco products, CC: conventional cigarette; EC: electronic cigarettes. p25-p75: 25th and 75th percentile Classification I – At recruitment, reported exposure of the child, NS, no EC/CC use by caregivers or residents and home smoking ban, EC exposure to EC inside, CC, exposure to CC inside. Classification II - Exposure of the child in same indoor room based on report of caregiver for 7 day period. Classification III -Residents reported use over 7 day period inside the home whether or not child was present. *sample size for all measures except wristband cotinine. ^a significantly higher levels than NS, (p<0.01); ^b significantly higher levels than EC, (p<0.01); ^c Group includes EC users only outside home (n=2) and EC users inside home (n= 12); ^d Group includes reported exposure to CC outside with no EC users (n= 2), CC outside with EC users outside (n=3), CC outside with EC users inside home (n= 5). ^e percentiles not reported for samples sizes of less than 4. Note: bolded values are significant (p< 0.05). Also note that the overall detection rate for air monitors was 36%; details are given in Table S1 in supporting online materials.

Table 3. Correlations of wristband, urine and air exposure measures in relation to conventional cigarettes (CC) and electronic cigarettes (EC) exposure and usage

Reported tobacco product exposure and use	Median (p25-p75)	7-Day Wristband Nicotine (ng/g silicone) <i>rho</i> (n)	7-Day Wristband Cotinine (ng/g silicone) <i>rho</i> (n)	Urinary cotinine (ng/ml) <i>rho</i> (n)	Air Nicotine (ng/m³) <i>rho</i> (n)
CC variables					
Child's exposure to CC inside (cigarettes/week)^a	0.0 (0.0-12.0)	0.706*** (38)	0.560* (15)	0.717*** (38)	0.723** (38)
Residents' use of CC inside (cigarettes/week)^b	0.9 (0.0-12.0)	0.725*** (29)	0.774** (10)	0.680*** (29)	0.779** (29)
EC variables					
Child's exposure to EC inside (minutes/week)^c	0.0 (0.0-14.8)	0.353* (38)	0.083 (18)	0.470** (38)	0.351* (38)
Residents' use of EC inside (minutes/week)^d	0.0 (0.0-17.8)	0.557** (26)	0.425 (13)	0.625*** (26)	0.455* (26)
Residents' use of EC inside (mL/week)^d	0.0 (0.0-3.3)	0.581** (26)	0.406 (13)	0.646*** (26)	0.500** (26)

CC: conventional cigarettes; EC: electronic cigarettes. p25-p75: 25th and 75th percentile. Spearman's correlations (*rho*): *(p<0.05), **(p<0.01), ***(p<0.001). ^a Exposure of the child in same indoor room based on report of caregiver for 7 day period, excluding exclusive EC use (a subset of Classification II from Table 2). ^b Residents reported use of CC over 7 day period inside the home whether or not child was present excluding exclusive EC use; (a subset of classification III from Table 2) ^c Exposure of the child in same indoor room based on report of caregiver for 7 day period, excluding any with CC usage (a subset of Classification II from Table 2).; ^d Reported caregiver use of EC over 7 day period inside the home whether or not child was present, excluding any with CC usage (a subset of classification III from Table 2. Note: bolded values are significant (p< 0.05).

Table 4. Tobacco-specific nitrosamines (TSNAs) in wristbands worn for 7 days in relation to child's exposure classification.

Exposure group by classification scheme (#)	Wristband Total TSNAs n	Wristband Total TSNAs n (%) detected*	Wristband Total TSNAs concentration, ng/g silicone (median, p25-p75)	NNK n (%) detected*	NAT n (%) detected*	NAB n (%) detected*	NNN n (%) detected*
Recruitment (I)							
NS- Non-exposed at recruitment	14	2 (14)	0.10 (0.0-0.10)	0 (0)	0 (0)	2 (14)	0 (0)
EC	19	5 (26)	0.10 (0.0-0.11)	3 (16)	0 (0)	2 (11)	1 (5)
CC	18	13 (72)	0.20 (0.05-0.53)^a	11 (61)	7 (39)	6 (33)	3 (17)
Child's reported exposure (II)							
NS -Non-exposed by caregiver report	23	6 (26)	0.05 (0.04-0.24)	3 (13)	1 (4)	3 (13)	1 (4)
EC	14	3 (21)	0.05 (0.04-0.19)	2 (14)	0 (0)	1 (7)	0 (0)
EC+CC	4	3 (75)	0.09 (0.04-0.53)	2 (50)	2 (50)	1 (25)	0 (0)
CC	9	8 (88)	0.40 (0.04-0.67)^a	7 (78)	4 (44)	5 (56)	3 (33)
Residents' tobacco product use (III)							
NS -No use by residents	13	2 (15)	0.05 (0.04-0.13)	0 (0)	0 (0)	2 (15)	0 (0)
EC	14	3 (21)	0.05 (0.04-0.21)	1 (7)	0 (0)	2 (14)	1 (7)
EC+CC	13	6 (46)	0.06 (0.04-0.53)	6 (46)	3 (23)	2 (15)	1 (8)
CC	11	9 (82)	0.25 (0.04-0.67)^{a,b}	7 (64)	4 (36)	4 (36)	2 (18)

Abbreviations: NS: no smoking and no e-cigarette use by caregivers or residents and home smoking ban, CC: Conventional Cigarette; EC: electronic cigarettes; TSNAs: Tobacco-specific nitrosamines; NAT: N'-nitrosoanatabine; NNN: N-nitrososnormicotine; NAB: N-nitrosoanabasine NNK: 4-(methylnitrosamino)-1-(3-pyridyl)-1-butanone ; p25-p75: 25th and 75th percentile. LOD: Limit of Detection (0.05 ng/g). Classification I – At recruitment, reported exposure of the child, NS, no smoking and no e-cigarette use by caregivers or residents and home smoking ban, EC exposure to EC inside, CC, exposure to CC inside. Classification II - Exposure of the child in same indoor room based on report of caregiver for 7 day period. Classification III -Residents reported use over 7 day period inside the home whether or not child was present. *n, (%) detected = wristbands that had listed compounds detected in laboratory analysis. ^a significantly higher levels than NS, (p<0.01); ^b significantly higher levels than EC, (p<0.01); Note: bolded values are significant different values (p< 0.05). Also note that one participant did not have a complete reported exposure variable, so n=50 instead of n=51 for this group.

Supplementary Material (online)

List of contents:

1.0 METHODS: Detailed Methods for Laboratory Analysis

2.0 RESULTS: Supplementary Tables

Table S1. Detection frequency and concentrations in wristbands, air and urine sample

Table S2. Detection frequency and concentrations in wristbands (ng/wristband) - not adjusted by wristband weight

Table S3. Child's exposure group related to nicotine and cotinine concentrations in wristbands worn for two days, nicotine in air and urinary cotinine

Table S4. Individual tobacco-specific nitrosamines (TSNAs) in wristbands worn for 7 days in relation to child's exposure classification

Table S5. Spearman correlations (*rho*) among analytes in wristband, urine and air samples

Table S6. Cotinine to nicotine ratio in wristbands, ng/g silicone, n=22

1.0 METHODS: Detailed Methods for Laboratory Analysis

Personnel performing extraction and analysis were blinded as to exposure status of participants. All sample containers and laboratory tools (scissors, tweezers, pipet tips, syringes, and syringe filters) were rinsed with solvent prior to use. Laboratory personnel wore disposable caps and laboratory coats when processing samples.

Materials. All solvents were LC/MS grade. Chemical standards of nicotine, nicotine-*d*₄, cotinine, and cotinine-*d*₃ were purchased from MilliporeSigma. Chemical standards of the tobacco-specific nitrosamines (TSNAs) 4-(methylnitrosamino)-1-(3-pyridyl)-1-butanone (NNK), N-nitrosoanatabine (NAT), and N-nitrosoanabasine (NAB) were purchased from Cambridge Isotope Laboratories. N-nitrosornicotine (NNN) was purchased from MilliporeSigma. The deuterated forms of the four TSNAs were purchased from Toronto Research Chemicals.

Wristband TSNA. Separate WB was used for TSNA. Each silicone wristband was cut into small pieces, and half were placed in a 50 mL centrifuge tube and spiked with 12.5 ng of each of the four internal standards (NNK-*d*₄, NAB-*d*₄, NAT-*d*₄, NNN-*d*₄). Five mL of acetonitrile was added, and the samples were vortexed for 30 min. The wristband pieces were removed, 300 mg of MgSO₄ was added, and the samples were vortexed for 3 min. Samples were centrifuged at 3000 rpm (900 × *g*) for 5 minutes. The organic layer was transferred to a new tube, evaporated to 1 mL, and added to a 2 mL vial containing the dSPE mixture (Agilent, QuEChERS Dispersive Kit, AOAC method, containing 50 mg primary-secondary amine, 50 mg C₁₈, and 150 mg MgSO₄). Samples were vortexed for 1 minute, then centrifuged at 10,000 rpm (5600 × *g*) for 1 minute. The liquid layer was removed and passed through the syringe filter. The final concentration of each of the deuterated TSNAs was 12.5 ng/mL. The instrumental method is described in ¹

Air Nicotine. Nicotine was extracted from the air badges by placing each badge in a 50 mL centrifuge tube, spiking with 10 ng of nicotine-*d*₄, and equilibrating for 15 min. Two mL of 0.1% formic acid was added and the samples were vortexed for 1 min. One mL 1M KOH was added and the samples were vortexed for 1 min. Two mL acetonitrile was added and the samples were vortexed for 30 min. The badge was removed, 2 g MgSO₄ and 0.5 g NaCl were added, and the samples vortexed for 1 min. Samples were then centrifuged at 3000 rpm (900 × *g*) for 5 min. One mL of the top (organic) layer was removed and passed through the syringe filter. The final concentration of nicotine-*d*₄ was 5 ng/mL. The instrumental method and calibration procedure described for wristband nicotine was used for quantification. The LOQ was 0.2 ng/badge, and the MDL was 0.13 ng/badge. Field blank values ranged from 0.7 - 4.4 ng nicotine (average 2.77), so the detection limit was more driven by field blank levels than LOQ of the method.

1. Matt GE, Quintana PJE, Zakarian JM, et al. When smokers quit: exposure to nicotine and carcinogens persists from thirdhand smoke pollution. *Tobacco control*. 2016;26(5):548-556.

2.0 RESULTS: Supplementary Tables

Table S1. Detection frequency and concentrations in wristbands, air and urine samples.

Analyte / measure	Detected n (%)	p25	p50	p75	p95
Urine Cotinine (ng/ml), n = 53	50 (94)	0.14	0.49	2.82	10.88
Air Nicotine (ng/m ³), n = 53	19 (36)	0.00	0.00	65.85	399.97
2-Day Wristband Nicotine (ng/g), n = 53	53 (100)	4.43	22.80	94.40	405.28
2 Day Wristband Cotinine (ng/g), n=22	21 (95)	0.52	3.98	9.05	39.72
7-Day Wristband Nicotine (ng/g), n = 52	52 (100)	5.44	31.63	107.71	562.37
7 Day Wristband Cotinine (ng/g), n=22	20 (91)	1.12	9.75	31.04	69.74
2 Day Wristband cotinine/nicotine ratio, (ng/g) n = 22	21 (95)	0.09	0.37	0.62	1.78
7 Day Wristband cotinine/nicotine ratio, (ng/g) n = 22	20 (91)	0.13	0.43	0.68	1.25
7 Day Wristband Total TSNAs (ng/g), n= 51	20 (39)	0.05	0.05	0.13	0.53
7 Day Wristband NAT, n =51	7 (14)	0.05	0.05	0.06	0.10
7 Day Wristband NAB, n=51	10 (20)	0.05	0.05	0.06	0.13
7 Day Wristband NNN, n = 51	4 (8)	0.05	0.05	0.05	0.07
7 Day Wristband NNK, n = 51	14 (27)	0.05	0.05	0.12	0.31

TSNAs: Tobacco-specific nitrosamines; NAT: N'-nitrosoanatabine; NNN: N-nitrosornicotine; NAB: N-nitrosoanabasine NNK: 4-(methylnitrosamino)-1-(3-pyridyl)-1-butanone; Note: only 22 (7-day) and 22 (2-day) wristbands were analyzed for cotinine. .p25 = 25th percentile, etc.

Table S2. Detection frequency and concentrations in wristbands (ng/wristband) - not adjusted by wristband weight.

Wristband measure	Detected n (%)	p25	p50	p75	p95
2-Day Wristband Nicotine (ng/WB), n = 53	53 (100)	16.30	85.96	362.25	1665.45
2-day Wristband Cotinine (ng/WB), n=22	21 (95)	1.95	15.21	34.20	151.44
7-Day Wristband Nicotine (ng/WB), n = 52	52 (100)	21.88	127.13	437.44	2162.48
7-day Wristband Cotinine (ng/WB), n=22	20 (91)	4.18	34.96	118.83	264.75

Table S3. Child's exposure group related to nicotine and cotinine concentrations in wristbands worn for two days, nicotine in air and urinary cotinine.

Exposure group by classification scheme (#)	n (%) [*]	2-day Wristband Nicotine Concentration (ng/g), n = 53 Median (p25–p75)	2-day Wristband Cotinine Concentration (ng/g), n = 22 Median (p25–p75), n	Urinary Cotinine Concentration (ng/ml), n = 53 Median (p25–p75)
Recruitment groups (I)				
NS-Non-exposed at recruitment	15 (28.3)	3.1 (1.3-4.4)	0.2 (0.1-0.9), 5	0.1 (0.0–0.1)
EC	19 (35.8)	24.8 (11.2-36.9)^a	4.0 (0.9–7.0)^a, 10	0.5 (0.3–1.2)^a
CC	19 (35.8)	149.9 (65.3-245.5)^{a,b}	14.6 (7.6–18.7)^{a,b}, 7	3.6 (1.4–9.9)^{a,b}
Child's reported exposure (II) (n=50)				
NS -Non-exposed by caregiver report	23 (46.0)	4.4 (1.9-25.5)	1.6 (0.2–7.6), 11	0.1 (0.1–0.3)
EC	14 (28.0)	14.9 (11.1-34.3)	3.6 (1.0–6.4), 7	0.5 (0.3–1.2)^a
EC+CC	4 (8.0)	136.8 (55.3-241.7)^a	16.2 (---) ^c , 1	2.4 (1.9–8.1)^a
CC	9 (18.0)	180.9 (109.1-391.1)^{a,b}	18.7 (---), 3	4.2 (3.2–7.6)^{a,b}
Residents' tobacco product use (III)				
NS -No reported use by residents	14 (26.4)	3.2 (1.2-4.4-26.3)	0.2 (0.1–0.9), 5	0.1 (0.0–0.2)
EC	14 (26.4)	17.4 (11.1-34.3)^a	4.0 (1.0-7.3)^a, 9	0.3 (0.2–0.6)^a
EC+CC	13 (24.5)	47.1 (13.9-132.6)^a	4.4 (1.2-10.8)^a, 4	1.4 (0.7–2.8)^a
CC	12 (22.6)	180.9 (81.2-242.9)^{a,b}	17.4 (14.9–37.2)^{a,b}, 4	4.2 (3.0–10.4)^{a,b}
Detailed residents' tobacco products use				
NS -No reported use by residents	14 (26.4)	3.2 (1.2-4.4)	0.2 (0.1–0.9), 5	0.1 (0.0–0.2)
EC, inside or only outside home, no CC ^c	14 (26.4)	17.4 (11.1-34.3)^a	4.4 (1.0–7.2)^a, 9	0.3 (0.2–0.6)^a
CC outside only, regardless of EC usage ^d	10 (18.9)	20.9 (11.5-48.6)^a	3.3 (0.5–11.9)^a, 3	1.1 (0.2–1.5)^a
CC and EC inside home	5 (9.4)	115.2 (49.2-286.6)^a	7.6 (---), 1	2.8 (1.4–8.3)^a
CC inside home, no EC	10 (18.9)	196.6 (137.3-334.9)^{a,b}	17.4 (14.9–37.2)^{a,b}, 4	4.8 (3.5–10.8)^{a,b}

NS: no exposure or no use of tobacco products, CC: conventional cigarette; EC: electronic cigarettes; Classification I – At recruitment, reported exposure of the child, NS, no smoking and no e-cigarette use by caregivers or residents and home smoking ban, EC exposure to EC inside, CC, exposure to CC inside. Classification II - Exposure of the child in same indoor room based on report of caregiver for 7-day period. Classification III -Residents reported use over 7-day period inside the home whether or not child was present. *n refers to all measures except wristband cotinine. ^a significantly higher levels than NS, (p<0.01); ^b significantly higher levels than EC, (p<0.01); ^c Group includes EC users only outside home (n=2) and EC users inside home (n= 12); ^d Group includes reported exposure to CC outside with no EC users (n= 2), CC outside with EC users outside (n=3), CC outside with EC users inside home (n= 5). ^epercentiles not reported for samples sizes of less than 4. Note: bolded values are significant (p< 0.05).

Table S4. Individual tobacco-specific nitrosamines (TSNAs) in wristbands worn for 7 days in relation to child's exposure classification.

Exposure group by classification scheme (#)	n (%)	NNK (ng/g)		NAT (ng/g)		NAB (ng/g)		NNN (ng/g)	
		Detected (%)	Median (min-max)	Detected (%)	(Median; min-max)	Detected (%)	(Median; min-max)	Detected (%)	(Median; min-max)
Recruitment groups (I)									
NS-Non-exposed at recruitment	14 (27.4)	0	< LOD	0	< LOD	15	LOD (LOD -0.12)	0	<LOD
EC	19 (37.3)	15	LOD (LOD-0.06)	0	< LOD	11	LOD (LOD -0.10)	5	LOD (LOD -0.06)
CC	18 (35.3)	61	0.15 (LOD-0.43)	39	0.06 (LOD-0.15)	33	LOD (LOD -0.13)	17	LOD (LOD -0.08)
Child's reported exposure (II)									
NS -Non-exposed by caregiver report	23 (46.0)	0	< LOD	0	< LOD	9	LOD (LOD -0.25)	0	<LOD
EC	14 (28.0)	14	LOD (LOD-0.19)	7	LOD (LOD -0.06)	7	LOD (LOD -0.10)	0	<LOD
EC+CC	4 (8.0)	25	0.09	50	0.06 (LOD -0.09)	25	0.05	0	<LOD
CC	9 (18.0)	77	0.25 (LOD-0.43)	44	0.06 (LOD -0.15)	33	LOD (0.04-0.14)	33	LOD (LOD -0.43)
Residents' tobacco product use (III)									
NS -No use by residents	14 (27.5)	0	< LOD	0	< LOD	14	LOD (LOD-0.13)	0	<LOD
EC	14 (27.5)	7	0.13	0	< LOD	14	LOD (LOD-0.10)	7	0.07
EC+CC	13 (25.5)	46	0.06 (LOD-0.31)	23	LOD (LOD-0.09)	25	LOD (LOD-0.14)	7	LOD (LOD-0.06)
CC	12 (23.5)	70	0.24 (LOD -0.43)	40	LOD (LOD-0.15)	40	LOD (LOD-0.14)	20	LOD (LOD-0.08)

NS: no exposure or no use of tobacco products, CC: conventional cigarette; EC: electronic cigarettes; Classification I – At recruitment, reported exposure of the child, NS, no smoking and no e-cigarette use by caregivers or residents and home smoking ban, EC exposure to EC inside, CC, exposure to CC inside. Classification II - Exposure of the child in same indoor room based on report of caregiver for 7-day period, Classification III -Residents reported use over 7-day period inside the home whether or not child was present. ^a significantly higher levels than NS, (p<0.01); ^b significantly higher levels than EC, (p<0.01); TSNAs: Tobacco-specific nitrosamines; NAT: N'-nitrosoanatabine; NNN: N-nitrosornicotine; NAB: N-nitrosoanabasine NNK: 4-(methylnitrosamino)-1-(3-pyridyl)-1-butanone ; LOD: Limit of Detection (approximately 0.05 ng/g, varies by wristband size).

Table S5. Spearman correlations (*rho*) among analytes in wristband, urine and air samples.

Environmental and Biomarker Samples	Urinary Cotinine (ng/ml), n = 53	Air Nicotine (ng/m ³), n = 53	2-Day Wristband Nicotine (ng/g), n=53	2 Day Wristband Cotinine (ng/g), n=22	7-Day Wristband Nicotine (ng/g), n = 52	7 Day Wristband Cotinine (ng/g), n=22	7 Day Wristband Total TSNAs (ng/g), n= 51
Urinary Cotinine (ng/ml), n = 53	-	0.72***	0.91***	0.87***	0.92***	0.84***	0.64***
Air Nicotine (ng/m ³), n = 53		-	0.66**	0.56**	0.71***	0.48*	0.46***
2-Day Wristband Nicotine (ng/g), n = 53			-	0.89***	0.94***	0.86***	0.60***
2 Day Wristband Cotinine (ng/g), n=22				-	0.91***	0.90***	0.48*
7-Day Wristband Nicotine (ng/g), n = 52					-	0.93***	0.59***
7 Day Wristband Cotinine (ng/g), n=22						-	0.44*
7 Day Wristband Total TSNAs (ng/g), n= 51							-

*(p<0.05) **(p<0.01) ***(p<0.001) TSNAs: Tobacco-specific nitrosamines, Note: bolded values are significant different values (p< 0.05). The sample size for correlations is the lower number of the pair.

Table S6. Cotinine to nicotine ratio in wristbands, ng/g silicone, n=22

Exposure group by classification scheme (#)	n (%)	7-day wristband	
		cotinine/ nicotine ratio (median, p25-p75)	p-value
Recruitment (I)			
NS-Non-exposed at recruitment	5 (19.0)	0.74 (0.3-1.11)	0.158
EC	10 (47.6)	0.50 (0.3-0.71)	
CC	7 (33.4)	0.20 (0.1-0.42)	
Child's reported exposure (II)			
NS-Non-exposed by caregiver report	11 (48.0)	0.60 (0.21-0.89)	0.391
EC	7 (33.3)	0.30 (0.11-0.58)	
EC+CC	1 (4.7)	0.092	
CC	3 (14.0)	0.20 (0.11-0.22)	
Residents' tobacco product use (III)			
NS-No reported use by residents	5 (19.0)	0.74 (0.0-1.27)	0.424
EC	9 (43.0)	0.45 (0.09-1.14)	
EC+CC	4 (19.0)	0.32 (0.13-0.61)	
CC	4 (19.0)	0.16 (0.09-0.66)	

Abbreviations: NS Child not exposed to tobacco smoke or electronic cigarette vapor, CC: conventional cigarette ; EC: electronic cigarettes;