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RESEARCH REPORT

ADDICTION

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Associations between nicotine vaping uptake and cigarette smoking cessation vary by smokers' plans to quit: longitudinal findings from the International Tobacco Control Four Country Smoking and Vaping Surveys

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Abstract

Background and Aims: Most population studies that evaluate the relationship between nicotine vaping and cigarette cessation focus on limited segments of the smoker population. We evaluated vaping uptake and smoking cessation considering differences in smokers' plans to quit.

Design: Longitudinal International Tobacco Control (ITC) Four Country Smoking and Vaping Surveys were conducted in 2016, 2018 and 2020.

Setting: This study was conducted in the United States, Canada, England and Australia.

Participants: Participants of this study were adult daily cigarette smokers who had not vaped in the past 6 months at baseline and had participated in two or more consecutive waves of the ITC Four Country Smoking and Vaping Surveys (n = 2815).

Measurements: Plans to quit cigarette smoking was assessed at baseline (within 6 months, beyond 6 months, not planning to quit) and at follow-up (within 6 months vs not within 6 months), cigarette smoking cessation was assessed at follow-up (smoking less than monthly [including complete cessation] vs daily/weekly/monthly smoking) and inter-wave vaping uptake was assesed between baseline and follow-up (none, only non-daily vaping and any daily vaping). Generalized estimating equations were used to evaluate whether inter-wave vaping uptake was associated with smoking cessation at follow-up and with planning to quit at follow-up, each stratified by plans to quit smoking at baseline.

Findings: Overall, 12.7% of smokers quit smoking. Smokers not initially planning to quit within 6 months experienced higher odds of smoking cessation when they took up daily vaping (32.4%) versus no vaping (6.8%; adjusted odds ratio [AOR], 8.58; 95% Cl, 5.06– 14.54). Among smokers planning to quit, smoking cessation rates were similar between those who did and did not take up daily vaping (25.1% vs 16.8%; AOR, 1.91; 95% Cl, 0.91–4.00), although we could not account for potential use of cessation aids. Daily vaping uptake was associated with planning to quit smoking at follow-up among those initially not planning to quit (AOR,6.32; 95% Cl, 4.17–9.59).

Conclusions: Uptake of nicotine vaping appears to be strongly associated with cigarette smoking cessation among smokers with no initial plans to quit smoking. Excluding

funding from the Society for the Study of Addiction. The sponsors had no role in the design and conduct of the study; collection, management, analysis, and interpretation of the data; preparation, review, or approval of the manuscript; and decision to submit for publication. smokers not planning to quit from studies on vaping and smoking cessation may underestimate potential benefit of daily vaping for daily smokers.

KEYWORDS

cessation, cigarette smoking, longitudinal, nicotine vaping, plans to quit, population

INTRODUCTION

Most population-based studies that have evaluated the relationship between use of nicotine vaping products (NVPs) and cigarette smoking cessation have focused on cigarette smokers who attempt to quit or who express interest in quitting smoking [1]. However, nicotine vaping may be related to longer-term progression toward smoking cessation at the population level (Figure 1). Studies that are limited to smokers who are already planning to quit, or who attempt to quit, exclude from consideration any positive or negative impact that nicotine vaping may have on earlier junctures of the smoking cessation process.

Indeed, using data from the Population Assessment of Tobacco and Health (PATH) Study, a longitudinal study in the United States (US), Kasza *et al.* [2] found that adult daily cigarette smokers who were not planning to ever quit smoking experienced nearly sixfold higher odds of planning to quit smoking in the future when they initiated daily vaping. Further, using PATH Study data, Kasza *et al.* [3] found that daily vaping uptake was associated with eightfold greater odds of smoking cessation among those who were not planning to ever quit. These findings are consistent with experimental studies that have reported that giving NVPs to unmotivated smokers is positively associated with change in quit intentions [4, 5] and reductions in cigarette smoking [6, 7]. However, no study has yet investigated whether the association between uptake of NVP use and cigarette smoking cessation differs as a function of smokers' initial plans to quit.

We hypothesized that the association between uptake of daily vaping and smoking cessation may be stronger among those who were initially not planning to quit than among those who were initially planning to quit, because there are more junctures in the cessation process that vaping can act on among the former group than among the latter group (Figure 1). That is, vaping may be positively associated with planning to quit, with making a quit attempt and/or with quitting among those who made a quit attempt. However, aside from the first juncture, where a person's guitting plans can be reported in the present, population studies that evaluate quit attempts or vaping effectiveness because a cigarette cessation aid (grey box in Figure 1) necessarily exclude from consideration all those who did not characterize themselves because having attempted to quit [8-10]. If recall of quit attempts systematically differs between those who did and did not use NVPs or other cessation aids, then efforts are needed to mitigate potential bias in analyses of making a quit attempt and analyses of cessation aid effectiveness when using population-based data [11-13]. Our approach was to determine whether current report of quit plans at baseline, which was not subject to recall, distinguishes a differential relationship between vaping uptake and smoking

cessation at the population level, extending findings recently reported from the US-only PATH Study [2, 3].

We used longitudinal data from the International Tobacco Control Four Country Smoking and Vaping Surveys (ITC 4CV) to evaluate the relationship between uptake of NVP use and smoking cessation among adult daily cigarette smokers in the United States, Canada, England and Australia, stratified by initial plans to quit smoking. We also evaluated the relationship between uptake of NVP use and change in plans to quit smoking, again stratified by initial plans to quit smoking.

METHODS

Population

Cohort data for this study come from the 2016-2020 ITC 4CV surveys (waves 1-3). The ITC 4CV includes four parallel online surveys of adult (ages 18+ years) cigarette smokers and recent quitters in the United States, Canada, England and Australia. Wave 1 data were collected from July 2016 to November 2016, wave 2 from February 2018 to July 2018 and wave 3 from February 2020 to June 2020. The retention rate for wave 2 was 45% [14], and the retention rate for wave 3 was 42% [15]. Detailed information on sample recruitment, retention, weighting, and so on can be found in the ITC 4CV technical reports [14, 15] and methods paper [16], and the full surveys and information on accessing the data are available online (https://itcproject.org).

We analysed data from daily cigarette smokers who had never vaped or had not vaped in the past 6 months at their baseline assessment, participated in the next follow-up assessment and had an interwave interval between baseline and follow-up of 18 to 24 months. These criteria of having not vaped in the past 6 months at baseline and having an inter-wave interval of 18 to 24 months were required to ascertain inter-wave vaping uptake that followed the baseline assessment of cigarette quit plans (further described in the Measures section below), which excluded 539 persons (please see Supporting information Figure S1), leaving a final sample of n = 2815 persons who contributed n = 3405 observations (i.e. 2815 persons provided one wave-pair observation and 590 persons provided two wave-pair observations). Compared with the 2815 persons included in analyses, the 539 persons excluded from analyses were more likely to be younger, male, heavier smokers and from Canada. Those included versus excluded did not differ on their plans to quit smoking, cessation rates, socioeconomic indicators or race/ethnicity (Supporting information Figure S1).

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FIGURE 1 Junctures at which nicotine vaping may relate to progression toward cigarette smoking cessation at the population level. NVP = nicotine vaping product

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Measures

Measures are described because being assessed at 'baseline' and/or at 'follow-up'. That is, we used three waves of data with wave 1 serving because baseline wave to wave 2 and with wave 2 serving as baseline wave to wave 3. such that the three waves were evaluated as two wave pairs. That is, a person could provide one wave-pair observation or could provide two wave-pair observations if the person was present in all three waves and met the baseline eligibility criteria at both wave 1 and wave 2. In the Statistical Analysis section below, we describe our analytic approach to account for multiple observations contributed by a person.

Cigarette quit plans at baseline

At each assessment, smokers were asked: 'Are you planning to quit smoking ...' with response options being within the next month; between 1 and 6 months from now; sometime in the future, beyond 6 months; not planning to quit; do not know. We combined response options, and we present results for those who were planning to quit within the next 6 months and for those who were not planning to quit within the next 6 months.

Descriptive characteristics at baseline

Respondents reported their sex (male, female), race/ethnicity (ethnic majority: White/English vs ethnic minority: Black/other minority), age (18-24 years, 25-39 years, 40-54 years, 55+ years), income (low, moderate, high, not reported, which incorporated country-specific differences in currency) and educational attainment (low, moderate, high, not reported, which incorporated country-specific differences in education systems), which were combined to indicate socioeconomic status (SES) as follows: low if both income and education were low, moderate if either income or education was low and high if neither income nor education were low (respondents who answered only one of the two items were included in the SES category called for by the answered item), country (United States, Canada, England and

Australia), cigarettes smoked per day (CPD) (1-10, 11-20, 21-30 and 31+), guitting self-efficacy using the item 'If you decided to give up smoking completely in the next 6 months, how sure are you that you would succeed? You do not need to be intending to guit to respond' (not at all sure/slightly sure, moderately sure, very sure/extremely sure, do not know) and desire to guit smoking using the item 'How much do you want to guit smoking?' (not at all, a little, somewhat, a lot, do not know).

Uptake of vaping between baseline and follow-up (inter-wave vaping uptake)

At each assessment, respondents were asked whether they ever vaped and if so, how often they currently vape (daily, less than daily but at least once a week, less than weekly but at least once a month, less than once a month but occasionally, not at all), whether they ever vaped daily and if so, how long ago they stopped vaping daily (less than 1 month ago, 1-3 months ago, 4-6 months ago, 7-12 months ago, 1-2 years ago, more than 2 years ago) and how long ago they last vaped (less than 1 week ago, 1-4 weeks ago, 1-3 months ago, 4-6 months ago, 7-12 months ago, 1-2 years ago, more than 2 years ago).

Among those who at baseline had not vaped within the past 6 months (that includes those who never vaped and those who last vaped more than six months before baseline), we derived a measure of inter-wave vaping uptake between baseline and follow-up as follows: (i) no vaping uptake (i.e. no vaping after baseline including at follow-up assessment), (ii) uptake of only non-daily vaping (i.e. any non-daily vaping after baseline including non-daily vaping at follow-up assessment; note that 'uptake of only non-daily vaping' includes those who vaped only once after baseline to make the qualitative distinction between no vaping and any vaping), and (iii) uptake of any daily vaping (i.e. any daily vaping after baseline including daily vaping at follow-up assessment). We also conducted a separate set of analyses in which we combined the 'uptake of only non-daily vaping' group with the 'uptake of any daily vaping' group to produce an 'uptake of any vaping' group, and we compared this combined group to the 'no vaping uptake' group.

Cigarette smoking cessation at follow-up (outcome)

We defined cigarette smoking cessation at follow-up as smoking cigarettes less than monthly, including complete cessation, versus daily/ weekly/monthly smoking at follow-up. We also conducted a set of sensitively analyses in which we coded less than monthly cigarette smokers as not having achieved cigarette smoking cessation.

Planning to quit cigarette smoking at follow-up (outcome)

The same item described above for cigarette quit plans at baseline was also used to assess whether plans to quit had changed at followup. We defined planning to quit cigarette smoking at follow-up as planning to quit in the next 6 months versus not planning to quit in the next 6 months. Those who responded 'do not know' regarding their plans to quit at follow-up were categorized as not planning to quit in the next 6 months. Those who had already quit at follow-up and were not asked about their plans to quit were included in the sample as 'planning to quit', and in a separate analysis, they were excluded from the sample. We did not impute missing data for any variables.

Statistical analysis

First, we assessed descriptive characteristics of our sample of daily cigarette smokers, stratified by cigarette quit plans at baseline. We compared groups using χ^2 tests. Next, we evaluated prevalence of vaping uptake between baseline and follow-up, stratified by cigarette quit plans at baseline. We evaluated prevalence of cigarette cessation at follow-up as a function of vaping uptake between baseline and follow-up, stratified by cigarette quit plans at baseline. We evaluated prevalence of cigarette cessation at follow-up as a function of vaping uptake between baseline. Last, we evaluated prevalence of planning to quit in the next 6 months at follow-up as a function of vaping uptake between baseline and follow-up, stratified by cigarette quit plans at baseline. Unweighted sample sizes are presented alongside weighted estimates.

We used generalized estimating equation (GEE) logistic regression analyses to evaluate the association between vaping uptake and smoking cessation (and separately, between vaping uptake and planning to quit in the next 6 months) using both assessment pairs (i.e. 2016–2018 and 2018–2020), stratified by baseline cigarette quit plans, and we evaluated the interactions between baseline cigarette quit plans and vaping uptake. We tested for statistical rather than 'biologic' interaction (see Knol *et al.*) [17] because our interest was in determining whether the estimate from the combination of the cigarette quit plans term and the vaping uptake term yielded a departure from the underlying form of our statistical model. We additionally attempted to evaluate whether these interactions varied by country, but many models failed to converge; some were not testable because of small cell sizes, and those that remained did not yield any significant country interactions for the main analyses and therefore, are not ADDICTION

reported here. GEE allows for the assessment of change between baseline and follow-up from both assessment pairs in a single analysis while statistically controlling for interdependence among observations contributed by the same individuals, increasing statistical power [18, 19]. We specified the unstructured covariance and within-person correlation matrices and the binomial distribution of the dependent variables using the logit link function.

All GEE analyses were adjusted for country, sex, race/ethnicity, age group, SES, CPD, time in sample (i.e. the number of waves the respondent had completed) and assessment pair. We decided a priori to adjust for these covariates and to categorize continuous variables such as age and CPD, consistent with prior ITC analyses and reporting of these descriptive data. All covariates were assessed at baseline of each assessment pair such that time-varying covariates varied by time. Those with missing data on covariates were excluded from analyses (n = 52). All analyses were weighted using longitudinal weights that were rescaled for country and for cohort so that the weighted results presented here represent the population of cigarette smokers in each country. All analyses were conducted using STATA V16 software (StataCorp LP). Analysis plans were not preregistered and results should be considered exploratory. This report follows the STROBE reporting guideline for cohort studies (https://www.equator-network. org/reporting-guidelines/strobe/).

RESULTS

Descriptive characteristics of adult daily cigarette smokers, stratified by cigarette quit plans at baseline

Among daily cigarette smokers who had not vaped in the past 6 months, 28% were planning to quit within the next 6 months, 34% were planning to quit sometime in the future beyond 6 months, 26% were not planning to quit at all and 13% did not know whether they planned to quit. As shown in Table 1, males, those age \geq 55 years, those with lower SES (including each SES component—income and education) and those smoking more cigarettes per day were overrepresented among those who were not planning to quit. Those with higher quitting self-efficacy and greater desire to quit were overrepresented among those who were planning to quit in the next 6 months.

Vaping uptake between baseline and follow-up as a function of cigarette quit plans at baseline

Overall, 13.5% (95% CI, 11.8–15.4) of daily cigarette smokers took up daily vaping, with daily vaping uptake highest among those who at baseline were planning to quit within the next 6 months (16.3% uptake; 95% CI, 13.0–20.3) and lowest among those who at baseline were not planning to quit at all (10.3% uptake; 95% CI, 7.4–14.2) or who did not know whether they planned to quit (10.8% uptake; 95% CI, 7.6–15.1) (Table 2).

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313-422 338 $290-389$ 278 $22-343$ 261 $192-345$ \mathbf{V} 6 $267-348$ 333 $295-373$ 272 $230-319$ 296 $238-362$ \mathbf{V} 7 $255-327$ 205 $272-340$ 418 $371-46.6$ 41.5 $534-479$ 4.5 0001 7 $244-330$ 31.5 $202-281$ 328 $272-36.6$ 322 $221-337$ 4.5 0001 7 $244-330$ 31.5 $202-281$ 328 $214-350$ 31.7 $272-36.6$ 322 $221-39.0$ 4.5 0001 7 $428-527$ 404 $361-448$ 31.3 $225-36.7$ 30.7 4.9 0001 935-49.4 453 31.7 $272-36.6$ 31.7 4.9 60001 935-49.4 453 401 31.3 $245-36.7$ 307 $249-31.8$ 6001 935-49.4 104 $01-12$ 01 $112-16.5$ 103 $11-26.8$ 5.7 60001	6 31.3 - 422 338 290 - 389 27.8 22.2 - 343 26.1 19.2 - 34.5 28. 6 26.7 - 348 333 29.5 - 373 27.2 230 - 319 29.6 238 - 36.2 238 - 36.2 238 - 36.2 238 - 36.2 25 330 31.5 25 330 31.5 22.2 - 330 31.5 22.1 - 330 31.5 22.1 - 330 31.5 22.1 - 330 31.5 22.1 - 330 31.5 20.011 32.8 - 30.6 31.7 27.2 - 36.6 32.2 26.1 - 390 31.7 27.2 - 36.1 - 390 31.3 26.5 - 36.7 30.7 - 30.6 31.3 26.5 - 36.7 30.7 - 30.6 31.3 26.5 - 36.7 30.7 - 30.6 31.3 26.5 - 36.7 30.7 - 30.6 31.3 26.5 - 36.7 30.1 - 30.7 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6 30.6	Ч	ς.	2.6-7.0	2.4	1.3-4.5	3.1	1.3-7.4	2.7	0.9-8.1	2.4	0.015
26.7-348 333 $295-37.3$ 27.2 $230-319$ 296 $238-362$ $251-323$ 305 $272-340$ 418 $371-466$ 415 $354-479$ 45 60001 $155-227$ 240 $202-2811$ 328 $278-366$ 322 $261-390$ 45 60001 $244-330$ 315 $274-359$ 317 $272-366$ 322 $261-390$ 45 60001 $244-330$ 315 $274-359$ 317 $272-366$ 322 $261-390$ 45 60001 $749-320$ 315 $27-65$ 317 $272-366$ 322 $261-300$ 491 $492-321-320$ 401 $402-341$ 49 60001 $395-494$ 453 $403-500$ 401 $349-456$ 4011 $345-411$ 49 60001 $30-81/2$ $37/2$ $403-500$ 401 $349-456$ 4111 $345-411$ 49 60001 $992-45/2$	6 $26.7-348$ 333 $295-37.3$ 272 $230-319$ 29.6 $238-36.2$ 6 $251-323$ 305 $272-340$ 418 $371-46.6$ 415 $354-479$ 45 6001 7 $247-330$ 315 $272-340$ 418 $371-46.6$ 415 $354-479$ 45 6001 7 $248-330$ 315 $207-281$ 328 $217-36.6$ 322 $261-39.0$ 45 400 416 $361-486$ 415 60001 7 $428-527$ 404 $361-448$ 313 $265-36.7$ 307 $249-372$ 400 4001 401 $41-480$ 401 $401-490$ 401 $401-490$ 401 $401-490$ 401 $401-490$ 4001 4001 $401-490$ 4001 $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$ $401-420$	e	6.6	31.3-42.2	33.8	29.0-38.9	27.8	22.2-34.3	26.1	19.2-34.5		
5 $251-323$ 30.5 $272-340$ 41.8 $371-46.6$ 41.5 $35.4-7.9$ 6 $155-227$ 240 $202-281$ 32.8 $278-381$ 275 $221-337$ 45 0001 7 $244-330$ 31.5 $274-359$ 31.7 $272-36.6$ 32.2 $261-39.0$ 45 $20-1-39.0$ 45 00001 31.3 $255-36.7$ 30.7 $249-37.2$ $261-39.0$ 41.6 $30.61.7$ $229-50.0$ 96 $52-17.0$ 4001 $30-81$ $402-54.1$ 402 4001 $402-54.1$ $402-54.1$ $402-54.1$ $402-54.1$ $402-54.1$ $402-54.1$ $402-54.1$ $402-54.1$ $402-54.1$ $402-54.1$ $402-64.1$ $402-64.1$ $402-64.1$ $402-64.1$ $402-64.1$ 400001 7 $30-80.1$ 45.3 4001 $349-45.6$ 4011 $345-48.1$ 40001 $402-54.1$ $402-54.1$ 40001 7 $200-60.6$ $401-1$ $347-45.6$		õ	0.6	26.7-34.8	33.3	29.5-37.3	27.2	23.0-31.9	29.6	23.8-36.2		
155-227 240 202-281 328 $278-381$ 275 $221-337$ 45 <0001 7 $244-330$ 315 $274-359$ 317 $272-366$ 322 $261-390$ 45 $20-372$ 307 $249-372$ 40.4 $361-448$ 313 $265-367$ 307 $249-372$ 40.972 $249-372$ 40.972 $249-372$ $492-527170$ 492 $292-1700$ 96 $52-1700$ 96 $52-1700$ 7 $30-81$ 422 $27-65$ 422 $29-60$ 96 $52-1700$ 96 $92-1700$ 7 $30-81$ 453 $400-496$ 401 $345-481$ 49 60001 7 $114-207$ 115 $110-165$ 103 $73-481$ 49 60001 7 $100-0.4$ 0.1 $117-182$ 115 $110-165$ 103 $53-481$ 49 6001 7 $000-04$ 0.4 $011-122$ 011	8 155-227 240 202-281 328 278-381 275 221-337 45 <0001 7 428-527 404 361-448 313 265-365 322 261-390 45 <0001	5	3.5	25.1-32.3	30.5	27.2-34.0	41.8	37.1-46.6	41.5	35.4-47.9		
244-330 31.5 $274-359$ 31.7 $272-366$ 32.2 $261-390$ $428-527$ 40.4 $36.1-448$ 31.3 $265-367$ 30.7 $249-372$ $30-81$ 4.2 $27-65$ 4.2 $29-60$ 9.6 $52-170$ $30-81$ 4.2 $27-65$ 4.2 $29-60$ 9.6 $52-170$ $30-81$ 4.2 $29-60$ 9.6 $52-170$ 4.9 60001 $395-49.4$ 45.3 $408-49.8$ 40.1 $349-45.6$ 41.1 $345-48.1$ 4.9 60001 $395-49.4$ 45.3 $144-20.7$ 13.5 $110-16.5$ 10.3 $34.4.3$ 40.1 $349-45.6$ 41.1 4.9 60001 $700-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ $15.1-26.8$ 5.7 60001 $700-0.04$ 0.7 $0.11-1.22$ 0.9 $0.4-1.9$ $15.1-26.8$ 5.7 60001 $700-0.04$ $0.$	5 $244-330$ 31.5 $274-359$ 31.7 $272-36.6$ 32.2 $261-39.0$ 7 $428-52.7$ 40.4 $361-44.8$ 31.3 $265-36.7$ 30.7 $249-37.2$ 9 $285-37.6$ 37.0 $228-41.4$ 45.6 $40.3-50.9$ 47.1 $40.2-54.1$ 49 <0001 4 $395-49.4$ 45.3 $408-49.8$ 40.1 $349-45.6$ 41.1 $345-48.1$ 49 <0001 7 $188-272$ 17.3 $144-20.7$ 13.5 $110-16.5$ 10.3 $73-43.1$ 4.9 <0001 7 $08-04.7$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ $15.$ $66-4.2$ 5060.6 5.7 <0001 8 $9.2-15.2$ 14.7 $11.7-18.2$ 20.1 $16.3-24.6$ 35.7 60.001 9 $00-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 5.7 <0001 9 $2.2.52$ 35.5 36.7 30.7 $36.7-20.6$ $36.7-20.6$ 96.42 2	18	8.	15.5-22.7	24.0	20.2-28.1	32.8	27.8-38.1	27.5	22.1-33.7	4.5	<0.001
7 $428-527$ 404 $361-448$ 31.3 $26.5-36.7$ 30.7 $249-37.2$ $30-81$ 4.2 $2.7-65$ 4.2 $2.7-6.6$ 9.6 $5.2-17.0$ $30-81$ 4.2 $2.7-6.5$ 4.2 $2.7-6.5$ 4.2 $2.9-6.0$ 9.6 $5.2-17.0$ $395-49.4$ 45.3 $30-81.4$ 45.6 $40.3-50.9$ 47.1 $402-54.1$ 4.9 $60-7.4$ $395-49.4$ 45.3 $40.8-49.8$ 40.1 $345-48.1$ 49.2 60.01 $395-49.4$ 45.3 $40.1-9.07$ 13.9 $110-16.5$ 10.3 $7.3-14.3$ 60001 $00-0.4$ 0.4 $0.4-1.9$ 11.5 $11.6-16.5$ 10.3 $7.3-14.3$ 60001 $00-0.4$ 0.4 $0.4-1.9$ 11.5 $0.6-4.2$ 5.7 60001 $1020-312$ $314-39.9$ 37.4 32.2 36.1 49.7 $600-0.7$ $000-0.4$ 0.0 0.1	7 $42.8-52.7$ 40.4 $36.1-44.8$ 31.3 $26.5-36.7$ 30.7 $24.9-37.2$ 9 $28.5-37.6$ 37.0 $22.7-6.5$ 4.2 $2.7-6.5$ 4.2 $2.7-6.7$ 4.9 60.001 4 $39.5-49.4$ 45.3 $40.8-49.8$ 40.1 $34.5-48.1$ 4.9 <0001 7 $18.8-27.2$ 17.3 $14.4-20.7$ 13.5 $11.0-16.5$ 10.3 $7.3-48.1$ 4.9 <0001 7 $18.8-27.2$ 14.7 $11.7-18.2$ 0.9 $0.4-1.9$ 1.5 $06.4.2$ <0001 8 $9.2-15.2$ 14.7 $11.7-18.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ <0001 7 $000-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ <0001 7 $508-60.6$ 49.8 42.4 $37.2-47.9$ 43.5 36.7 <0001 7 $508-60.6$ 49.8 42.4 $37.2-47.9$ 43.5 36.7 <0001 7 $000-0.$	28.	Ŋ	24.4-33.0	31.5	27.4-35.9	31.7	27.2-36.6	32.2	26.1-39.0		
30-81 4.2 $2.7-6.5$ 4.2 $2.9-6.0$ 9.6 $5.2-170$ $285-37.6$ 370 $328-41.4$ 45.6 $403-50.9$ 471 $40.2-54.1$ 4.9 <0001 $100-0.4$ 15.3 $14.4-20.7$ 13.5 $110-16.5$ 10.3 $7.3-14.3$ <0001 $118.8-272$ 17.3 $14.4-20.7$ 13.5 10.3 $7.3-14.3$ <0001 $00-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ $15.1-26.8$ 5.7 <0001 $100-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ $15.3-24.6$ $36.7-50.6$ 5.7 <0001 $280-372$ 35.5 $314-39.9$ 42.4 $37.2-47.6$ 36.7 $36.7-42.2$ $5.7 <0001 280-372 35.5 314-39.9 42.4 37.2-47.6 36.7 36.7-50.6 5.7 50001 174-24.9 20.7 20.3 35.7-32.6 36.7-32.6 36.7-42.7 50.6$	30-81 4.2 $2.7-6.5$ 4.2 $2.7-6.5$ 4.2 $2.7-6.7$ 9.6 $5.2-17.0$ 4 $39.5-49.4$ 45.3 $408-49.8$ 40.1 $349-45.6$ 41.1 $34.5-48.1$ 4.9 <0001 4 $39.5-49.4$ 45.3 $408-49.8$ 40.1 $349-45.6$ 41.1 $34.5-48.1$ 4.9 <0001 7 $188-27.2$ 17.3 $144-20.7$ 135 $110-16.5$ 10.3 $7.3-4.8.1$ 4.9 <0001 8 $9.2-15.2$ 14.7 $0.1-1.2$ 0.9 $0.4-1.9$ 15.7 <0.001 9 $9.2-15.2$ 14.7 $0.1-1.2$ 0.9 $0.4-1.9$ 15.7 <0.001 9 $9.2-15.2$ 14.7 $0.11.2$ 0.9 $0.4-2.7$ $0.00-0.4$ $0.0-0.7$ $0.0-0.7$ 0.0 $0.0-0.7$ $0.00-0.7$ $0.00-0.7$ $0.00-0.7$ $0.00-0.7$ $0.00-0.7$ $0.00-0.7$ $0.00-0.7$ $0.00-0.7$ $0.00-0.7$ $0.00-0.7$ $0.0-0.7$ $0.0-0.7$ $0.00-0.7$ $0.0-0.7$	47.		42.8-52.7	40.4	36.1-44.8	31.3	26.5-36.7	30.7	24.9-37.2		
285-37.6 37.0 $328-41.4$ 45.6 $403-50.9$ 47.1 $402-54.1$ 4.9 <0001 $395-49.4$ 45.3 $408-49.8$ 40.1 $349-45.6$ 41.1 $345-48.1$ 4.9 <0001 $188-27.2$ 17.3 $144-207$ 13.5 $110-16.5$ 10.3 $7.3-14.3$ $<000-0.4$ 0.1 $11.7-18.2$ 0.9 $0.4-1.9$ $15.1-26.8$ 5.7 <0001 $00-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ 0.001 $00-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ 0.001 $280-37.2$ 35.5 $314-39.9$ 42.4 $372-47.9$ 43.5 $36.7-50.6$ 5.7 0.001 $00-0.4$ 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ $0.00-0.7$ 0.0 $0.00-0.7$ $0.00-0.7$ 0.01 $0.00-0.7$ $0.00-0.7$ 0.01 $0.00-0.7$ 0.01 $0.00-0.7$ 0.01 $0.00-0.7$ $0.000-0.7$ $0.000-0.7$	285-376 37.0 $32.8-41.4$ 45.6 $403-50.9$ 47.1 $40.2-54.1$ 4.9 <0001 $395-49.4$ 45.3 $408-49.8$ 401 $34.9-45.6$ 41.1 $345-481$ 4.9 <0001 $188-272$ 17.3 $144-20.7$ 13.5 $11.0-16.5$ 10.3 $7.3-14.3$ <0001 $0.0-0.4$ 0.4 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ <0001 $9.2-15.2$ 14.7 $11.7-18.2$ 0.2 0.1 1.5 $0.6-4.2$ <0001 $9.2-15.2$ $314-39.9$ 42.4 $37.2-47.9$ 35.7 $36.7-50.6$ $5.7 <0001 280-30.6 49.8 45.3-54.3 37.4 32.2-42.6 36.7-50.6 5.7 <0001 0.0-0.4 0.0 0.1 0.1-0.5 0.1 0.0-0.7 0.001 0.0 0.0 0.0 0.0 0.0 0.0 0.0 0.0 0.0 0.0 0.0 0.0 0.0 0.0 0$	4.9		3.0-8.1	4.2	2.7-6.5	4.2	2.9-6.0	9.6	5.2-17.0		
395-49,4 45.3 $408-49,8$ 401 $349-45,6$ 411 $345-48,1$ $188-272$ 17.3 $144-20.7$ 13.5 $110-16.5$ 10.3 $7.3-48,1$ $188-272$ 17.3 $144-20.7$ 13.5 $110-16.5$ 10.3 $7.3-48,1$ $0.0-0.4$ 0.4 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $92-15.2$ 14.7 $11.7-18.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $92.0-37.2$ 35.5 $314-39.9$ 42.4 $37.2-47.9$ 43.5 $3.57-6.6$ 5.7 60001 $280-30.6$ 49.8 $45.3-54.3$ 37.4 $37.2-47.9$ $33.5-50.6$ 5.7 60001 $200-0.4$ 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ $0.0-0.7$ $0.0-0.7$ 0.7427 0.0 0.0 0.1 $0.1-0.5$ 0.1 $0.0-0.7$ 0.001 $0.0-0.7$ 0.0 $0.10.$	395-49,4 45.3 $408-49,8$ 401 $34,9-45,6$ 411 $34,5-481$ $188-272$ 17.3 $14,4-20.7$ 13.5 $110-16.5$ 10.3 $7.3-14.3$ $188-272$ 17.3 $14,4-20.7$ 13.5 $110-16.5$ 10.3 $7.3-14.3$ $00-0.44$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $92-152$ 14.7 $117-182$ 20.1 $16.3-24.6$ 20.3 $151-26.8$ 5.7 $280-372$ 35.5 $314-39.9$ 42.4 $372-47.9$ 43.5 $36.7-50.6$ $508-60.6$ 49.8 $45.3-54.3$ 37.4 $372-47.9$ 43.5 $36.7-50.6$ $508-60.6$ 49.8 $45.3-54.3$ 37.4 $372-47.9$ 43.5 $36.7-50.6$ $500-0.4$ 00 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ $174-24.9$ 26.7 $230-30.6$ 31.9 $274-36.8$ 35.3 $28.8-42.4$ 70 $300-38.7$ 28.4 $249-32.1$ 14.9 $274-36.8$ 35.3 $28.8-42.4$ 70 $500-38.7$ 28.4 $249-25.1$ 14.9 $249-27.6$ 34.9 $274-36.8$ 35.3 $28.8-42.4$ 70 $500-38.7$ 28.4 $249-25.0$ $230-30.6$ 31.9 $274-36.8$ 35.3 $28.8-42.4$ 70 70001 $164-26.0$ 28.4 $249-26.7$ $230-30.6$ 31.9 $274-36.8$ 25.2 $246.2-60.9$ $164-26.0$ 28.7 $230-2.21.2$ 32.7 <td< td=""><td>32.9</td><td></td><td>28.5-37.6</td><td>37.0</td><td>32.8-41.4</td><td>45.6</td><td>40.3-50.9</td><td>47.1</td><td>40.2-54.1</td><td>4.9</td><td><0.001</td></td<>	32.9		28.5-37.6	37.0	32.8-41.4	45.6	40.3-50.9	47.1	40.2-54.1	4.9	<0.001
188-772 173 $144-207$ 135 $110-16.5$ 103 $7.3-14.3$ $00-0.4$ 0.4 $0.1-12$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $9.2-15.2$ 14.7 $117-18.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $9.2-15.2$ 14.7 $117-18.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $280-37.2$ 35.5 $314-39.9$ 42.4 $372-47.6$ 43.5 $36.7-50.6$ $50.8-60.6$ 49.8 $453-54.3$ 37.4 $32.5-42.6$ 36.1 3042.7 $0.0-0.4$ 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ $174-24.9$ 26.7 $230-30.6$ 31.9 $274-36.8$ 36.1 3042.7 $300-38.7$ 28.4 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ $300-38.7$ 28.8 20.7 31.9 $274-36.8$ 35.3 $28.8-42.4$ 7.0 6.0001 $300-38.7$ 28.4 20.7 210.2 214.2 <td>188-272 173 $144-207$ 135 $110-165$ 103 $7.3-14.3$ $00-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $9.2-152$ 14.7 $11.7-182$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $9.2-152$ 14.7 $11.7-182$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $280-372$ 35.5 $314-39.9$ 424 $372-47.9$ 435 5.7 <0001 $280-30.6$ 49.8 $45.3-54.3$ 37.4 $322-47.6$ 36.1 $30.0-42.7$ 0.001 $174-249$ 26.7 $230-30.6$ 31.9 $274-36.8$ 35.3 $28-42.4$ 7.0 <0001 $174-240$ 0.0 0.1 $0.1-0.5$ 0.1 $0.0-0.7$ $0.0-2.2$ $230-30.6$ 31.9 $274-36.8$ 35.3 $28-7-20.6$ 0.001 $174-249$ 2.03 $214-36.8$ 35.3 $28-42.4$ 7.0 <0.001 $300-387$ 284 $249.2-36.8$ $274-36.8$ 3</td> <td>44.4</td> <td></td> <td>39.5-49.4</td> <td>45.3</td> <td>40.8-49.8</td> <td>40.1</td> <td>34.9-45.6</td> <td>41.1</td> <td>34.5-48.1</td> <td></td> <td></td>	188-272 173 $144-207$ 135 $110-165$ 103 $7.3-14.3$ $00-0.4$ 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $9.2-152$ 14.7 $11.7-182$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $9.2-152$ 14.7 $11.7-182$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $280-372$ 35.5 $314-39.9$ 424 $372-47.9$ 435 5.7 <0001 $280-30.6$ 49.8 $45.3-54.3$ 37.4 $322-47.6$ 36.1 $30.0-42.7$ 0.001 $174-249$ 26.7 $230-30.6$ 31.9 $274-36.8$ 35.3 $28-42.4$ 7.0 <0001 $174-240$ 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ $0.0-2.2$ $230-30.6$ 31.9 $274-36.8$ 35.3 $28-7-20.6$ 0.001 $174-249$ 2.03 $214-36.8$ 35.3 $28-42.4$ 7.0 <0.001 $300-387$ 284 $249.2-36.8$ $274-36.8$ 3	44.4		39.5-49.4	45.3	40.8-49.8	40.1	34.9-45.6	41.1	34.5-48.1		
0.0-0.4 0.4 $0.1-1.2$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $9.2-15.2$ 14.7 $11.7-18.2$ 20.1 $16.3-24.6$ 20.3 $15.1-26.8$ 5.7 <0001 $280-37.2$ 35.5 $31.4-39.9$ 42.4 $37.2-47.9$ 43.5 $36.7-50.6$ <0001 $280-60.6$ 49.8 $45.3-54.3$ 37.7 $37.2-47.9$ 43.5 $36.7-50.6$ <0001 $508-60.6$ 49.8 $45.3-54.3$ $37.2-47.9$ 43.5 $36.7-50.6$ <0001 $0.0-0.4$ 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-42.7$ $174-24.9$ 26.7 $23.0-30.6$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 $0.0-0.4$ 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ $0.0-0.7$ $174-24.9$ 26.7 $23.0-30.6$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 $300-38.7$ 28.4 $24.9-32.1$ 14.9 $12.0-18.3$ 23.8 $19.0-29.4$ $16.4-26.0$ 23.2 $19.4-27.6$ 34.8 $29.7-40.2$ 24.0 $18.3-30.7$ $20.1-28.5$ 21.8 $18.1-26.0$ 18.5 $14.6-23.2$ 16.9 $12.4-22.7$ $20.1-28.5$ 21.8 $18.1-26.0$ 18.5 $27.6-37.7$ 22.7 $24.2-38.8$ 3.0 $20.1-28.5$ 28.7 $28.8-12.6$ 38.7 $24.2-5.3$ $24.2-6.9$ $24.2-6.7$ $24.2-6.9$ $20.1-28.5$ 28.7 28.7 $27.6-37.7$	00-04 04 $0.1-12$ 0.9 $0.4-1.9$ 1.5 $0.6-4.2$ $9.2-15.2$ 14.7 $11.7-18.2$ 20.1 $16.3-24.6$ 20.3 $15.1-26.8$ 5.7 <0.001 $280-37.2$ 35.5 $314-39.9$ 42.4 $372-47.9$ 43.5 $36.7-50.6$ 5.7 <0.001 $280-37.2$ 35.5 $314-39.9$ 42.4 $372-47.9$ 43.5 $36.7-50.6$ 5.7 <0.001 $280-60.6$ 49.8 $45.3-54.3$ 37.4 $32.5-42.6$ 36.1 3042.7 <0.001 $17.4-24.9$ 26.7 $23.0-30.6$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 <0.001 $300-387$ 28.4 0.0 $ 0.1-0.5$ 0.1 $0.0-0.7$ $0.0-0.7$ $300-387$ 28.4 0.0 $2.30-30.6$ 31.9 $120-29.4$ 7.0 <0.001 $300-30.6$ 23.7 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 </td <td>22.7</td> <td></td> <td>18.8-27.2</td> <td>17.3</td> <td>14.4-20.7</td> <td>13.5</td> <td>11.0-16.5</td> <td>10.3</td> <td>7.3-14.3</td> <td></td> <td></td>	22.7		18.8-27.2	17.3	14.4-20.7	13.5	11.0-16.5	10.3	7.3-14.3		
9.2-15.2 14.7 $11.7-18.2$ 20.1 $16.3-24.6$ 20.3 $15.1-26.8$ 5.7 <0001 $28.0-37.2$ 35.5 $31.4-39.9$ 42.4 $37.2-47.9$ 43.5 $36.7-50.6$ <0001 $28.0-37.2$ 35.5 $31.4-39.9$ 42.4 $37.2-47.9$ 43.5 $36.7-50.6$ <0001 $50.8-60.6$ 49.8 $45.3-54.3$ 37.4 $32.5-42.6$ 36.1 $30.0-42.7$ <0001 $0.0-0.4$ 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ <0.001 $174-24.9$ 26.7 $23.0-30.6$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 <0.001 $174-24.9$ 26.7 $23.0-30.6$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 <0.001 $30.0-38.7$ 28.4 26.7 $23.0-30.6$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 <0.001 $30.0-38.7$ 28.4 26.7 $23.0-30.6$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 <0.001 $30.0-38.7$ 28.4 $24.9-32.1$ 14.9 $12.0-18.3$ 23.8 $19.0-29.4$ 7.0 <0.001 $16.4-26.0$ 23.2 $19.4-27.6$ 34.8 $297-40.2$ 24.0 $18.3-30.7$ $<0.00120.1-28.521.818.1-26.018.514.6-23.216.912.4-22.7<0.00121.2-77238.734.323.527.6-37.732.326.4-38.83.0$	9.2-15.2 14.7 11.7-18.2 20.1 16.3-24.6 20.3 15.1-26.8 5.7 <0.001	0.1		0.0-0.4	0.4	0.1-1.2	0.9	0.4-1.9	1.5	0.6-4.2		
$\begin{array}{llllllllllllllllllllllllllllllllllll$	$ \begin{array}{llllllllllllllllllllllllllllllllllll$	11.8		9.2-15.2	14.7	11.7-18.2	20.1	16.3-24.6	20.3	15.1-26.8	5.7	<0.001
508-60.6 49.8 $45.3-54.3$ 37.4 $32.5-42.6$ 36.1 $30.0-42.7$ $0.0-0.4$ 0.0 - 0.1 $0.1-0.5$ 0.1 $0.0-0.7$ $174-24.9$ 26.7 $23.0-30.6$ 31.9 $274-36.8$ 35.3 $28.8-42.4$ 7.0 $30.0-38.7$ 28.4 $23.0-30.6$ 31.9 $274-36.8$ 35.3 $28.8-42.4$ 7.0 $30.0-38.7$ 28.4 $24.9-32.1$ 14.9 $12.0-18.3$ 23.8 $19.0-29.4$ $30.0-38.7$ 23.2 $19.4-27.6$ 34.8 $29.7-40.2$ 24.0 $18.3-30.7$ $20.1-28.5$ 21.8 $18.1-26.0$ 18.5 $14.6-23.2$ 16.9 $12.4-22.7$ $20.1-28.5$ 21.8 $18.1-26.0$ 18.5 $14.6-23.2$ 16.9 $12.4-22.7$ $41.2-51.2$ 38.7 $34.3-43.2$ 32.5 $27.6-37.7$ 32.3 $26.4-38.8$ 3.0 $37.5-47.2$ 48.6 $44.2-53.1$ 49.5 $44.2-54.9$ 53.2 $46.3-60.0$ $7.8-12.6$ 10.3 $8.4-12.6$ 14.9 $11.1-19.6$ 11.4 $8.4-15.2$	508-60.6 49.8 $45.3-54.3$ 37.4 $32.5-42.6$ 36.1 $30.0-42.7$ $0.0-0.4$ 0.0 $ 0.1$ $0.1-0.5$ 0.1 $0.0-0.7$ $17.4-24.9$ 26.7 $230-30.6$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 $30.0-38.7$ 28.4 $2.0.0$ 31.9 $27.4-36.8$ 35.3 $28.8-42.4$ 7.0 $30.0-38.7$ 28.4 $2.9,-32.1$ 14.9 $12.0-18.3$ $28.8-42.4$ 7.0 $30.0-38.7$ 28.4 $2.4.9-32.1$ 14.9 $12.0-18.3$ $28.8-42.4$ 7.0 $30.0-38.7$ 28.4 $2.9,-32.1$ 14.9 $12.0-18.3$ $28.8-42.4$ 7.0 $30.0-38.7$ 28.4 $2.9,-32.1$ 14.9 $12.0-18.3$ $28.8-42.4$ 7.0 $30.0-38.7$ 28.4 $24.9-36.0$ 23.2 $19.4-27.6$ 34.8 $29.7-40.2$ 24.0 $18.3-30.7$ $20.1-28.5$ 21.8 $18.1-26.0$ 18.5 $14.6-23.2$ 16.9 $12.4-22.7$ 30.0002 $37.5-47.2$ 48.6 $44.2-54.9$ 53.2 $46.3-60.0$ 30.0002 $7.8-12.6$ 10.3 $8.4-12.6$ $14.2-54.9$ 53.2 $46.3-60.0$ $7.9-31.1$ 2.4 $1.4-42$ 3.2 $2.1-4.8$ 3.1 $1.7-5.8$ $0.9-3.1$ 2.4 $1.4-42$ 3.2 $2.1-4.8$ 3.1 $1.7-5.8$ $0.9-3.1$ 2.4 3.1 $1.7-5.8$ 0.0002	32.4		28.0-37.2	35.5	31.4-39.9	42.4	37.2-47.9	43.5	36.7-50.6		
$\begin{array}{cccccccccccccccccccccccccccccccccccc$	0.0-0.4 0.0 - 0.1 0.1-0.5 0.1 0.0-0.7 17.4-24.9 26.7 23.0-30.6 31.9 27.4-36.8 35.3 28.8-42.4 7.0 <0.001	55.7		50.8-60.6	49.8	45.3-54.3	37.4	32.5-42.6	36.1	30.0-42.7		
	0 17.4-24.9 26.7 23.0-30.6 31.9 27.4-36.8 35.3 28.8-42.4 7.0 <0.001	0.1		0.0-0.4	0.0	I	0.1	0.1-0.5	0.1	0.0-0.7		
2 30.0-38.7 28.4 24.9-32.1 14.9 12.0-18.3 23.8 19.0-29.4 3 16.4-26.0 23.2 19.4-27.6 34.8 29.7-40.2 24.0 18.3-30.7 1 20.1-28.5 21.8 19.4-27.6 34.8 29.7-40.2 24.0 18.3-30.7 1 20.1-28.5 21.8 18.1-26.0 18.5 14.6-23.2 16.9 12.4-22.7 2 41.2-51.2 38.7 34.3-43.2 32.5 27.6-37.7 32.3 26.4-38.8 3.0 0.002 3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 7.8-12.6 7.8-12.6 14.9 11.1-19.6 11.4 8.4-15.2 7.8	2 30.0-38.7 28.4 24.9-32.1 14.9 12.0-18.3 23.8 19.0-29.4 8 16.4-26.0 23.2 19.4-27.6 34.8 29.7-40.2 24.0 18.3-30.7 1 20.1-28.5 21.8 19.4-27.6 34.8 29.7-40.2 24.0 18.3-30.7 1 20.1-28.5 21.8 18.1-26.0 18.5 14.6-23.2 16.9 12.4-22.7 2 41.2-51.2 38.7 34.3-43.2 32.5 27.6-37.7 32.3 26.4-38.8 3.0 0.002 3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 3 7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 84-15.2 0.9-3.1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 1.4.2.1.1.1	20.	6	17.4-24.9	26.7	23.0-30.6	31.9	27.4-36.8	35.3	28.8-42.4	7.0	<0.001
3 16.4-26.0 23.2 19.4-27.6 34.8 29.7-40.2 24.0 18.3-30.7 1 20.1-28.5 21.8 18.1-26.0 18.5 14.6-23.2 16.9 12.4-22.7 2 41.2-51.2 38.7 34.3-43.2 32.5 27.6-37.7 32.3 26.4-38.8 3.0 0.002 3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2	8 16.4-26.0 23.2 19.4-27.6 34.8 29.7-40.2 24.0 18.3-30.7 1 20.1-28.5 21.8 18.1-26.0 18.5 14.6-23.2 16.9 12.4-22.7 2 41.2-51.2 38.7 34.3-43.2 32.5 27.6-37.7 32.3 26.4-38.8 3.0 0.002 3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 3 7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2 0.9-3.1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 1.4.1.1.1.1.1.1.1.1.1.1.1.1.1.1.1.1.1.1	34.	2	30.0-38.7	28.4	24.9-32.1	14.9	12.0-18.3	23.8	19.0-29.4		
1 20.1–28.5 21.8 18.1–26.0 18.5 14.6–23.2 16.9 12.4–22.7 2 41.2–51.2 38.7 34.3–43.2 32.5 27.6–37.7 32.3 26.4–38.8 3.0 0.002 3 37.5–47.2 48.6 44.2–53.1 49.5 44.2–54.9 53.2 46.3–60.0 7.8–12.6 10.3 8.4–12.6 14.9 11.1–19.6 11.4 8.4–15.2	1 20.1-28.5 21.8 18.1-26.0 18.5 14.6-23.2 16.9 12.4-22.7 2 41.2-51.2 38.7 34.3-43.2 32.5 27.6-37.7 32.3 26.4-38.8 3.0 0.002 3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 7 7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2 0.9-3.1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8	20.8	~	16.4–26.0	23.2	19.4-27.6	34.8	29.7-40.2	24.0	18.3-30.7		
2 41.2-51.2 38.7 34.3-43.2 32.5 27.6-37.7 32.3 26.4-38.8 3.0 0.002 3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 7 7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2	2 41.2-51.2 38.7 34.3-43.2 32.5 27.6-37.7 32.3 26.4-38.8 3.0 0.002 3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2 0.9-3.1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8	24.	H	20.1-28.5	21.8	18.1-26.0	18.5	14.6-23.2	16.9	12.4-22.7		
3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2 $\frac{5}{55}$	3 37.5-47.2 48.6 44.2-53.1 49.5 44.2-54.9 53.2 46.3-60.0 7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2 0.9-3.1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 275	46.	7	41.2-51.2	38.7	34.3-43.2	32.5	27.6-37.7	32.3	26.4-38.8	3.0	0.002
7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2	7.8-12.6 10.3 8.4-12.6 14.9 11.1-19.6 11.4 8.4-15.2 N 0.9-3.1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N N <t< td=""><td>42</td><td>3</td><td>37.5-47.2</td><td>48.6</td><td>44.2-53.1</td><td>49.5</td><td>44.2-54.9</td><td>53.2</td><td>46.3-60.0</td><td></td><td></td></t<>	42	3	37.5-47.2	48.6	44.2-53.1	49.5	44.2-54.9	53.2	46.3-60.0		
	0.9-3.1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 [V 1 1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 [V 1 1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 [V 1 1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 [V 1 2.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 [V 1 3.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 [V 1 4.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 [V 1 4.4 1.4-4.2 3.2 2.1-4.8 3.1 1.7-5.8 [V 1 4.4 1.4 1.4 1.4 1.4 1.4 1.4 1.4 1.4 1.	.6	6	7.8-12.6	10.3	8.4-12.6	14.9	11.1-19.6	11.4	8.4-15.2		

TABLE 1 Descriptive characteristics of adult daily cigarette smokers in the United States, Canada, England and Australia who had not vaped in the past 6 months at baseline, by cigarette guit

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		Cigarette qui	t plans at baseline								
		Planning to q 6 months n =	uit within the next 897	Planning to qui beyond 6 montl	t sometime in future, hs n = 1102	Not plan n = 937	ning to quit	Unknown n = 469	plans to quit		
Descriptive characterist	tics at baseline	%	95% CI	%	95% CI	%	95% CI	%	95% CI	x ²	P-value
Quitting self-efficacy	Not at all/slightly sure $(n = 1943)$	39.7	34.8-44.8	59.8	55.3-64.1	64.7	59.3-69.7	57.9	51.0-64.5	20.7	<0.001
	Moderately sure (n = 855)	38.6	34.0-43.4	28.4	24.4-32.7	13.7	10.7-17.5	18.0	13.6-23.5		
	Very/extremely sure (n = 394)	20.4	16.8-24.7	9.4	7.0-12.5	10.4	7.8-13.9	8.7	5.0-14.7		
	Do not know ($n = 210$)	1.3	0.7-2.2	2.4	1.7-3.5	11.2	7.7-16.1	15.4	11.6-20.2		
Desire to quit	Not at all (<i>n</i> = 476)	0.6	0.2-2.1	0.8	0.4-1.7	43.8	38.7-49.2	4.8	2.5-9.2	83.6	<0.001
	A little (<i>n</i> = 649)	3.3	1.6-6.7	18.5	15.0-22.5	32.5	27.9-37.6	22.3	17.4-28.1		
	Somewhat $(n = 1140)$	26.3	22.2-30.9	52.7	48.2-57.1	19.2	14.9-24.3	42.0	35.6-48.8		
	A lot (n = 1062)	69.7	64.9-74.2	27.3	23.7-31.2	1.8	1.0-3.1	18.4	13.3-24.8		
	Do not know $(n = 71)$	0.1	0.0-0.4	0.8	0.4-1.6	2.7	1.3-5.6	12.5	7.4-20.3		
SES = socioeconomic sta	atus										

Cigarette smoking cessation at follow-up as a function of vaping uptake, stratified by cigarette quit plans at baseline

ADDICTION

Overall, cigarette smoking cessation rates at follow-up were highest among those who were initially planning to quit within the next 6 months (19.3% quit; 95% CI, 15.4–24.0) and were lowest among those who were initially not planning to quit at all (7.1% quit; 95% CI, 4.9–10.1) (Table 3).

A significant interaction was found between cigarette guit plans at baseline and vaping uptake in the association with cigarette smoking cessation. Among smokers who at baseline were planning to guit within the next 6 months, daily vaping uptake was associated with a nearly twofold higher odds of smoking cessation although this estimate did not reach statistical significance (adjusted odds ratio [AOR]. 1.91; 95% CI, 0.91-4.00), whereas among those who at baseline were not planning to quit within the next 6 months, those who took up daily vaping experienced over eightfold higher odds of quitting smoking compared with those who did not take up vaping (AOR, 8.58; 95% Cl. 5.06-14.54) (Table 3). When disaggregating those who were not planning to guit within the next 6 months, the strongest associations between uptake of daily vaping and smoking cessation were found among those who at baseline were planning to guit sometime in the future beyond 6 months (AOR, 10.88; 95% CI, 5.31-22.30) and those who at baseline did not know whether they planned to guit (AOR, 8.64; 95% CI, 3.05-24.46) (Table 3).

Uptake on non-daily vaping was negatively associated with smoking cessation among those who at baseline were not planning to quit at all (AOR, 0.18; 95% CI, 0.06–0.55); no other significant differences were observed (Table 3). Findings were consistent in sensitivity analyses in which the definition of smoking cessation considered those who smoked less than monthly as having not achieved cessation, which changed the value of the cessation outcome for only 16 people.

Planning to quit cigarette smoking at follow-up as a function of vaping uptake, stratified by cigarette quit plans at baseline

A significant interaction between cigarette quit plans at baseline and vaping uptake was found in the association with change in plans to quit smoking. Among smokers who at baseline were planning to quit within the next 6 months, there was no statistically significant association between daily vaping uptake and change in plans to quit smoking (AOR, 0.82; 95% CI, 0.44–1.55). By contrast, among those who at baseline were not planning to quit in the next 6 months, those who took up daily vaping had an over sixfold higher odds of change to be planning to quit smoking compared with those who did not take up vaping (AOR, 6.32; 95% CI, 4.17–9.59). When disaggregating those who were not planning to quit within the next 6 months, the strongest association between uptake of daily vaping and change to be planning to quit smoking was found among those

TABLE 2 Vaping uptake between baseline and follow-up as a function of cigarette quit plans at baseline among adult daily cigarette smokers in the United States, Canada, England and Australia who had not vaped in the past 6 months at baseline

	Vaping	uptake	between baseli	ine and i	follow-up)				
	No vap	ing upta	ke	Non-a	daily vap	ing uptake	Daily	vaping u	ptake	
Cigarette quit plans at baseline	n	%	95% CI	n	%	95% CI	n	%	95% CI	χ^2 P-value
Overall: N = 3405	2149	62.9	60.4-65.4	803	23.6	21.5-25.9	453	13.5	11.8-15.4	-
Planning to quit within the next 6 months: n = 897 (27% of sample)	540	59.0	54.1-63.8	215	24.7	20.7-29.1	142	16.3	13.0-20.3	2.4 0.089
Not planning to quit within next 6 months (note: this group is the aggregate of the groups beneath it): <i>n</i> = 2508 (73% of sample)	1609	64.4	61.4-67.3	588	23.2	20.7-25.9	311	12.4	10.5-14.6	
Planning to quit sometime in the future, beyond 6 months: <i>n</i> = 1102 (34% of sample)	631	57.8	53.3-62.2	308	27.6	23.7-31.8	163	14.6	11.7-18.1	4.3 < 0.001
Not planning to quit: $n = 937$ (26% of sample)	682	72.6	67.5-77.0	161	17.1	13.5-21.4	94	10.3	7.4-14.2	
Unknown plans to quit: <i>n</i> = 469 (13% of sample)	269	65.3	58.7-71.3	119	24.0	18.8-30.1	54	10.8	7.6-15.1	

ns are unweighted and reflect the number of observations; %s and 95% Cl are weighted.

who at baseline were not planning to quit smoking at all (AOR, 10.12; 95% CI, 4.46–22.93).

Finally, in a sensitivity analysis excluding those who had quit smoking at follow-up, we still found a statistically significant positive association between daily vaping uptake and change in plans to quit smoking among those who at baseline were not planning to quit in the next 6 months (AOR, 3.61; 95% Cl, 1.92–6.79) and among the subgroup who at baseline were not planning to quit at all (AOR, 11.65; 95% Cl, 3.48–38.99) (Table 4).

DISCUSSION

The key finding from this study is that there is a strong positive association between vaping uptake and cigarette smoking cessation among smokers with no initial plans to quit smoking. Specifically, those not planning to quit in the next 6 months who started vaping daily experienced a 32% cigarette quit rate compared with a 7% cigarette quit rate among their counterparts who did not take up vaping. These findings are consistent with the US PATH Study findings [2, 3] and indicate that studies that focus on smokers who are already planning or attempting to quit may underestimate a potential benefit of daily vaping for smoking cessation at the population-level.

One mechanism by which vaping may be positively associated with smoking cessation could be by changing motivation and selfefficacy for quitting among those initially not interested in quitting smoking. Our findings show that vaping uptake was associated with change in plans to quit smoking among those initially not planning to quit, which is consistent with the one other population-based study that investigated this question [2]. Further, in a sensitivity analysis among those who at baseline did not plan to quit at all and continued to smoke at follow-up, we found that they experienced an 11-fold greater odds of planning to quit at follow-up when they took up daily vaping, suggesting that there could be further increases in smoking cessation rates among this 'hard-to-reach' group of smokers in the longer term. Importantly, our findings show that these smokers tended to be male, older, smoked more cigarettes per day and were of lower SES than their counterparts who were initially planning to quit.

It is also possible that smokers in our study changed their plans to quit after baseline and subsequently began vaping as a method of quitting. We had between 18- and 24-month intervals between surveys, and we could not assess whether quit intentions changed first or whether vaping status changed first. Regardless of the directionality, our findings show contemporaneous associations between moving from daily smoking to no smoking alongside moving from no vaping to daily vaping specifically among those who were initially not planning to quit smoking. These findings are consistent with the one other population study that evaluated this question among smokers not planning to quit [3], as well as experimental studies that have shown that giving NVPs to smokers unmotivated to quit increases their motivation to quit [4–6] and is associated with reductions in their cigarette smoking [6,7].

Consistent with other studies in the literature, we found no association between uptake of non-daily vaping and smoking cessation [20–22]. This may reflect that non-daily vaping was not associated with change in quit intentions and/or may reflect that non-daily vaping did not provide sufficient nicotine to serve as a substitute for those who were daily cigarette smokers, as discussed previously by Gravely *et al.* [20]. Importantly, we found that uptake of non-daily vaping was nearly twice as common as daily vaping uptake, meaning that the majority of vaping uptake occurring among smokers in the population is not expected to yield smoking cessation gains.

When considering differences in the magnitudes of associations between daily vaping uptake and smoking cessation when

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TABLE 3 Cigarette smoking cessation at follow-up as a function of vaping uptake between baseline and follow-up among adult daily cigarette smokers in the United States, Canada, England, and Australia who had not vaped in the past 6 months at baseline, stratified by cigarette quit plans at baseline

		Cigarette	smoking cessation at follow	-up ^a
Cigarette quit plans at baseline	Vaping uptake between baseline and follow-up (inter-wave vaping uptake)	n	% (95% CI)	AOR ^b (95% CI)
Overall	Overall: N = 3405	393	12.7 (10.9,14.6)	_
Planning to quit within the	Overall: n = 897	155	19.3 (15.4,24.0)	-
next 6 months	No vaping: <i>n</i> = 540	80	16.8 (11.8,23.3)	Ref
	Any vaping: n = 357	75	23.0 (17.0,30.3)	1.65 (0.93,2.92)
	No vaping: $n = 540$	80	16.8 (11.8,23.3)	Ref
	Non-daily vaping: n = 215	37	21.6 (14.1,31.7)	1.50 (0.78,2.87)
	Daily vaping: $n = 142$	38	25.1 (16.2,36.6)	1.91 (0.91,4.00)
Not planning to quit within	Overall: <i>n</i> = 2508	238	10.1 (8.4,12.1)	-
the next 6 months (note: this	No vaping: <i>n</i> = 1609	114	6.8 (5.3,8.8)	Ref
three groups beneath it)	Any vaping: $n = 471$	76	19.1 (13.7,25.9)	3.20 (2.03,5.06) ^c
	No vaping: <i>n</i> = 1609	114	6.8 (5.3,8.8)	Ref
	Non-daily vaping: n = 588	44	7.3 (4.5,11.7)	1.24 (0.68,2.27)
	Daily vaping: $n = 311$	80	32.4 (24.6,41.3)	8.58 (5.06,14.54) ^c
Planning to quit sometime in the future,	Overall: <i>n</i> = 1102	129	12.4 (9.7,15.8)	-
beyond 6 months	No vaping: <i>n</i> = 631	53	7.6 (5.5,10.5)	Ref
	Any vaping: $n = 471$	76	19.1 (13.7,25.9)	4.02 (2.19,7.39)
	No vaping: $n = 631$	53	7.6 (5.5,10.5)	Ref
	Non-daily vaping: <i>n</i> = 308	30	10.1 (5.5,17.6)	1.78 (0.87,3.65)
	Daily vaping: $n = 163$	46	36.1 (25.0,49.0)	10.88 (5.31,22.30)
Not planning to quit	Overall: n = 937	68	7.1 (4.9,10.1)	-
	No vaping: $n = 682$	40	6.1 (3.7,10.1)	Ref
	Any vaping: $n = 255$	28	9.5 (5.9,15.1)	1.73 (0.84,3.56)
	No vaping: $n = 682$	40	6.1 (3.7,10.1)	Ref
	Non-daily vaping: <i>n</i> = 161	5	1.0 (0.4,2.6)	0.18 (0.06,0.55) ^c
	Daily vaping: $n = 94$	23	23.5 (13.8,37.3)	5.81 (2.60,13.00) ^d
Unknown plans to quit ^e	Overall: <i>n</i> = 469	41	10.2 (6.9,14.9)	-
	No vaping: $n = 269$	21	6.7 (3.8,11.4)	Ref
	Any vaping: n = 173	20	16.9 (9.9,27.3)	3.23(1.24,8.41)
	No vaping: $n = 269$	21	6.7 (3.8,11.4)	Ref
	Non-daily vaping: <i>n</i> = 119	9	8.1 (3.4,17.9)	1.46 (0.47,4.53)
	Daily vaping: $n = 54$	11	36.5 (20.1,56.7)	8.64 (3.05,24.46) ^c

ns are unweighted and reflect the number of observations; %s, and adjusted odds ratios (AOR) are weighted. Generalized estimating equation (GEE) logistic regression analyses were used to assess the association between uptake of vaping (between baseline and follow-up) and cigarette cessation (at follow-up) among adult daily cigarette smokers who had not vaped in the past 6 months at baseline, stratified by cigarette quit plans at baseline, over two periods of time (i.e. wave 1 to wave 2 and wave 2 to wave 3), including up to two sets of observations per individual and statistically controlling for the correlation among observations contributed by the same individuals.

^aCigarette cessation at follow-up was defined as less than monthly cigarette smoking (vs monthly/weekly/daily cigarette smoking) among those who were daily cigarette smokers at baseline.

^bAnalyses were adjusted for country, biological sex, race/ethnicity (White, non-White), age group (18–24, 25–39, 40–54, 55+ years), socioeconomic status (SES) (low, moderate, high), cigarettes smoked per day, time in sample, and wave; all covariates were assessed at baseline; GEE models were fitted specifying the unstructured covariance and within-person correlation matrices.

^cAssociation is significantly different from analogous association among those planning to quit within the next 6 months (P = <0.01 for interaction). ^dAssociation is significantly different from analogous association among those planning to quit within the next 6 months (P = <0.06 for interaction). ^eModel convergence not achieved for those with unknown plans to quit (for either model).

		Planning follow-u	to quit smoking with (quitters included) ^a	in the next 6 months at		Plannin follow-u	g to quit smoking with p (quitters excluded) ^b	in the next 6 months at	DIC
Cigarette quit plans at baseline ^c	Vaping uptake between baseline and follow-up (inter-wave vaping uptake)	2	% (95% Cl)	AOR ^d (95% Cl)	Vaping uptake between baseline and follow-up (inter-wave vaping uptake)	2	% (95% CI)	AOR ^d (95% Cl)	TION
Overall	Overall: N = 3405	1130	34.7 (32.2,37.2)	1	Overall: <i>n</i> = 3012	742	25.3 (23.0,27.7)	1	
Planning to quit within	Overall: $n = 897$	612	70.4 (65.8,74.7)	ı	Overall: n = 742	457	63.3 (58.0,68.3)	I	
the next 6 months	No vaping: $n = 540$	359	69.9 (64.1,75.2)	Ref	No vaping: <i>n</i> = 460	279	63.9 (57.3,69.9)	Ref	
	Any vaping: <i>n</i> = 357	253	71.1 (63.3,77.9)	1.05 (0.66,1.67)	Any vaping: $n = 282$	178	62.5 (53.5,70.7)	0.92 (0.57,1.48)	SS
	No vaping: <i>n</i> = 540	359	69.9 (64.1,75.2)	Ref	No vaping: <i>n</i> = 460	279	63.9 (57.3,69.9)	Ref	jA
	Non-daily vaping: $n = 215$	150	74.2 (65.0,81.7)	1.23 (0.72,2.09)	Non-daily vaping: <i>n</i> = 178	113	67.1 (56.5,76.2)	1.12 (0.65,1.93)	—
	Daily vaping: <i>n</i> = 142	103	66.4 (52.6,77.9)	0.82 (0.44,1.55)	Daily vaping: <i>n</i> = 104	65	55.2 (40.1,69.3)	0.66 (0.34,1.28)	
Not planning to quit within the next	Overall: $n = 2508$	518	21.1 (18.8,23.7)	I	Overall: <i>n</i> = 2270	285	14.4 (10.5,14.5)	I	
6 months (note: this group is the	No vaping: <i>n</i> = 1609	276	15.9 (13.6,18.6)	Ref	No vaping: <i>n</i> = 1495	164	9.8 (8.0,12.0)	Ref	
aggregate of the three groups beneath it)	Any vaping: <i>n</i> = 899	242	30.6 (25.9,35.6)	2.62 (1.90,3.63) ^e	Any vaping: $n = 775$	121	17.4 (13.5,22.2)	2.03 (1.36,3.03) ^f	
	No vaping: <i>n</i> = 1609	276	15.9 (13.6,18.6)	Ref	No vaping: <i>n</i> = 1495	164	9.8 (8.0,12.0)	Ref	
	Non-daily vaping: <i>n</i> = 588	119	20.3 (15.9,25.6)	1.45 (1.01,2.10)^f	Non-daily vaping: <i>n</i> = 544	77	14.1 (10.7,18.5)	1.55 (1.03,2.32)	
	Daily vaping: <i>n</i> = 311	123	49.8 (41.1,58.4)	6.32 (4.17,9.59) ^e	Daily vaping: <i>n</i> = 231	44	25.9 (16.5,38.2)	3.61 (1.92,6.79) ^e	
Planning to quit sometime in the	Overall: $n = 1102$	341	30.8 (26.9,35.0)	I	Overall: $n = 973$	213	21.0 (17.6,24.9)	I	
future, beyond 6 months	No vaping: <i>n</i> = 631	171	24.8 (20.5,29.6)	Ref	No vaping: <i>n</i> = 578	118	18.6 (14.7,23.2)	Ref	
	Any vaping: <i>n</i> = 471	170	39.1 (32.4,46.2)	2.18 (1.45,3.27) ^f	Any vaping: <i>n</i> = 395	95	24.8 (18.9,31.7)	1.43 (0.90,2.28)	
	No vaping: <i>n</i> = 631	171	24.8 (20.5,29.6)	Ref	No vaping: <i>n</i> = 578	118	18.6 (14.7,23.2)	Ref	
	Non-daily vaping: <i>n</i> = 308	91	29.9 (22.8,38.1)	1.41 (0.89,2.23)	Non-daily vaping: <i>n</i> = 278	62	22.1 (16.1,29.4)	1.24 (0.75,2.04)	
	Daily vaping: <i>n</i> = 163	79	56.5 (45.0,67.3)	4.52 (2.63,7.77) ^e	Daily vaping: <i>n</i> = 117	33	31.9 (19.5,47.5)	2.02 (0.98,4.18) ^f	
Not planning to quit	Overall: $n = 937$	95	11.0 (8.0,15.0)	I	Overall: <i>n</i> = 869	31	4.5 (2.5,8.0)	I	
	No vaping: <i>n</i> = 682	58	8.4 (5.5,12.4)	Ref	No vaping: <i>n</i> = 642	20	2.6 (1.4,4.7)	Ref	
	Any vaping: <i>n</i> = 255	37	18.0 (11.0,28.1)	3.05 (1.44,6.47) ^f	Any vaping: <i>n</i> = 227	11	9.8 (3.9,22.4)	5.07 (1.84,13.93) ^f	
	No vaping: <i>n</i> = 682	58	8.4 (5.5,12.4)	Ref	No vaping: <i>n</i> = 642	20	2.6 (1.4,4.7)	Ref	
	Non-daily vaping: <i>n</i> = 161	6	4.5 (1.9,10.5)	0.72 (0.25,2.06)	Non-daily vaping: <i>n</i> = 156	5	3.7 (1.3,10.1)	2.41 (0.73,7.98)	
	Daily vaping: <i>n</i> = 94	28	40.4 (24.7,58.5)	10.12 (4.46,22.93) ^e	Daily vaping: <i>n</i> = 71	6	22.9 (7.5,52.0)	11.65 (3.48,38.99) ^e	
Unknown plans to quit ^g	Overall: <i>n</i> = 469	82	16.3 (12.2,21.3)	I	Overall: <i>n</i> = 428	41	6.7 (4.5,9.9)	I	К/
								(Continues)	ASZA

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1360043, 2023, 2. Downloaded from https://onlinelibrary.wiley.com/doi/10.1111/add.16050 by University Of Waterloo Dama Parter Library, Wiley Online Library on [1007/2023]. See the Terms and Conditions (https://onlinelibrary.wiley.com/ems-and-conditions) on Wiley Online Library for rules of use; OA articles are governed by the applicable Creative Commons License

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No vaping: $n = 296$ 47 124 ($8.5,17$ 8)RefNo vaping: $n = 275$ 26 $6.1(3,7,9,9)$ RefAny vaping: $n = 173$ 35 2.35 ($15.6,3.3,9$) 2.35 (112.494)Any vaping: $n = 153$ 15 8.0 ($4.2.14.8$) 128 ($0.4,3.24$)No vaping: $n = 179$ 35 2.35 ($15.6,3.3,9$) 2.35 (112.434)Any vaping: $n = 133$ 15 8.0 ($4.2.14.8$) 128 ($0.4,3.26$)Non-daily vaping: $n = 296$ 47 12.4 ($8.1,24.5$) 1.34 ($0.21,295$)Non-daily vaping: $n = 110$ 10 6.9 ($3.3.14.1$) 1.22 ($0.46,3.26$)Daily vaping: $n = 54$ 16 4.39 ($2.96.62.4$) 5.5 ($5.128,1358$) $Daily vaping: n = 110106.9 (3.3.14.1)1.22 (0.46,3.26)Promovel of observations: 8.5 and adjusted odds ratios (0.01,2.95)Non-daily vaping: n = 110106.9 (3.3.14.1)1.22 (0.46,3.26)Promovel of observations: 8.5 and adjusted odds ratios (0.01) are weighted.8.6 (3.3.14.1)1.20 (0.4,3.20)Promovel of observations: 8.5 and adjusted odds ratios (0.01) are weighted.8.13 (3.9.2, 9.14 (0.2,7.27)*Promovel of observations: 8.5 and adjusted odds ratios (0.01) are weighted.8.13 (3.9.2, 9.14 (0.29,7.27)*Promovel of observations: 8.5 and adjusted odds ratios (0.01) are weighted.8.33 (3.14.1)1.22 (0.4,5.2.26)Promovel of observations: 8.5 and adjusted odds ratios (0.01) are weighted.8.32 (3.218,128)1.44 (0.27,272Promovel of observations promovel of outservaled in the out of the out of the ou$	Cigarette quit plans at baseline ^c	Vaping uptake between baseline and follow-up (inter-wave vaping uptake)	2	% (95% Cl)	AOR ^d (95% Cl)	Vaping uptake between baseline and follow-up (inter-wave vaping uptake)	2	% (95% CI)	AOR ^d (95% CI)
Any vaping: $n = 173$ 35235 (13.6,33.9)235 (13.6,43.4)Any vaping: $n = 133$ 1580 (42.148)128 (0.49.3.34)No vaping: $n = 296$ 4712.4 (8.5,178)RefNovaping: $n = 275$ 266.1 (3.7,99)RefNon-daily vaping: $n = 296$ 4712.4 (8.1,12.4.5)1.34 (0.6,1.2.95)Non-daily vaping: $n = 110$ 106.9 (3.3,14.1)1.22 (0.46.3.26)Daily vaping: $n = 54$ 164.39 (2.6.9,6.2.4)5.45 (2.18,13.50)*Daily vaping: $n = 130$ 1.06.9 (3.3,14.1)1.22 (0.46.3.26)reare unweighted and reflect the number of observations; %s, and adjusted odds ratios (AOR) are veripted. Generalized stimating equation (GEE) logistic regression analyses were used to assess the association between uptake of vaping (between baseline and follow-up) and planning to quit cigarette smoking at follow-up among aduit daily cigarette smokers who had not vaped in the past 6 months at baseline, tower two periods of time (i.e. wave 1 to wave 2 to wave 2), including up to two sets of observations per individual and statistically controlThose who bad quit smoking than the next 6 months at follow-up was coled vas for the corner of observations gen in the past 6 months at baseline; GE models were not smoking at least monthy and were not asked about their quitting plans (to retain in the analysis thosThose who were adjusted for country, biological sex, race/ethnicity (White, non-White), age group (18-24, 25-39, 40-54, 55+ years), socioeconomic status (GE) (with correlin), cigarettes smoked pout their quitting plans."Those who were adjusted for country, biological sex, race/ethnicity (White, non-White), age group (18-24, 25-39, 40-54, 55+ years), socioeconomic status (GE) (with edmetaes and whote and wave allow-anal		No vaping: <i>n</i> = 296	47	12.4 (8.5,17.8)	Ref	No vaping: <i>n</i> = 275	26	6.1 (3.7,9.9)	Ref
No vaping: $n = 296$ 4712.4 (6.5, 17.8)RefNo vaping: $n = 275$ 266.1 (3.7, 9.9)RefNon-daily vaping: $n = 119$ 1914.4 (8, 1, 24.5)1.34 (0.6.1, 2.95)Non-daily vaping: $n = 110$ 106.9 (3.3, 14.1)1.22 (0.46.3.26)Daily vaping: $n = 54$ 1.64.39 (26.9, 62.4)5.45 (2.18, 13.59)°Daily vaping: $n = 132$ 1.24 (0.297, 27)°rase unweighted and reflect the number of observations; %s, and adjusted odds ratios (AOR) are weighted. Generalized estimating equation (GEE) logistic regression analyses were used to assess the asseline and follow-up) and planning to quit cigarette smoking at follow-up anorg dolt daily (grarette smokers who had not vaped in the past 6 months at baseline. Aver two periods of time (i.e. wave 1 to wave 2 to wave 3), including up to two sets of observations per individual and statistically control for the correlation among observations contributed by the same individuals.'Planning to quit smoking.'Planning to quit smoking at baseline, over two periods of time (i.e. wave 1 to wave 2 to wave 3), including up to two sets of observations per individual and statistically control for the correlation among observations contributed by the same individuals.'Planning to quit smoking.'Planning to quit smoking at baseline and follow-up was coded ves for those who were not smoking at least monthly and were not asked about their quitting plans (to retain in the analysis thos who had quit smoking.'Planning to quit smoking.'Planning to quit within the next 6 months at baseline includes those who were planning to quit within nementator of the outcome for th		Any vaping: $n = 173$	35	23.5 (15.6,33.9)	2.35 (1.12,4.94)	Any vaping: <i>n</i> = 153	15	8.0 (4.2,14.8)	1.28 (0.49,3.34)
Non-daily vaping: $n = 119$ 19144 (8.1.24.5)1.34 (0.6.1.295)Non-daily vaping: $n = 110$ 106.9 (3.3.1.4.1)1.22 (0.46.3.26)Daily vaping: $n = 54$ 1643.9 (26.9.62.4)5.45 (2.18.13.58)*Daily vaping: $n = 43$ 51.16 (13.4.32.8)1.44 (0.29.7.27)*ras are unweighted and reflect the number of observations; %s, and adjusted odds ratios (AOR) are weighted. Generalized estimating equation (GEE) logistic regression analyses were used to assess the association between uptake of vaping (between baseline, over two periods of time (i.e. wave 1 to wave 2 to wave 3), including up to two sets of observations per individuals1.24 (0.29.7.27)*Planning to quit smoking within the next 6 months at baseline, over two periods of time (i.e. wave 1 to wave 2 and wave 2 to wave 3), including up to two sets of observations per individuals1.24 (0.24.5)Planning to quit smoking within the next 6 months at baseline, over two periods of time (i.e. wave 1 to wave 2 and wave 2 to wave 3), including up to two sets of observations per individuals.1.46 (0.29.7.20)*Planning to quit smoking within the next 6 months at baseline includes those who were not smoking at least monthly and were not asked about their quitting plans (to retain in the analysis thos who had quit smoking).1.46 (0.29.7.20)*Planning to quit smoking.1.40 (0.29.7.20)*1.40 (0.29.2.2.5)*1.44 (0.29.2.5)*Planning to quit smoking within the next 6 months at baseline includes those who were planning to quit within one month at baseline end those who were planning to quit between 1 and 6 month at baseline, the numerator of the outcome for the outcome for the other baseline end in the next 6 months at baseline includes those were were blanning to quit within the next 6 months		No vaping: <i>n</i> = 296	47	12.4 (8.5,17.8)	Ref	No vaping: <i>n</i> = 275	26	6.1 (3.7,9.9)	Ref
Daily vaping: $n = 54$ 16439 (26.9,62.4)5.45 (2.18.13.50)*Daily vaping: $n = 43$ 511.6 (3.4,32.8)1.44 (0.29.7.27)*rowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspacerowspace		Non-daily vaping: <i>n</i> = 119	19	14.4 (8.1,24.5)	1.34 (0.61,2.95)	Non-daily vaping: $n = 110$	10	6.9 (3.3,14.1)	1.22 (0.46,3.26)
In are unweighted and reflect the number of observations; %s, and adjusted odds ratios (AOR) are weighted. Generalized estimating equation (GEE) logistic regression analyses were used to assess the association between uptake of vaping (between baseline and follow-up) and planning to quit cigarette smoking at follow-up among adult daily cigarette smokers who had not vaped in the past 6 months at baseline, stratified by plans to quit smoking at baseline, over two periods of time (i.e. wave 1 to wave 2 and wave 2 to wave 3), including up to two sets of observations per individual and statistically control for the correlation among observations contributed by the same individuals. "Planning to quit smoking within the next 6 months at follow-up was coded yes for those who were not smoking at least monthly and were not asked about their quitting plans (to retain in the analysis thos who had quit smoking). "Those who wee planning to quit within the next 6 months at baseline includes those who were planning to quit within one month at baseline and those who were planning to quit plans. Those who were planning to quit within one month at baseline and those who were planning to quit plans. Those who were planning to quit within none month at baseline and those who were planning to quit plans. The numerator of the outcome for the outcome show and who were planning to quit within the next 6 months at assessed at baseline. Generating manyes were adjusted for country, biological sex. Tac/ethnicity (White, non-White), age group (18-24, 25-39, 40-54, 55+ years), socioeconomic status (SES) (low, moderate, high), cigarettes smoked per time in sample, and wave: all covariates were assessed at baseline. Generation anong those planning to quit within the next 6 months at baseline includes who were planning to quit within the next 6 month at baseline and those who were planning to quit baseline and those who were assesse		Daily vaping: $n = 54$	16	43.9 (26.9,62.4)	5.45 (2.18,13.58) ^e	Daily vaping: <i>n</i> = 43	5	11.6 (3.4,32.8)	1.44 (0.29,7.27) ^e
^c Those who were planning to quit within the next 6 months at baseline includes those who were planning to quit within the next 6 month at baseline and those who were planning to quit within the next 6 months quit plans. The numerators of the outcome for the other baseline quit intentions group reflect changing to have quit plans. The numerator of the outcome for this group reflect smaintaining quit plans. The numerators of the outcome for the other baseline quit intentions group reflect changing to have quit plans. ^d Analyses were adjusted for country, biological sex, race/ethnicity (White, non-White), age group (18–24, 25–39, 40–54, 55+ years), socioeconomic status (SES) (low, moderate, high), cigarettes smoked per time in sample, and wave; all covariates were assessed at baseline; GEE models were fitted specifying the unstructured covariance and within-person correlation matrices. ⁴ Association is significantly different from analogous association among those planning to quit within the next 6 months (<i>P</i> = <0.01 for interaction). ^f Association is significantly different from analogous association among those planning to quit within the next 6 months (<i>P</i> = <0.05 for interaction). ^f Association is significantly different from analogous association among those planning to quit within the next 6 months (<i>P</i> = <0.05 for interaction).	<i>ns</i> are unweighted and reflect the nu association between uptake of vapin baseline, stratified by plans to quit sr for the correlation among observatio ^a Planning to quit smoking within the who had quit smoking.	mber of observations; %s, and ad g (between baseline and follow-u moking at baseline, over two perio ins contributed by the same indivi- next 6 months at follow-up was- vortured from analysis	ijusted od ip) and pla ods of tim iduals. coded yes	ds ratios (AOR) are w anning to quit cigarett e (i.e. wave 1 to wave : for those who were	reighted. Generalized esti te smoking at follow-up a e 2 and wave 2 to wave 3 e 2 mor smoking at least mor not smoking at least mor	mating equation (GEE) logistic mong adult daily cigarette sm the including up to two sets of (ithly and were not asked abou	c regressi lokers wh observati ut their qu	on analyses were use o had not vaped in th ons per individual an uitting plans (to retair	d to assess the re past 6 months at d statistically controlling in the analysis those
$^{\circ}$ Association is significantly different from analogous association among those planning to quit within the next 6 months ($P = <0.01$ for interaction). ⁶ Association is significantly different from analogous association among those planning to quit within the next 6 months ($P = <0.05$ for interaction). ⁸ Model convergence not achieved for those with unknown plans to quit (for any model).	^c Those who were planning to quit wi baseline; the numerator of the outco ^d Analyses were adjusted for country, time in samule and wave: all covaria	thin the next 6 months at baselin me for this group reflects maintai , biological sex, race/ethnicity (W tes wore assessed at haselinar (GF	ining quit hite, non-	s those who were pla plans. The numerator White), age group (18 were fitted specifyin	inning to quit within one rs of the outcome for the 3-24, 25-39, 40-54, 55+ othe unstructured covari	month at baseline and those v other baseline quit intentions · years), socioeconomic status	who were s group re (SES) (lov	planning to quit beth sflect changing to hav <i>w</i> , moderate, high), ci	veen 1 and 6 months at 'e quit plans. garettes smoked per day,
	*Association is significantly different Association is significantly different fAssociation is significantly different 8Model convergence not achieved fo	from analogous association amor from analogous association amor from analogous association amor or those with unknown plans to qi	ng those p ng those p ng those p uit (for any	vere nuce specifyin lanning to quit within <i>y</i> model).	Is the next 6 months ($P = \cdot$) the next 6 months ($P = \cdot$)	40.05 for interaction).			

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disaggregating the groups that did not have plans to quit in the next 6 months, we found that those who were least definitive in their plans (i.e. those who were planning to quit beyond 6 months or did not know) experienced the greatest relative increases in smoking cessation when they took up daily vaping. Although additional studies are needed to understand our findings more fully, smokers without definitive plans to quit tended to want to quit, but had low quitting selfefficacy before vaping uptake; therefore, it is plausible that uptake of vaping among this group may help them to overcome expectancies that living without cigarettes is too challenging [23].

Although studies on use of nicotine replacement therapy (NRT) and smoking cessation typically focus on cessation aid effectiveness when used during a quit attempt [13, 24, 25] (see also Figure 1), our findings suggest that evaluation of use of NVPs and smoking cessation should also include those not planning to guit. With regard to our findings among smokers who were initially planning to quit, it is possible that those who did not take up vaping may have been more likely to use NRT or other cessation aids than their counterparts who took up daily vaping. We found that those planning to guit smoking who did not take up vaping experienced a 17% smoking cessation rate. whereas only 7% of those not planning to guit who did not take up vaping went on to guit smoking. That is, the difference in smoking cessation rates between those who do and do not plan to guit smoking appears to be driven by differences in cessation rates among those who do not take up vaping. In other words, daily vaping uptake is associated with similar smoking cessation rates for those who do and do not plan to quit.

Limitations

In this population-based study, we were not able to ascertain the temporal ordering of vaping uptake and change in plans to quit following baseline assessment, and our findings should not be interpreted as indicating causal associations. We were also not able to ascertain a measure of inter-wave NRT/other cessation aid uptake consistent with our measure of inter-wave vaping uptake, and therefore, we could not evaluate whether/to what extent other aid use may confound our findings for vaping use. Sample sizes were also generally small, owing in part to the follow-up timeframe eligibility requirement, which was necessary to ascertain inter-wave vaping uptake when including non-recent vapers in the sample given the timeframes for which respondents were asked about their prior vaping experiences; however, this reduced our statistical power. Last, many models to test for country-by-vaping uptake interactions failed to converge, and others were not statistically testable owing to some very small cell sizes. Future research with additional waves of data will increase statistical power to test for country interactions and to detect additional significant associations if they exist and will also allow for the assessment of smoking cessation in the longer term. Future research can also evaluate type of vaping products used and amount of nicotine consumed, which may be particularly informative for NVP regulation.

CONCLUSIONS

For daily cigarette smokers who are not planning to quit smoking, uptake of daily vaping is associated with higher smoking cessation rates compared with no uptake of vaping. Smokers not planning to quit tend to be older, heavier smokers of lower SES. Studies focused exclusively on smokers who are already planning or attempting to quit may underestimate a potential benefit of daily vaping for smoking cessation at the population level, including potential to reduce disparities in smoking rates. Last, only daily vaping was associated with smoking cessation, suggesting that vaping company marketing practices, vaping policies/restrictions and public health education campaigns could be undertaken to support smokers' use of vaping products on a daily basis.

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DECLARATION OF INTERESTS

K.M.C. has in the past and continues to serve as a paid expert witness in litigation filed against cigarette manufacturers. D.H. has served as a paid expert witness on behalf of governments and public health authorities in legal challenges against tobacco and vaping companies. G.T.F. has served as an expert witness or consultant for governments defending their country's policies or regulations in litigation. G.T.F. and S.G. served as paid expert consultants to the Ministry of Health of Singapore in reviewing the evidence on plain/standardized packaging. The views expressed in this article are those of the authors and not necessarily those of the National Health and Medical Research Council or the other funding agencies. None of the other authors has any conflict of interest to declare.

AUTHOR CONTRIBUTIONS

Karin A. Kasza: Conceptualization; formal analysis; investigation; methodology. David Hammond: Investigation; methodology. Shannon
Gravely: Investigation; methodology. Richard J. O'Connor: Investigation; methodology. Gang Meng: Investigation; methodology.
Katherine East: Investigation; methodology. Ron Borland: Funding acquisition; investigation; methodology. K. Michael Cummings: Funding acquisition; investigation; methodology. Geoffrey T. Fong:
Funding acquisition; investigation; methodology. Andrew Hyland: Conceptualization; funding acquisition; investigation; investigation; methodology.

ETHICS STATEMENT

Study questionnaires and materials were reviewed and provided clearance by the Research Ethics Committees at the following institutions: University of Waterloo (Canada, ORE20803/30570, ORE21609/ 30878), King's College London, UK (RESCM-17/18-2240), Cancer Council Victoria, Australia (HREC1603), University of Queensland,

Australia (2016000330/HREC1603) and Medical University of South Carolina (waived because of minimal risk). All participants provided consent to participate.

DATA AVAILABILITY STATEMENT

In each country participating in the International Tobacco Control Policy Evaluation (ITC) Project, the data are jointly owned by the lead researcher(s) in that country and the ITC Project at the University of Waterloo. Data from the ITC Project are available to approved researchers 2 years after the date of issuance of cleaned data sets by the ITC Data Management Centre. Researchers interested in using ITC data are required to apply for approval by submitting an International Tobacco Control Data Repository (ITCDR) request application and subsequently to sign an ITCDR Data Usage Agreement. The criteria for data usage approval and the contents of the Data Usage Agreement are described online (http://www.itcproject.org).

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SUPPORTING INFORMATION

Additional supporting information can be found online in the Supporting Information section at the end of this article.

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