

CURRENT ISSUES WITH ELECTRONIC CIGARETTES AND VAPING

MATTHEW J CAMPEN, PHD, MSPH

REGENTS' PROFESSOR OF PHARMACEUTICAL SCIENCES

COLLEGE OF PHARMACY, UNIVERSITY OF NEW MEXICO HEALTH SCIENCES CENTER



HEALTH AND SOCIOLOGICAL ISSUES

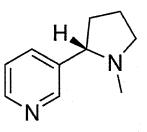
- FDA REGULATION
 - MARKETING
- MEDICAL USES
- HEALTH EFFECTS
 - · ADDICTION
 - CARDIOPULMONARY
 - NEUROLOGICAL





REGULATORY FRAMEWORK

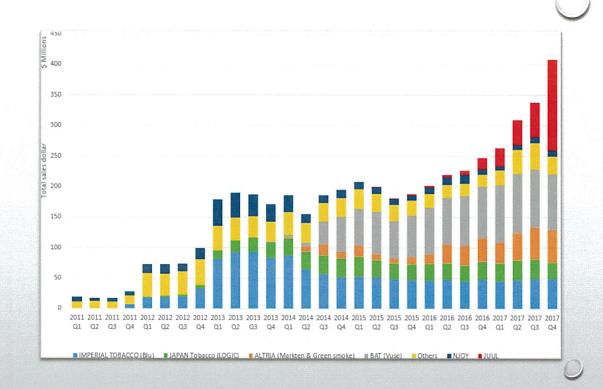
- ELECTRONIC NICOTINE DELIVERY SYSTEMS (ENDS) OR E-CIGS ARE A COMPLICATED ENTITY
- NICOTINE IS AMONG THE MOST ADDICTIVE CHEMICALS KNOWN
 - NICOTINE IS A MILD STIMULANT
- IS NICOTINE A MEDICINE?
 - YES, BUT ONLY TO TREAT ADDICTION TO NICOTINE IN TOBACCO PRODUCTS
- WHO REGULATES AN ADDICTIVE STIMULANT THAT IS OCCASIONALLY A MEDICINE?
 - FDA, VIA THE FOOD, DRUG AND COSMETICS ACT.
 - "WARNING: THIS PRODUCT CONTAINS NICOTINE. NICOTINE IS AN ADDICTIVE CHEMICAL."





VAPING TRENDS

- OVER THE PAST DECADE, ELECTRONIC NICOTINE DELIVERY SYSTEMS (ENDS)
 HAVE DRAMATICALLY INCREASED IN TERMS OF USAGE AND SALES
- BIG TOBACCO LED THE WAY, BUT INDEPENDENT COMPANIES (JUUL) ARE EMERGING
- LOCAL DISPENSARIES ARE ALSO
 EMERGING, BUT HARDER TO QUANTIFY



POTENTIAL CARDIOVASCULAR EFFECTS OF E-CIGARETTES

Electronic Cigarettes Increased Cardiovascular Risk

Mechanisms

Sympathetic Nerve Activation

↑ Heart rate^{14,16}

Abnormal HRV^{14,15}

↑ Blood pressure^{14,16}

Oxidative Stress/Inflammation

Endothelial damage/dysfunction

Platelet activation

- ↑ LDL Oxidizability¹⁵
- ↑ Plasma Myeloperoxidase³⁴
- ↓ Vitamin E levels³³
- ↑ 8-iso-prostaglandin F2α³³
- ↓ Nitric oxide bioavailability³³
- ↑ Inflammation on FDG-PET38

Biomarkers

- ↓ FMD³³
- ↑ PWV^{34,46}
- ↑ EPCs⁴⁴

↑ Platelet aggregation⁴⁷

Potential Clinical Sequelae

- Arrhythmias
 - Atrial
 - Ventricular
- Inflammatory Atherosclerosis
- Vasospasm
- Plaque Instability
- Acute Ischemia
- Platelet activation

- Inflammatory Atherosclerosis
- · Plaque Instability
- Acute Ischemia
- Platelet Activation

- · Inflammatory Atherosclerosis
- Plaque Instability
- Acute Ischemia
- **Acute Thrombosis** · Acute Ischemia



WHO SHOULD QUIT SMOKING?

WHO IS ACTUALLY PRESSURED TO QUIT SMOKING?

- PREGNANT MOMS
 - NICOTINE REPLACEMENT (PATCH OR ECIG) IS OFTEN RELIED UPON
 - NICOTINE IS PREGNANCY CATEGORY D = THERE IS EVIDENCE OF HUMAN FETAL RISK, BUT THE POTENTIAL BENEFITS FROM THE USE OF THE DRUG IN PREGNANT WOMEN MAY BE ACCEPTABLE DESPITE ITS POTENTIAL RISKS.
 - CHANTIX, OTHER PHARMACOLOGICAL TOOLS FOR SMOKING CESSATION



WHO SHOULD QUIT SMOKING?

WHO IS ACTUALLY PRESSURED TO QUIT SMOKING?

- HEART FAILURE PATIENTS
 - TYPICALLY FIRST NOTED IN PATIENTS WITH HIGH BLOOD PRESSURE
 - NICOTINE MAY INCREASED BLOOD PRESSURE AND WORSEN OTHER CARDIOVASCULAR SYMPTOMS

E-CIGARETTES MAY HELP PEOPLE QUIT TOBACCO USE

Variable	1 Wk after	Quit Date	Mean Difference (95% CI)	4 Wk after Quit Date		Mean Difference (95% CI)
	E-Cigarettes (N=158)	Nicotine Replacement (N=131)		E-Cigarettes (N = 186)	Nicotine Replacement (N=132)	
Score for frequency of urge	2.5±1.1	2.8±0.9	-0.4 (-0.6 to -0.1)	1.9±0.9	2.2±0.8	-0.3 (-0.5 to -0.1)
Score for strength of urge	2.7±1.1	3.2±1.0	-0.5 (-0.7 to -0.2)	2.1±1.1	2.4±1.0	-0.3 (-0.6 to -0.1)
Composite urge score	2.6±1.0	3.0±0.9	-0.4 (-0.6 to -0.2)	2.0±1.0	2.3±0.9	-0.3 (-0.5 to -0.1)

^{*} Plus-minus values are means ±SD. Scores for frequency of urge ranged from 1 (not at all) to 6 (all the time). Scores for strength of urge ranged from 1 (no urges) to 6 (extremely strong). The composite score (range, 1 to 6, with higher scores indicating more severe urges) is an average of the frequency and strength scores.

N Engl J Med 2019; 380:629-637









E-CIGARETTES MAY HELP PEOPLE QUIT TOBACCO USE

CORRESPONDENCE

E-Cigarettes versus Nicotine-Replacement Therapy for Smoking Cessation

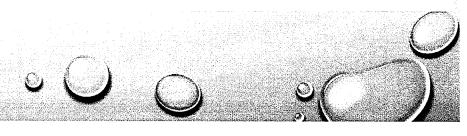
Normalizing vaping and modeling addictive behavior might escalate the extent of the epidemic among youth.² Nicotine adversely affects brain development, and as noted in the editorial by Drazen et al.⁴ in the same issue of the *Journal*, teens who vape are more likely to start smoking than those who do not.⁵ Caution is advised when choosing e-cigarettes, with consideration of their comparable effectiveness to FDA-approved medications but uncertain long-term harms.

Yuqing Zhang, M.D., Ph.D.

Guang'anmen Hospital China Academy of Chinese Medical Sciences Beijing, China

Dona Upson, M.D.

University of New Mexico School of Medicine Albuquerque, NM dona.upson@va.gov





VAPORS CONTAIN MORE THAN JUST NICOTINE

- E-LIQUID IS A FORMULATION OF PHOSPHOLIPIDS AND/OR GLYCEROL
- SEVERAL WELL-KNOWN CARCINOGENS ARE GENERATED IN THE PROCESS OF

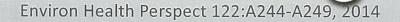
VAPORIZATION

- OTHER CONTAMINAL
- ADDITIONALLY, INDEI
 E-LIQUIDS

Toxic compound	Tobacco cigarette (µg in mainstream smoke)	E-cigarette (µg per 15 puffs*)	Average ratio (conventional vs electronic cigarette)	
ormaldehyde 1.6-52		0.20-5.61	9	
Acetaldehyde	52-140	0.11-1.36	450	
Acrolein	2.4-62	0.07-4.19	15	
Toluene	8.3-70	0.02-0.63	120	
NNN++	0.005-0.19	0.00008-0.00043	380	
INK** 0.012-0.11		0.00011-0.00283	40	

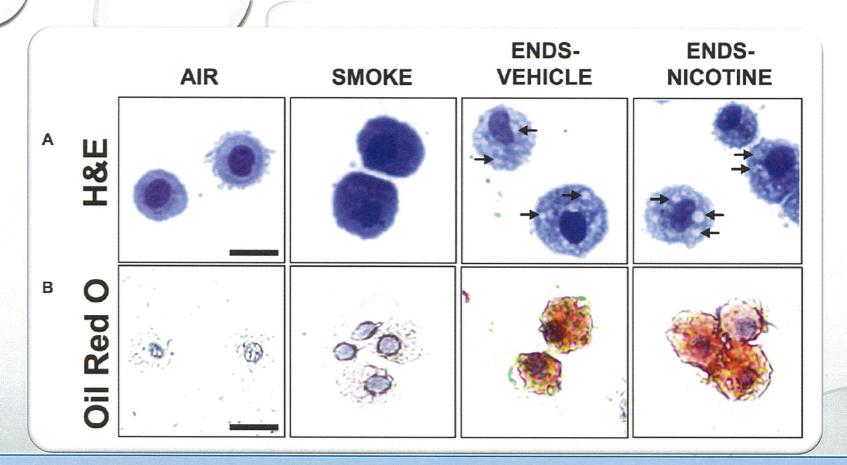
The authors assumed smokers of e-cigarettes would take an average of 15 puffs per vaping session, corresponding to smoking one tobacco cigarette.

^{**} Tobacco-specific nitrosamine, a carcinogenic compound that originates in the curing and processing of tobacco. Adapted from Goniewicz et al. (2014)⁴

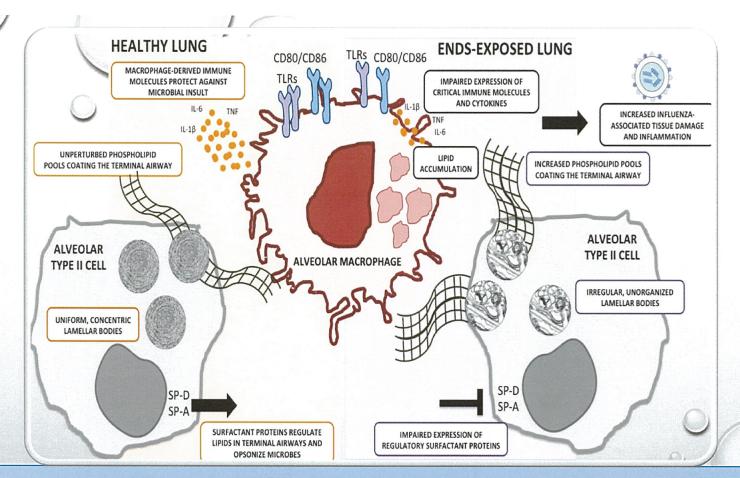


ONGOING EFFORTS TO STUDY E-CIGARETTES AT UNM ARE LIMITED

- MARY WALKER, PH.D., COLLEGE OF PHARMACY, MAINLY EXAMINING CIGARETTE SMOKING
- MIND NETWORK DOES LIMITED STUDIES OF NEUROLOGICAL EFFECTS
- CAMPEN WORKS ON A SUBCONTRACT
 WITH BAYLOR COLLEAGUES STUDYING THE
 PULMONARY AND VASCULAR TOXICITY
- UNM HAS LIMITED RESOURCES FOR EXPOSURE AND CHEMICAL CHARACTERIZATION OF POTENTIAL CONTAMINANTS
- STRONG FUNDING IN OTHER AREAS
 (OPIATES, ENVIRONMENTAL TOXICANTS)
 PULLS MANY INVESTIGATORS AWAY FROM
 THIS FIELD



LONG-TERM E-CIG EXPOSURE ALTERS LIPID HANDLING IN LUNG IMMUNE CELLS (J CLIN INVEST, IN PRESS, 2019)



THIS TOXIC EFFECT WAS DRIVEN BY CHRONIC INHALATION OF THE E-LIQUID, NOT THE NICOTINE



NEXT BIG THINGS: VAPORIZED MARIJUANA

- VAPING OF MARIJUANA AND ACTIVE COMPONENTS (CANNABINOIDS) IS ALSO A GROWING TREND
- AS WITH TOBACCO, VAPING LIKELY DELIVERS FEWER TOXIC COMPONENTS THAN THE COMBUSTED FORM.
- POPULARITY OF THE DEVICES AND FORMULATIONS OF THC MAY BECOME PROBLEMATIC
 - EASE FOR ABUSE
 - TARGETING MINORS