

Next Generation Products Science & Technology

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Important notice (continued)

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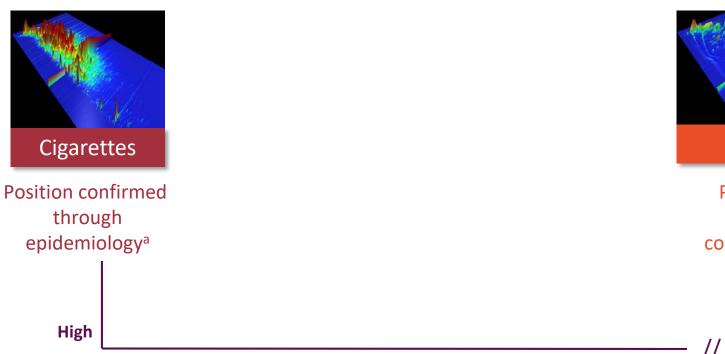








Increasing acknowledgement of the risk continuum model



THP

Position
to be confirmed*
confirmed*
epidemiology

Position proposed
By Public Health
Authorities*

Low

*Clinical data suggests reduced exposure^b
*UK pubic health suggests reduced risk^d

- a. Doll & Hill (1954) BMJ 1954: 1451-1455
- b. Fearon et al 2017 Poster 113 GFN Conference http://bit.ly/2xRPVvD
- c. Lee (2013) *Harm Reduct J* **10:** 36, doi: 10.1186/1477-7517-10-36
- d. PHE Report 2015 bit.ly/1lyp57N; RCP Report 2016 http://bit.ly/244lizV; CRUK 2016 http://bit.ly/2yVZlq0

These modellings do not necessarily mean that our products are less harmful than cigarettes



Increasing acknowledgement of the risk continuum model

Our portfolio approach supports the risk continuum model



These modellings do not necessarily mean that our products are less harmful than cigarettes



A three-step scientific journey to assessing the risk profile of Next Generation Products^a

01 EMISSIONS

- Science showing no combustion
- > Untargeted emissions
- Targeted emissions
- > Environmental emissions

02 EXPOSURE

- > Puffing behaviour
- Average daily consumption
- > Clinical PK
- > Clinical BoE

03 RISK

- > Clinical BoBE
- > Risk perception
- > Post-market surveillance

Reduced toxicity in lab models

in vitro reg tox

in vitro disease models

in vitro systems science

a. Murphy J et al (2017) Reg Tox Pharm doi.org/10.1016/j.yrtph.2017.09.008





Driving consumer conversion

Addressing safety, quality, potential reduced risk nature and consumer satisfaction









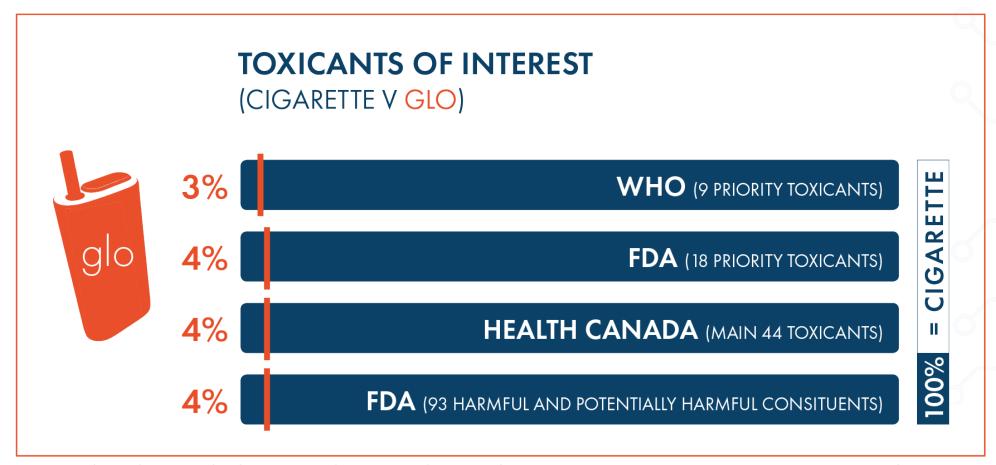
The glo development journey





Targeted emissions: glo

In laboratory tests, **glo** has substantially reduced levels of targeted toxicants in comparison to a scientific reference cigarette, 3R4F



Comparison of smoke from a scientific reference cigarette (approx. 9 mg tar) and vapour from glo. This quality does not necessarily mean this product is less harmful than other tobacco products

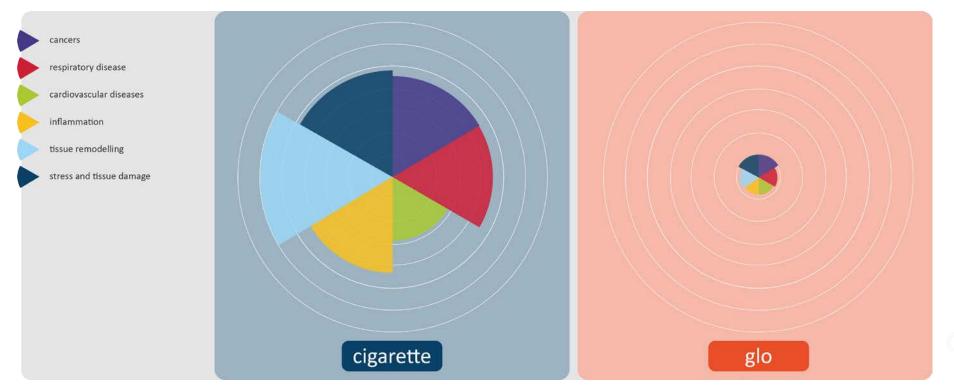


Systems biology: glo

In laboratory tests, **glo** promotes substantially less disease relevant gene changes in comparison to a scientific reference cigarette, 3R4F

In lab based tests, glo[™] promotes substantially less disease relevant gene responses in comparison to a cigarette^{a,b}

- > 3-D cellular system (Mucilair[™]) exposed to glo[™] heated tobacco vapour and cigarette smoke
- > Ingenuity Pathway Analysis^c of >22,000 gene responses grouped into disease relevant categories

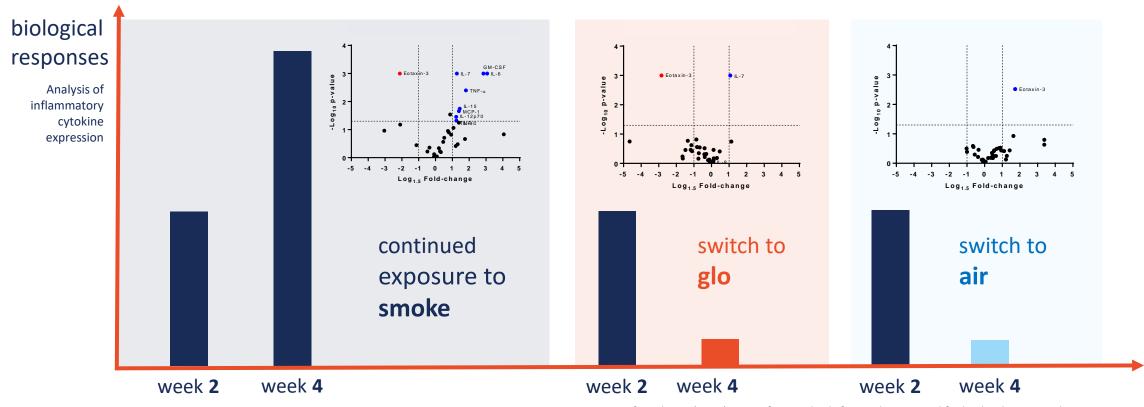


^aMethods adapted from Banerjee et al (2017) Appl in vitro Toxicol. 3(1) 84-98; doi:10.1089/aivt.2016.0024; ^bglo and a scientific reference cigarette (3R4F) puffed on machines using the Health Canada Intense (HCI) regime: The contents of glo vapour differ significantly from cigarette smoke, however, these qualities and the above responses do not necessarily mean this product produces less adverse health effects than other tobacco products; ^cIngenuity Pathway Analysis (IPA) has broadly been adopted by the life science research community and is widely cited for the analysis, integration, and interpretation of data derived from 'omics experiments



glo laboratory switching studies in support of Institute of Medicine 'gold standard'a

Following two week exposure to smoke, there was no statistically significant difference in cells which were exposed to **glo** vapour or air^b



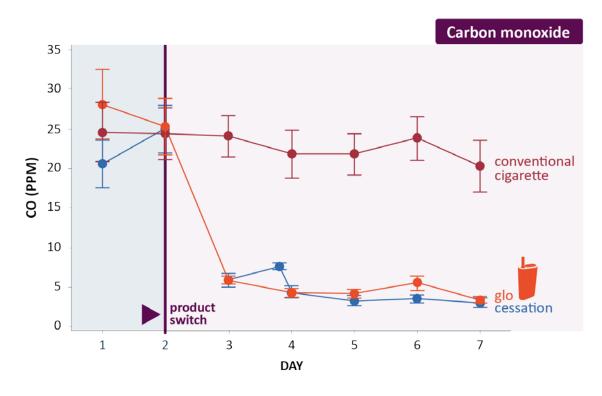
These results do not necessarily mean this product is less harmful than other tobacco products

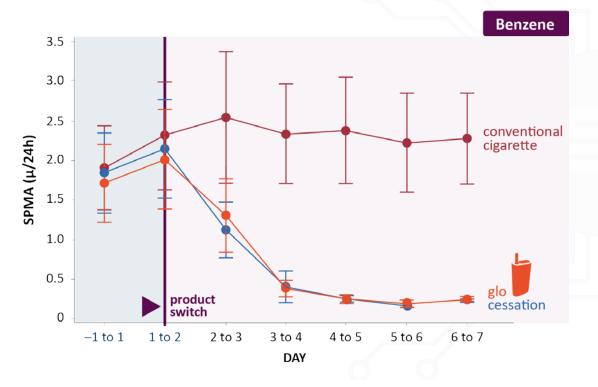
^aInstitute of Medicine (2012) Scientific Standards for studies on modified risk tobacco products ^bThe 3-D cell construct Mucilair™ was used in this study



glo clinical switching studies in support of Institute of Medicine 'gold standard'

No statistically significant difference in BoEs between smokers who switch to **glo** and those who quit^{b,c,d}



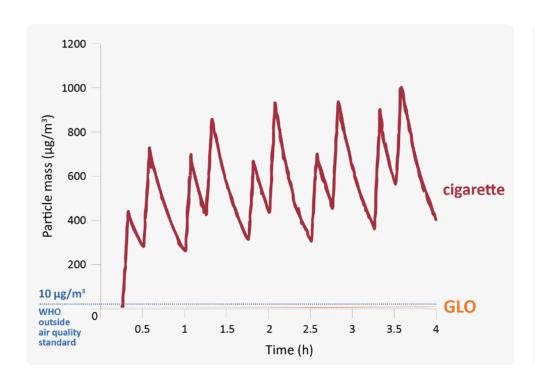


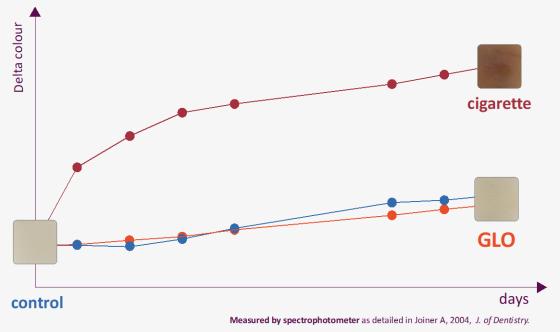
^aInstitute of Medicine (2012) Scientific Standards for studies on modified risk tobacco products ^bThese results do not necessarily mean this product is less harmful than other tobacco products ^cGale *et al* (2017) *BMC Public Health* doi.org/10.1186/s12889-017-4678-9 ^dFearon *et al* (2017) Poster 113 GFN Conference http://bit.ly/2xRPVvD



Science update: glo

Science based studies to assess personal hygiene and social consideration of glo^a





Social consideration: 'cleaner air' when using glo relative to smoking^b

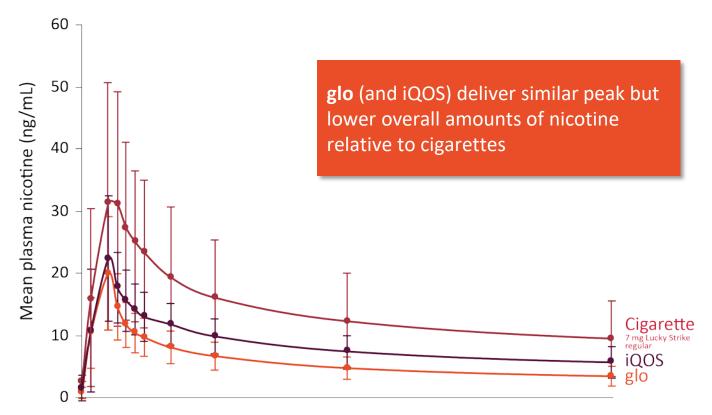
Personal hygiene: teeth staining is reduced when exposed to glo vapour in comparison to cigarette smoke and parity with untreated control

^aThese results do not necessarily mean this product is less harmful than other tobacco products ^bForster *et al* (2017) *Reg Tox Pharm*, in press



Pharmacokinetic data: glo

Uptake of nicotine amongst Japanese consumers



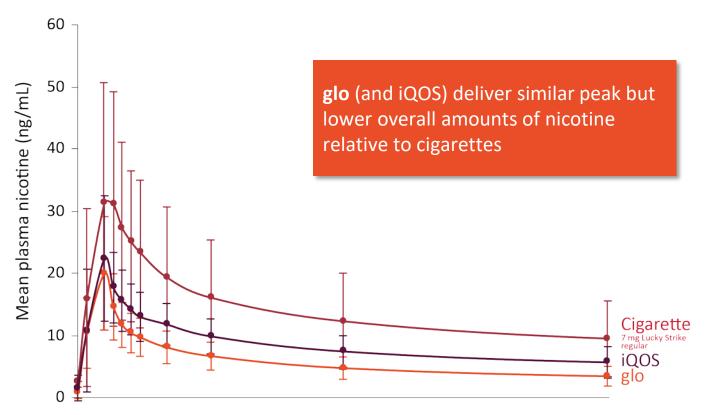
Time relative to first puff

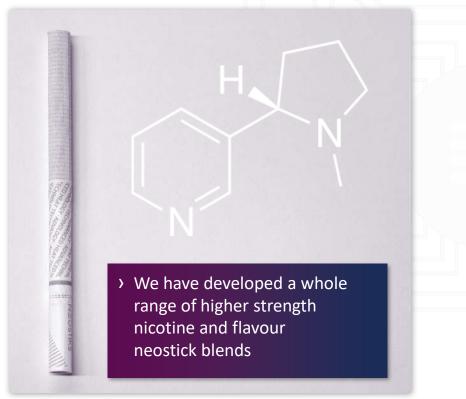
Gale et al (2017) BMC Public Health doi.org/10.1186/s12889-017-4678-9
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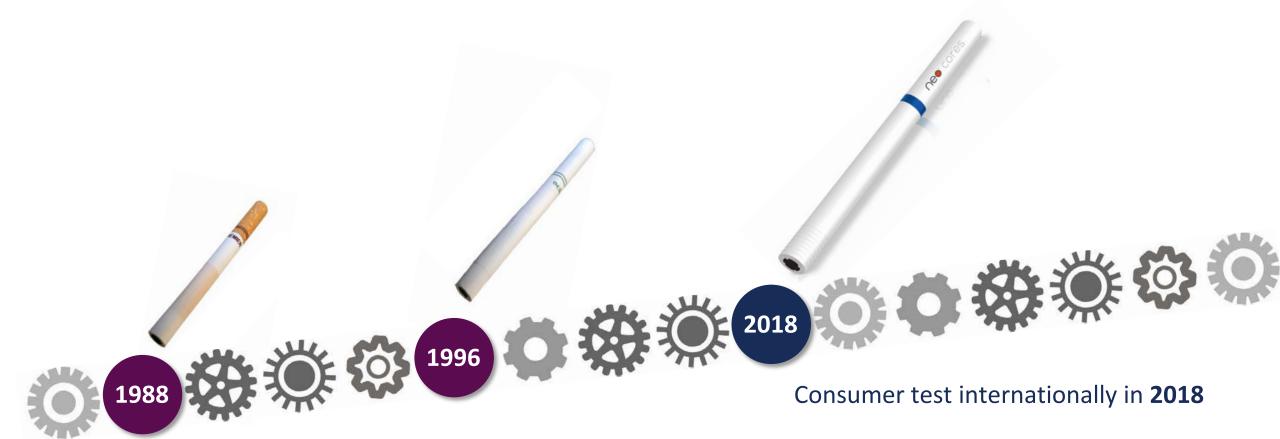






Third generation carbon tip technology









Strong performance in taste, satisfaction and usability

- More impact
- Improved taste of neocores vs. previous carbon tip heated products
- Easier to light up

- > Reduced toxicants
- > Reduced odour *
- > Reduced ash

Comparison of smoke from an industry standard reference cigarette (approx. 9 mg tar) and vapour from Neocores CTHP, in terms of the 9 harmful components the World Health Organisation recommends to reduce in cigarette smoke

This quality does not necessarily mean this product is less harmful than other tobacco products *Compared to a conventional cigarette when smoked





Extensive scientific evaluation and publications on the carbon tip platform **over three decades**

- Targeted emissions
- Side-stream emissions
- In vitro and in vivo regulatory toxicology
- Topography
- Urine mutagenicity
- Biomarker studies







Hybrid Gen 1 science and technology



^{*}This is average reduction in toxicants as defined by WHO Tobreg in the iFuse aerosol relative to 3R4F scientific reference cigarette smoke





Hybrid Gen 2 science and technology



This quality does not necessarily mean this product is less harmful than other tobacco products







A novel oral smokeless product addressing currently unmet consumer needs

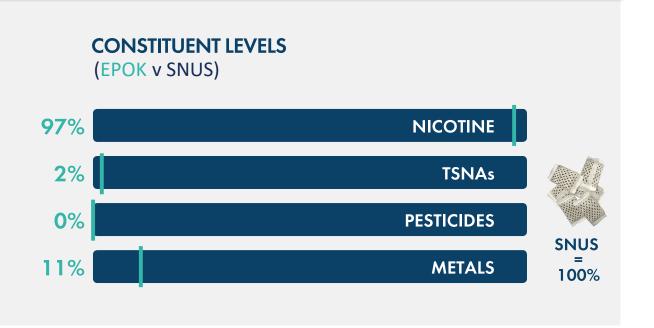
An extended portfolio to incorporate new to world smokeless products

Snus: Leading portfolios of smokeless tobacco products in US and Northern Europe

EPOK: new to world oral smokeless product

- Cellulose based low tobacco inclusion nicotine product containing flavours
- Delivers same consumer satisfaction as Snus, with improved chemical profile
- > Less visible oral hygiene impact*

Introducing a tobacco free variant containing nicotine and flavours in the future



This quality does not necessarily mean this product is less harmful than tobacco products



^{*}Compared to use of traditional snus products





Game changing technology: Dynamic Vapour Technology

Many advantages vs. coil/wire & wick systems

- > 46x larger surface area
- > 3x aerosol particle size
- > Advanced temperature control
- > Dynamic wicking
- > Improved sensory performance
- > Improved consumer satisfaction
- Less e-liquid nicotine required





	Competitor Coil/Wire & Wick Closed Systems*	Raptor
Performance		
Aerosol Particle Size (nm)	450 to 550	Up to 2000
Purity		
Formaldehyde (ug/puff)	0.8 to 1.2	0.1

^{*}Logic Pro and MESH

This quality does not necessarily mean this product is less harmful than other tobacco products



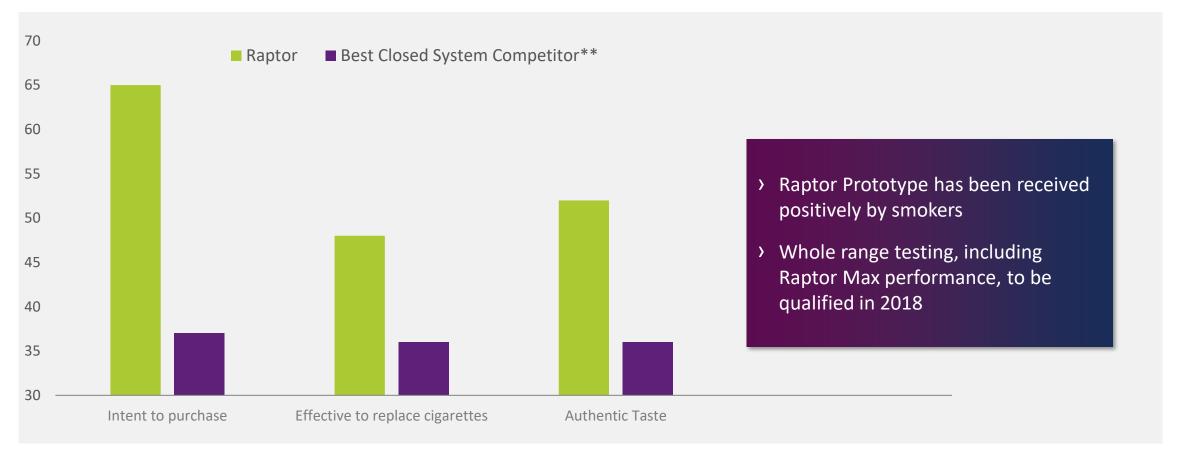
Developing and tuning a new to the world technology





Raptor: A new level of performance for smokers

Total Offer Evaluation with German smokers (Raptor 3mg/ml eLiquid)*



^{*40} to 50 smokers in each panel

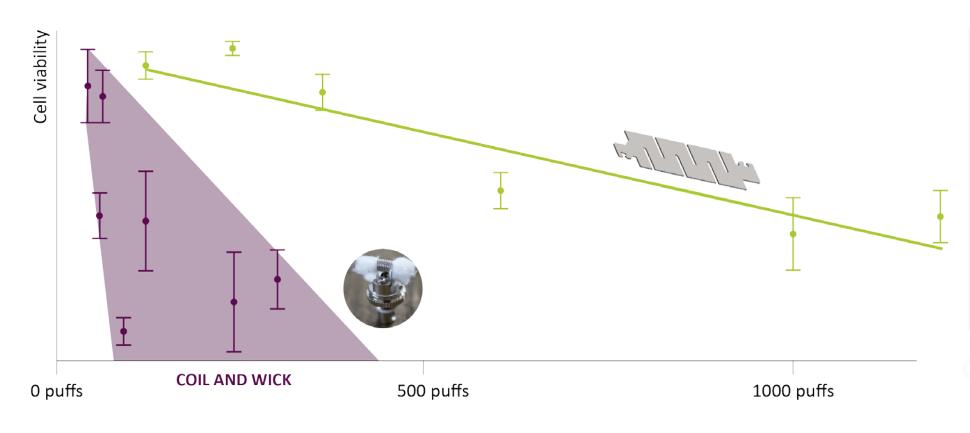
This quality does not necessarily mean this product is less harmful than other tobacco products



^{**}Logic Pro

Raptor: A new level of performance in toxicity testing

Pre-clinical toxicological assessment shows potential of **DV Tech** versus 'coil and wick' based e-cigarettes



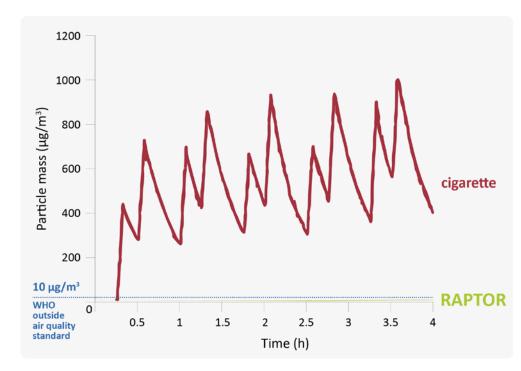
'Coil and wick' based vapour products have substantially reduced toxicological responses relative to cigarettes. However, Raptor, incorporating 'DV Tech' shows even greater reductions, even under extreme puffing conditions

This quality does not necessarily mean this product is less harmful than other tobacco products

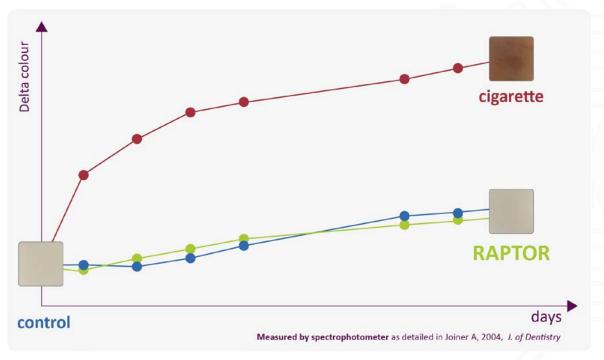


Science update: Raptor

Science based studies show improved personal hygiene and social consideration of Raptor^a



Social consideration: 'cleaner air' when using Raptor relative to smoking^b



Personal hygiene: teeth staining is reduced when exposed to Raptor vapour in comparison to cigarette smoke and parity with untreated control

a. These results do not necessarily mean this product is less harmful than other tobacco products



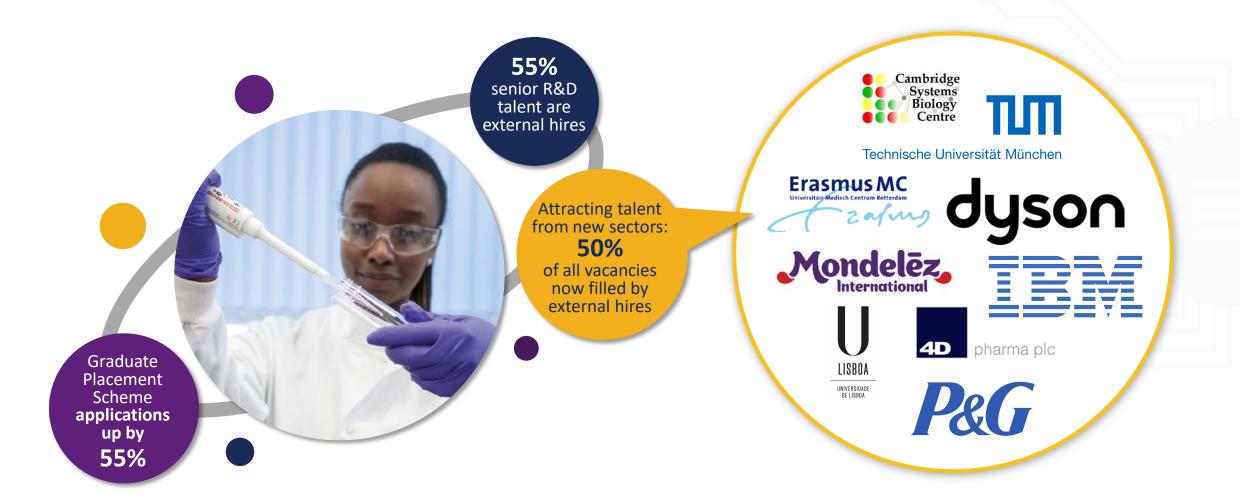








Transformation of our R&D through talent acquisition





BAT/RAI: creation of the industry's leading R&D





- Combining the capabilities of both organisations enhances our global R&D strength
- Combined R&D by numbers:
 - > 1500 people across science, technology and innovation
 - > Total R&D investment over last six years: \$2bn
 - Combined patent portfolio over last three years:
 more than 3300 filed; more than 1600 granted
 - Combined publications in last three years: 109
 - Innovation, product development and compliance expertise in all regions across the globe



R&D OF THE FUTURE NOW







Next Generation Products

Science & Technology







