



Medical Detection Dogs

Dr Claire Guest

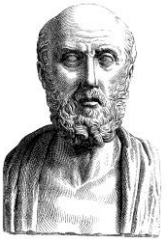
CEO and Chief Scientific Officer, Medical Detection Dogs

**Medical Detection Dogs
Founded in 2008**

Patron: HRH The Duchess of Cornwall

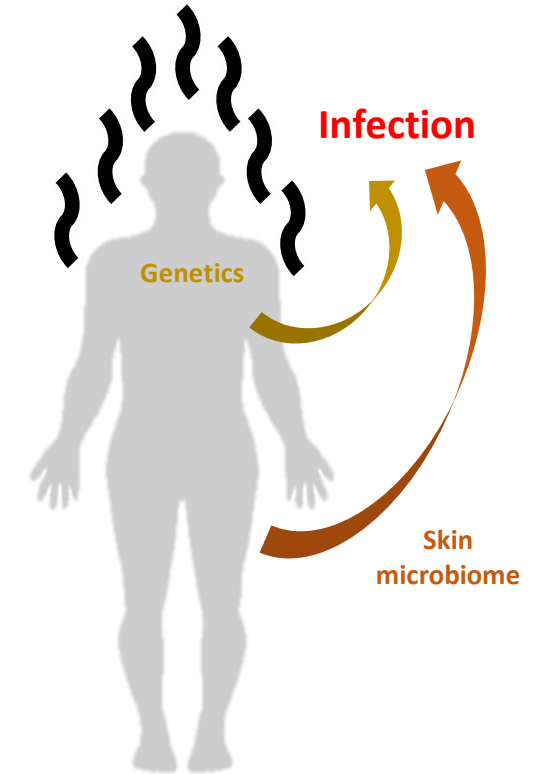
Volatile Organic Compounds: biomarkers of disease

- Humans emit a 500-600 VOCs
- Varies with age, diet, sex, physiological status, genetic background



Physicians have used smell to diagnose patients for hundreds of years: Hippocrates (400 BC) recognized the diagnostic usefulness of body odours and reported on several disease-specific odours

- Infection associated VOCs found in body odour (sweat), breath, urine, faeces



Typhoid



TB



Yellow fever



Medical Detection Dogs approach: dogs detecting human disease

Diagnosis
Bio-Detection Dogs



Assistance
Medical Alert Assistance Dogs



Measure of a Diagnostic

- Accuracy
- Affordability
- Accessibility (scalability)



Application and Development

1 to 1

Medical Alert Assistance Dogs working with one person who has complex health condition(s) such as Type 1 Diabetes, POTS, Addison's disease, Mast Cell Activation Syndrome or severe allergies. It is likely that we can train dogs to support a wide range of other conditions.

1 to Some

Medical Detection Dogs in a clinical setting working on samples rather than individuals including:

Clinical: Dogs in a Hospital, Rehabilitation or Care Home supporting clinicians by detecting for specific odours from samples, e.g. the presence of bacteria which cause UTIs but also providing 1 to 1 animal assisted therapy, and

Research: intended to help clinicians with faster, non-invasive means of diagnosis including Cancers, Neurological diseases, bacteria but not yet to scale.

1 to Many

Medical Detection Dogs working to help a wider group of people. This could be through the development of research projects which lead to dogs teaching an artificial intelligence device to learn the odour pattern, the 'tune' rather than the notes leading to a reliable electronic nose that can achieve the scale necessary to screen populations e.g. for cancer, or through dogs screening large numbers of people as they pass through ports of entry, e.g. for malaria.

MDD Method

- Dogs are selected based on internal MDD assessment of performance and suitability for the task
- Dogs are trained to identify the disease odour following a positive reinforcement training process
- Dogs are trained to present a 'yes' or 'no' behaviour to indicate if a sample is positive or negative
- Samples are presented from both target disease and controls and are tested using double blind protocol
- Rigorous data collection and performance analysis

Sample presentation

- Integrity of samples
- Carefully annotated samples
- Strict SOPs for handling and storage
- Dogs learn VOC pattern 'tune' of true odour through complex discrimination
- Requirement of multiple new samples without confounders
- Avoid repeated sample exposure



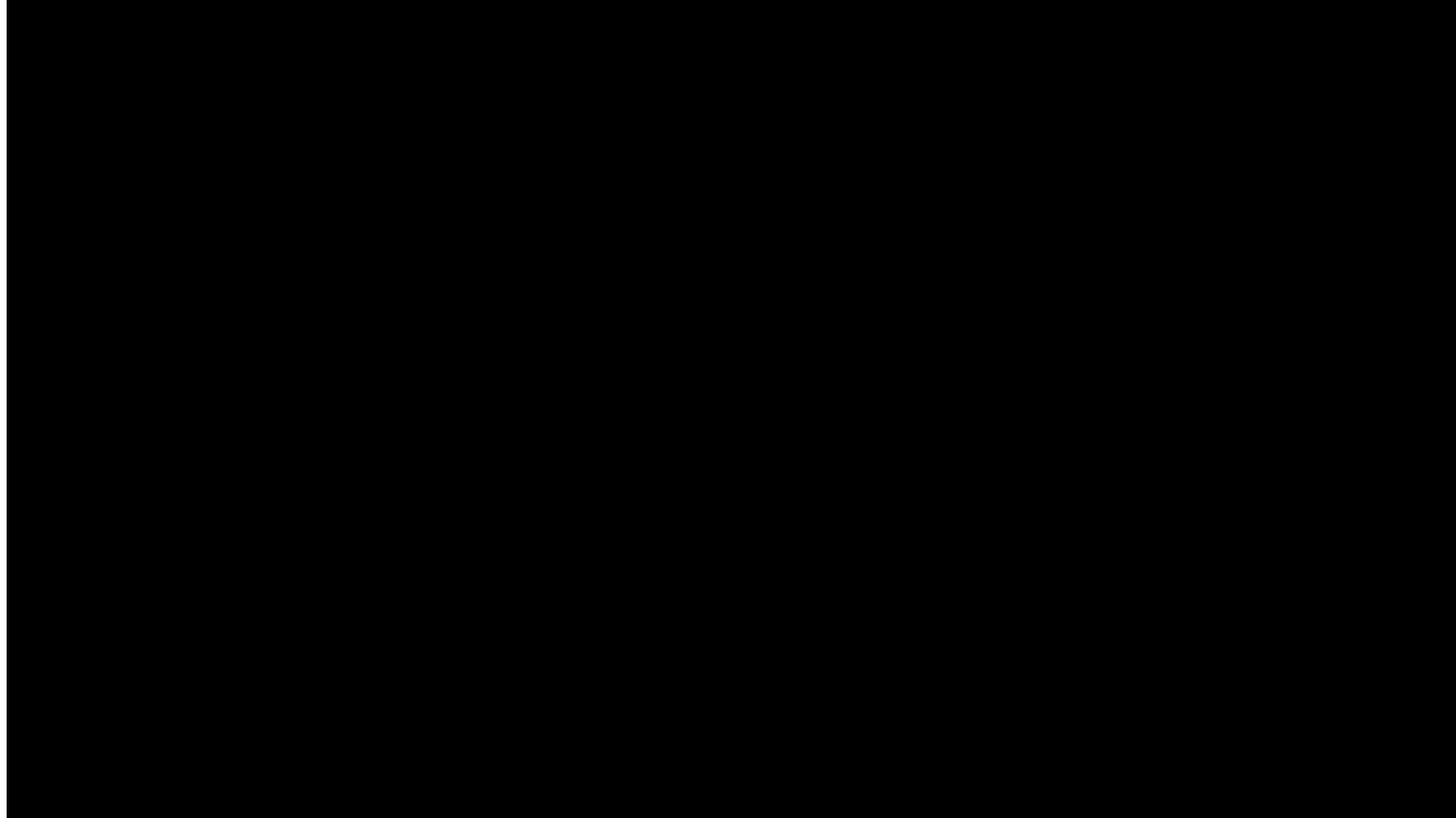
Target Run



Blank Run



Client Partnership: Claire and Magic: Type 1 diabetes



Rooney, N.J., Morant, S. and Guest, C., (8/2013). "Investigation into the value of trained glycaemia alert dogs to clients with type I diabetes". *PLoS one*, 8(8), p.e69921.

Wilson, C., Morant, S., Kane, S., Pesterfield, C., Guest, C., N.J. Rooney., (3/2019) "An Owner-Independent Investigation of Diabetes Alert Dog Performance". *Frontiers in Veterinary Science* 6:91. doi: 10.3389/fvets.2019.00091

Rooney, N.J., Guest, C.M., Swanson, L.C. and Morant, S.V., (1/2019). "How effective are trained dogs at alerting their owners to changes in blood glycaemic levels?: Variations in performance of glycaemia alert dogs." *PLoS ONE*, 14(1), p.e0210092.

Data-Evaluate

MDD Biodeflection data entry (B:\Projects\Cancer projects\Urological Cancer\Training and Samples databases and Apps\Biodeflection dat... Claire Guest


Session
Handler: RH
Helper: SW
Temperature: 19
Humidity: 54
System: Carousel
Positions: 4
Blinding: Trial

Sample set
Select file
Sample set 03_12_18.xlsx
Set No.: 1
Sequence No.: 1 [Reset]
Re-order samples

Run
Dog: Florin
Run No.: 1
Session No.: 120
Start video
Reveal results
Save Run
View database

| Position | Pass 1 | Pass 2 | Pass 3 | Pass 4 |
|------------|---|--|--|--|
| Position 1 | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input checked="" type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA |
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| Position 3 | <input checked="" type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA |
| Position 4 | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input checked="" type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA | <input type="radio"/> IND <input type="radio"/> HES <input type="radio"/> INT <input type="radio"/> NI <input type="radio"/> NS <input type="radio"/> CSA |

Overall result



OK

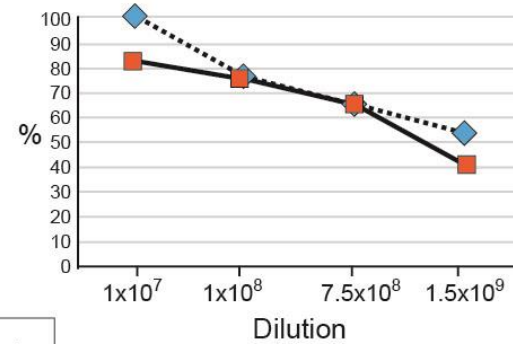
No decision
Reveal IDs

Evaluate: olfactory thresholds

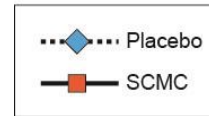
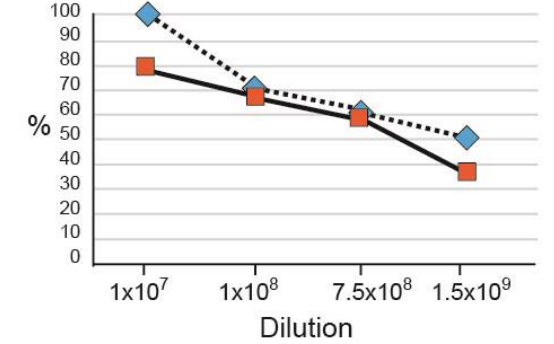
- We can influence the canine sensitivity and specificity through reinforcement schedules



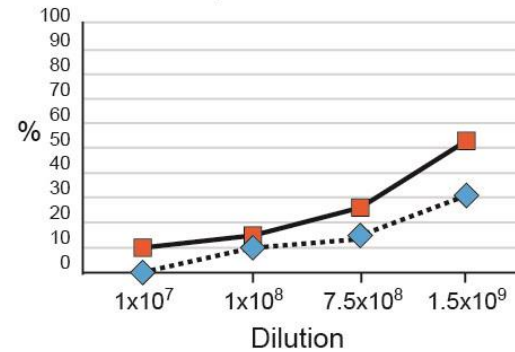
Proportion Hits - Total Exposure



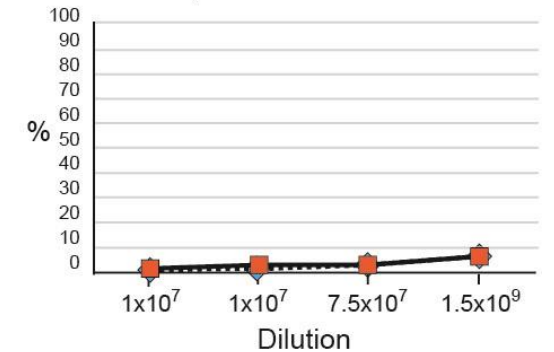
Proportion Hits - First Pass



Proportion Misses



Proportion False Alert



Concha, A., Mills, D.S., Feugier, A., Zulch, H., Guest, C., Harris, R., Pike, T.W: Using sniffing behaviour to differentiate true negative from false negative responses in trained scent-detection dogs. Chemical Senses, Sept. 2014.

Canine interface for detection dogs

- The olfactory performance in detection dogs may be influenced by training factors such as handler errors (Back and McLean 2003; Lasserter et al. 2003; Wasser et al. 2004; Lit et al. 2011)
- The impact of external factors on the olfactory detection performance can only be measure based on the presence/absence of the trained alert response
- No trained alert for equivocal response –is ‘grey’ yes or no..

Improved Understanding of Canine Training and Decision Making, and Communication to Empower Our Dogs



Interactive stands

- We are developing new technology with a sensor pad designed to sense the level of pressure the dog exerts whilst sniffing
- The level of pressure is recorded by a computer that is attached to the stand, this indicates the level of certainty that the dog has the particular disease is present



Interactive stands



Mancini, C., Harris, R., Aengenheister, B., Guest, C. (4/2015). "Re-Centering Multispecies Practices: a Canine Interface for Cancer Detection Dogs", 33rd International ACM CHI Conference on Human Factors in Computing Systems, ACM CHI'15, ACM Press, pp. 2673

Bio- detection projects: Malaria

- Trained dogs identified asymptomatic malaria infection with a high degree of accuracy in children - 81% sensitivity and 92% specificity.
- In line with the WHO's criteria for the procurement of rapid diagnostic tests.



THE LANCET
Infectious Diseases

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CORRESPONDENCE | VOLUME 19, ISSUE 6, P578-580, JUNE 01, 2019

PDF [56 KB]

Trained dogs identify people with malaria parasites by their odour

Claire Guest · Margaret Pinder · Mark Doggett · Chelci Squires · Muna Affara · Balla Kandeh · et al. [Show all authors](#)

Published: June, 2019 · DOI: [https://doi.org/10.1016/S1473-3099\(19\)30220-8](https://doi.org/10.1016/S1473-3099(19)30220-8)

Supplementary Material

Eliminating malaria would be simpler if a non-invasive method was available for detecting infected individuals in populations with low malaria prevalence. Infected individuals could then be treated with

Bio- detection projects: Pseudomonas



Davies JC, Alton E, Simbo A, Murphy R, Seth I, Williams K, Somerville M, Jolly L, Morant S, and Guest C., (11/2019) Training dogs to differentiate *Pseudomonas aeruginosa* from other cystic fibrosis bacterial pathogens: not to be sniffed at?. *Eur Respir J* 2019

Bio- detection projects: Cancer

Church, J. and Williams, H., (2001). “Another sniffer dog for the clinic?” *The Lancet*, 358(9285), p.930.

First robust published paper, BMJ 2004:

Willis, C.M., Church, S.M., Guest, C.M., Cook, W.A., McCarthy, N., Bransbury, A.J., Church, M.R. and Church, J.C., (2004). “**Olfactory detection of human bladder cancer by dogs: proof of principle study**”. *BMJ*, 329(7468), p.712.

Willis, C.M., Britton, L.E., Harris, R., Wallace, J. and Guest, C.M., (2011). “**Volatile organic compounds as biomarkers of bladder cancer: Sensitivity and specificity using trained sniffer dogs**”. *Cancer Biomarkers*, 8(3), pp.145-153.

C. Guest, R. Harris, K. Sfanos, A. Partin, B. Trock, L. Mangold, H. Steen, R. Bader, A. Kozak, J. Simons, H. Soule, T. Johnson, W-Y Lee, F. Turlomousis, T. Karydis, P. Stathatou, S. Thaler, A Spiliotopoulos, M. Weinstein, B. Chen and A. Mershin “**GC-MS and Canine/Machine Olfaction for Early Prostate Cancer Diagnostics**” poster presented: (2019) 26th Annual Prostate Cancer Foundation Scientific Retreat



Dog detection to electronic application



Development of Bio- Nose

Smell, the final frontier. To boldly go where no man has gone before.



Guest C, Harris R, Sfanos KS, Shrestha E, Partin AW, Trock B, et al. (2/2021) "Feasibility of integrating canine olfaction with chemical and microbial profiling of urine to detect lethal prostate cancer".

PLoS ONE 16(2): e0245530

COVID-19 Detection: Phase 1



Millie, aged 4, Golden Retriever



Tala, aged 3, Labrador



Marlow, aged 4, Labrador



Kyd, aged 4, Labrador X Golden Retriever



Asher, aged 8, Working Cocker Spaniel



Lexi, aged 5, Labrador

COVID-19 Detection: Phase 1

- COVID-19 infection does have a distinct odour
- Dogs can detect it with incredible speed and accuracy, achieving a sensitivity range of 82.1 to 94.3% and a specificity range of 76.4 to 92.0%
- The accuracy of dogs is consistent – detecting asymptomatic cases, and when the viral load is low
- Dog screening plus a PCR test could prevent more onward transmission than isolating symptomatic individuals only, or testing people with a LFT plus a PCR test

What Next? – Development

| Screening by Sample | Indirect Screening | Passive Screening |
|--|---|--|
|  |  |  |

Image Credit: Joe Raedle/Getty

Passive Search: Association of Covid-19 and Ball Reward

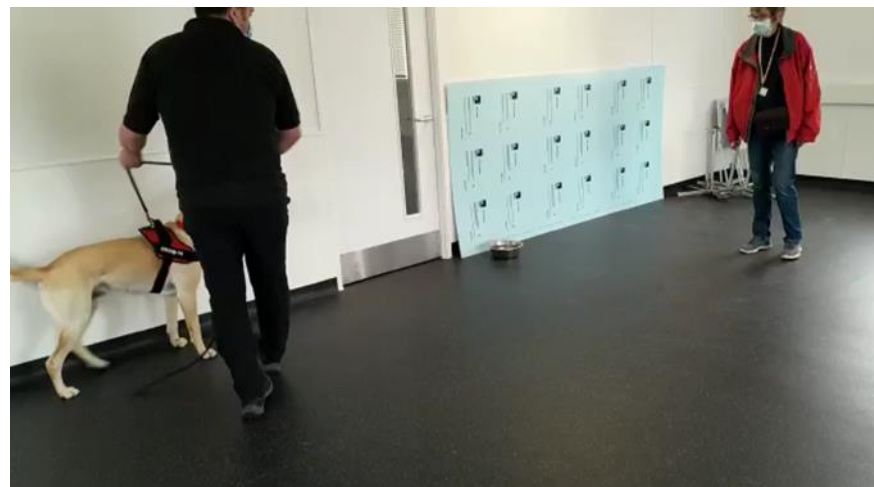


Restrictive Factors: Passive Screening

| Samples | Training | External |
|--|--|--|
| Procurement of continuation training samples | Random or inconsistent exposure to training sample materials | Limited access to training & test locations |
| Sample collection materials: storage, uses & shelf-life | Odour 'Tags' developed during training | Changes in UK covid legislations including removal of testing & self-isolation |
| Procurement of Covid-19 positive participants for transition & continuation training | Transition from collected samples to direct detection | Varied rates of infection |



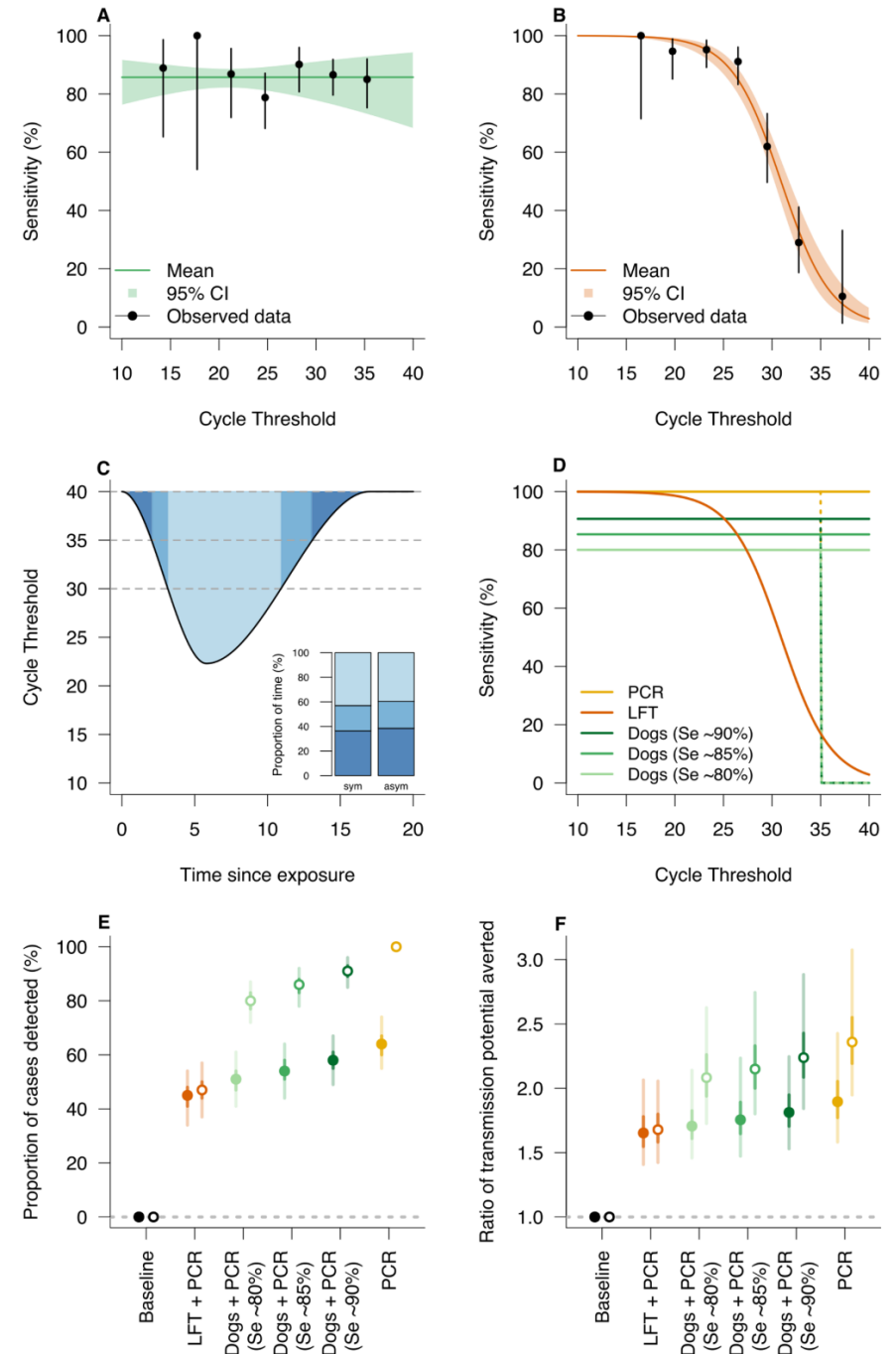
Target: Position 3
Confirmed Positive by LFT (Day 1)
Dog: Storm
Run: 1
Indication: **NO**



Target: Position 2
Confirmed Positive by LFT (Day 4)
Dog: Storm
Run: 1
Indication: **YES**

Mathematical modelling

- Dogs could be highly effective in detecting cases and averting transmission.
- Dogs detect low viral loads.
- 91% of cases detected with dog screening plus PCR test strategy.
- LFT plus PCR test strategy less effective in comparison.
- Dogs are significantly quicker



Mathematical modelling

- Two dogs could screen 300 people in 30 minutes.
- PCR testing only needed on those identified as positive by the dogs.
- Dogs may also serve as a visual deterrent.
- Possibilities for use at other mass gatherings.

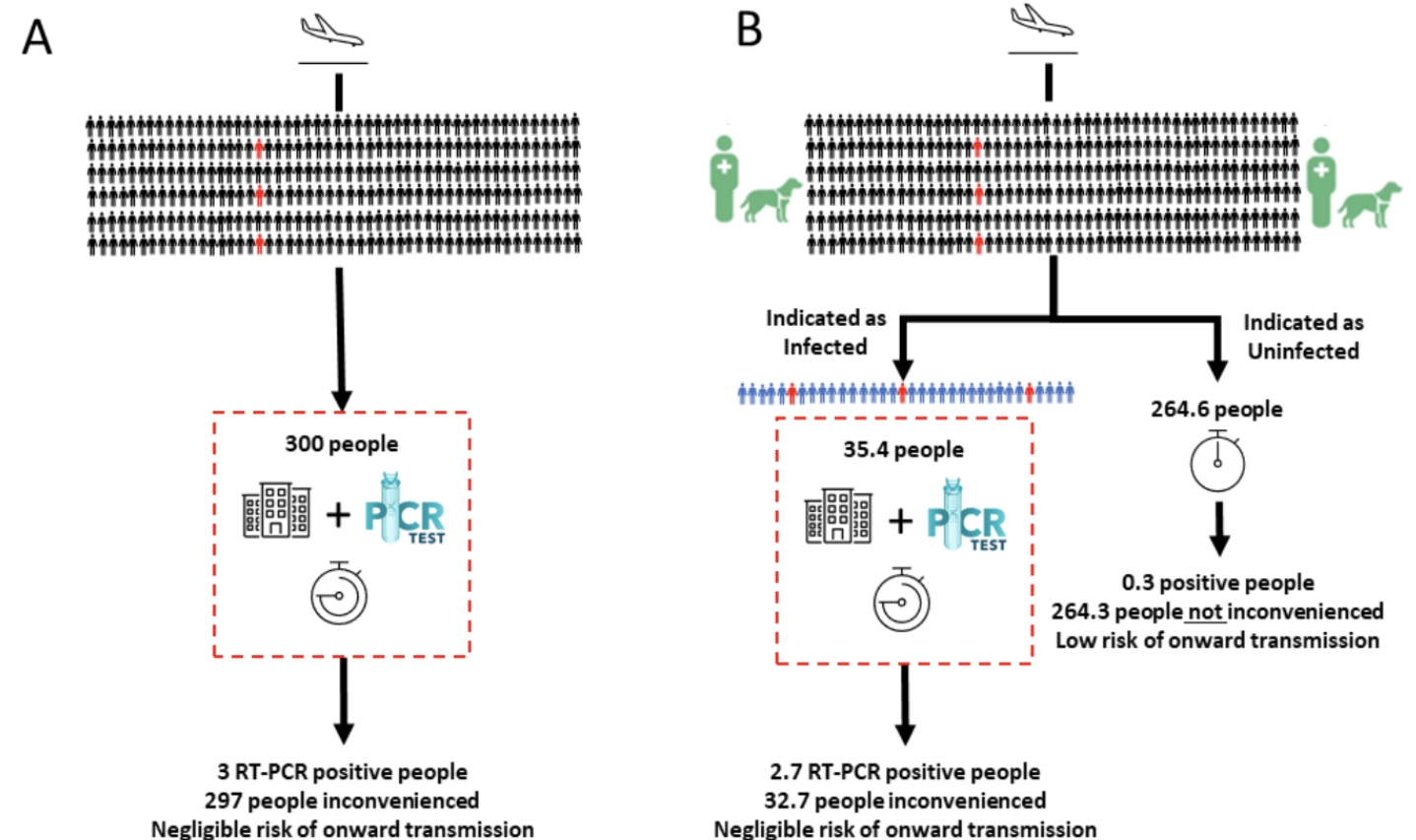
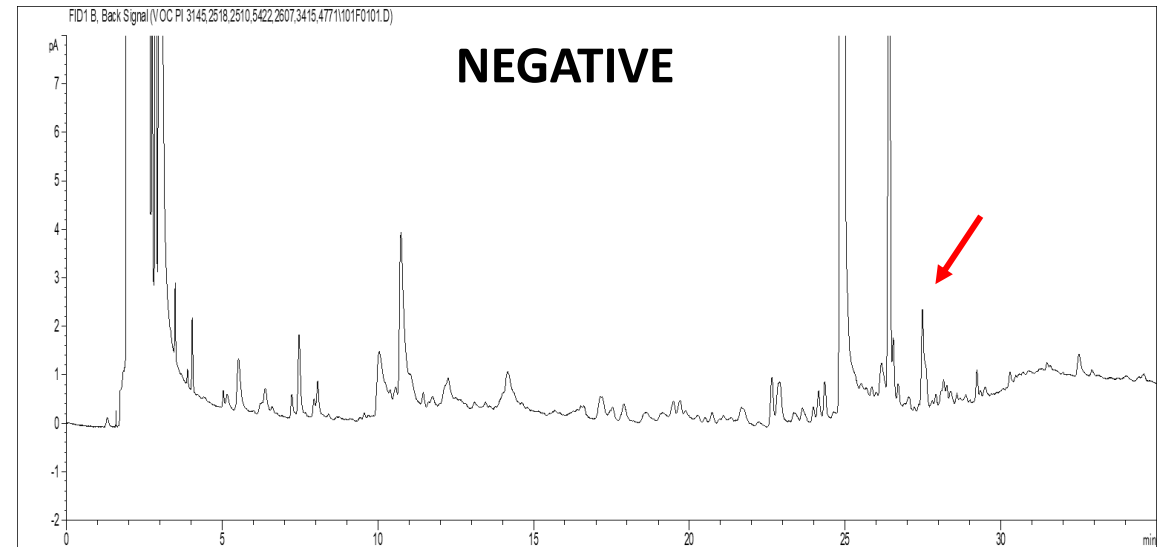
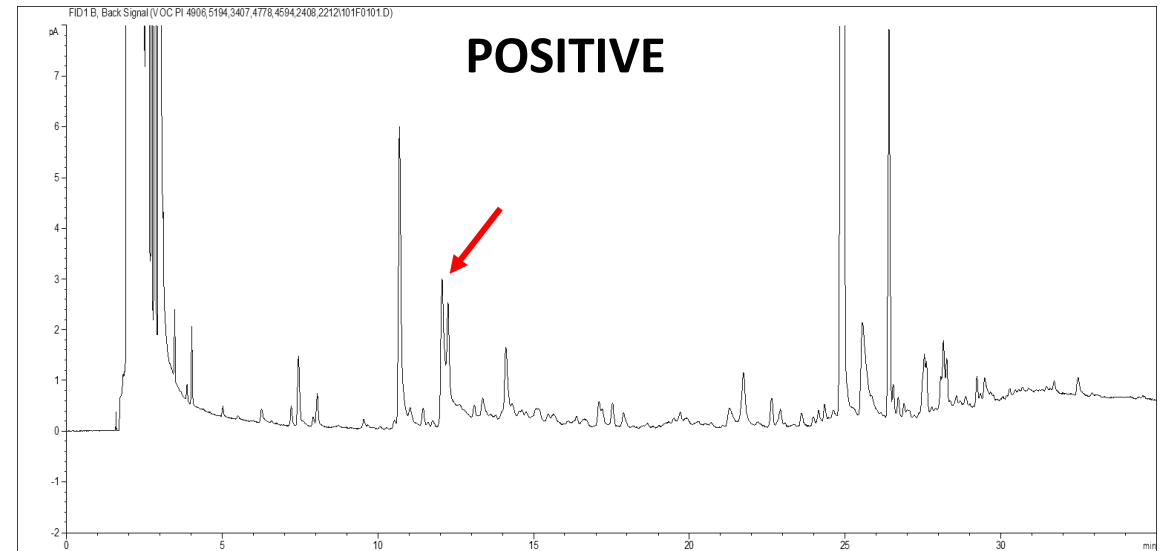


Figure 4: Exemplar of (A) Current SARS-CoV-2 Strategy and (B) Proposed Rapid Screen and Test Strategy. Schematic outlining the number of true negatives (black) and true

VOC analysis

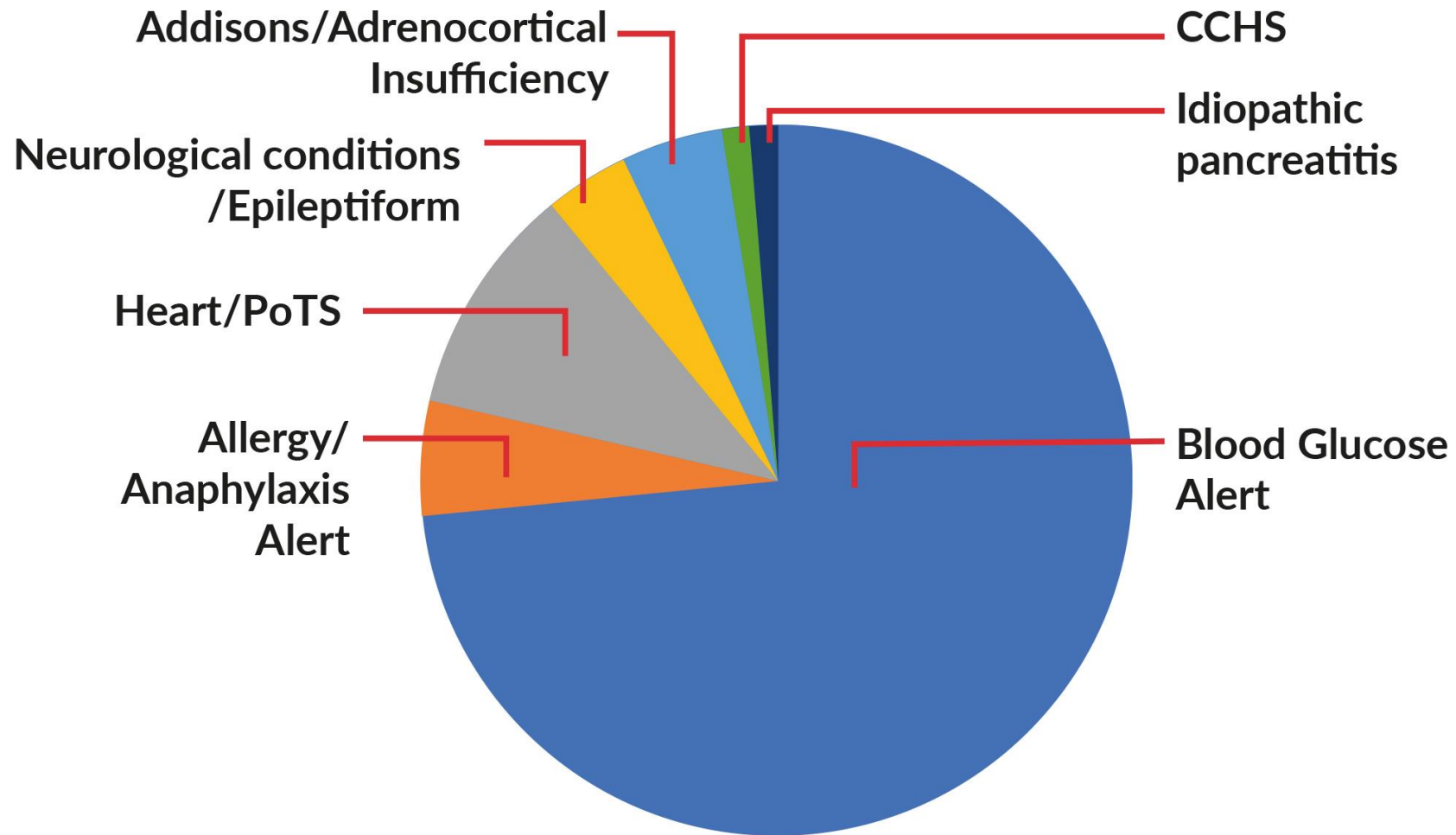
- Approximately 15 compounds significantly different.
- Hexanal, 5-methylhexanal, heptanal, 2-octanone, 7-octenal, nonanal, and 2-methyldodecane (tentative identification)
- Sensors



Client Partnerships



Qualified Partnerships by Condition



Summary

- Canine bio sensor
- Robust evidence based data
- Advanced training and communication methods
- Canine welfare
- Appropriate application of canine for maximum impact
- Scalable responses to disease detection

