Antimicrobial Use of Nitric Oxide to Reduce Problems Associated with Catheterisation

M.J.Duncan¹, S.J.Warrender¹, E.M.Coghill², I.L.Megson², B.Millar³, G.Menary³, M.J.Swann⁴, F.Watson⁴, S.L.Percival^{4*} and R.E.Morris¹

1. School of Chemistry, University of St Andrews, St Andrews, 2. Biomedical Sciences, University of Highland and Islands, Inverness, 3. School of Mechanical and Aerospace Engineering, Queen's University Belfast, Belfast,



BACKGROUND

4. 5D Bioscience. 5D Health Protection Group Ltd. Accelerator Building. 1 Daulby Street, Liverpool L7 8XZ, UK,

Cardiovascular Catheters

- coronary heart disease is the leading cause of death worldwide
- · treatment involves implantation of stents using catheters
- problems associated with catheterisation

Infection





localised and / or systemic infection

to this blood clot

catheter blocked due incidence rate ranges from 4 - 20 %

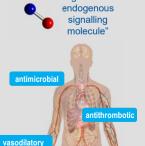
• 10 million percutaneous interventional procedures conducted per year

METHOD

Preventing Problems Associated With Catheterisation

· release nitric oxide from the surface

- of catheters
 - "gaseous



- **ISSUES** · difficult to administer toxic in high concentrations
- INOI dependant response
- NO loading

SOLUTION

· deliver from a MOF

suitable structure CPO 27

release of NO from the surface of the tubing

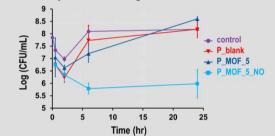
- · elicits a marked and prolonged reduction in bacterial growth
- · shows small colony variants viable bacteria are highly stressed

ANTIMICROBIAL RESULTS PROTOCOL inoculate inside of tube with MRSA/TSB 25 µL ID 1.25 mm x107 CFU/mL • incubate at 37 °C, up to 24 hr



RESULTS

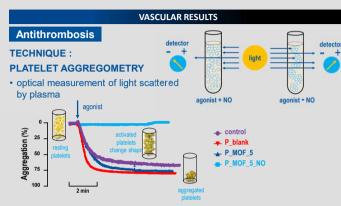
bacterial cell viability from inside tubing measured over time course



• spread plates at 6 hr time point



tubing inhibits infection

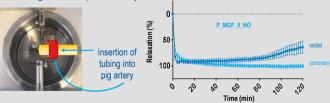


•sufficient NO released to stop cells in the blood from sticking together

Vasodilation

TECHNIQUE: MYOGRAPHY

measuring the force produced by the blood vessel when under contraction



· coronary arteries more receptive to NO than radial arteries

CONCLUSIONS

- demonstrated that MOFs can be extruded into polymer tubes
- verified NO loaded MOF/polymer formulations are biocompatible
- · capacity to tune the NO elution profile
- · ability to deliver an efficacious dose of NO which meets the clinical challenges associated with cardiac catheterisation