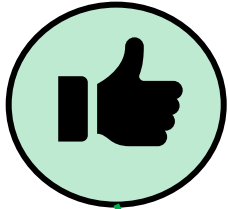


SURFACE DECONTAMINATION

Jean-Yves Maillard
Cardiff School of Pharmacy and
Pharmaceutical Sciences
Cardiff University

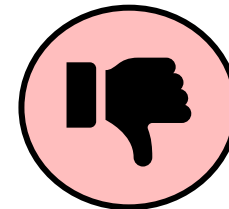


COVERED

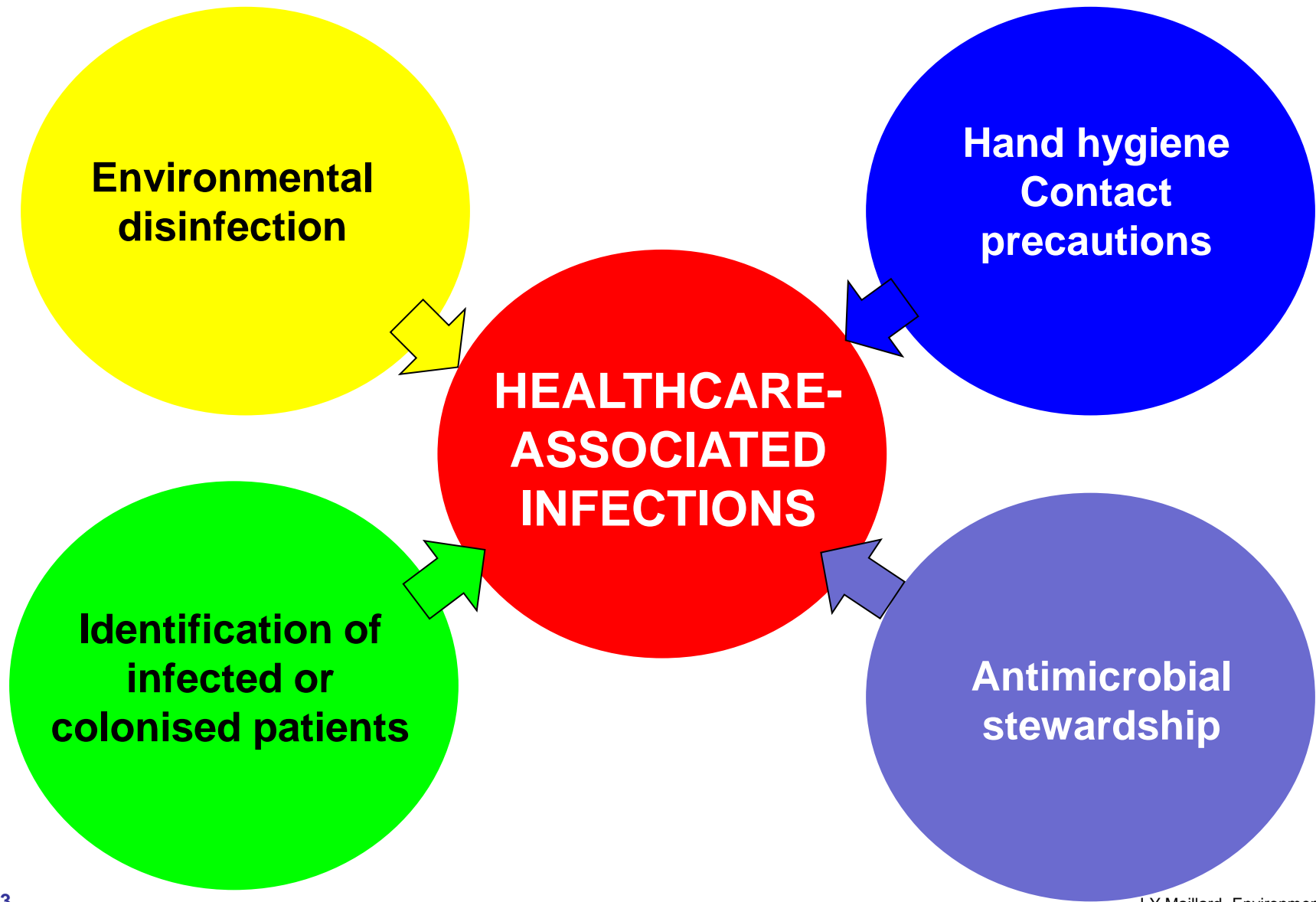
- Chemical disinfection
- Factors affecting activity
- Biocide delivery

NOT COVERED

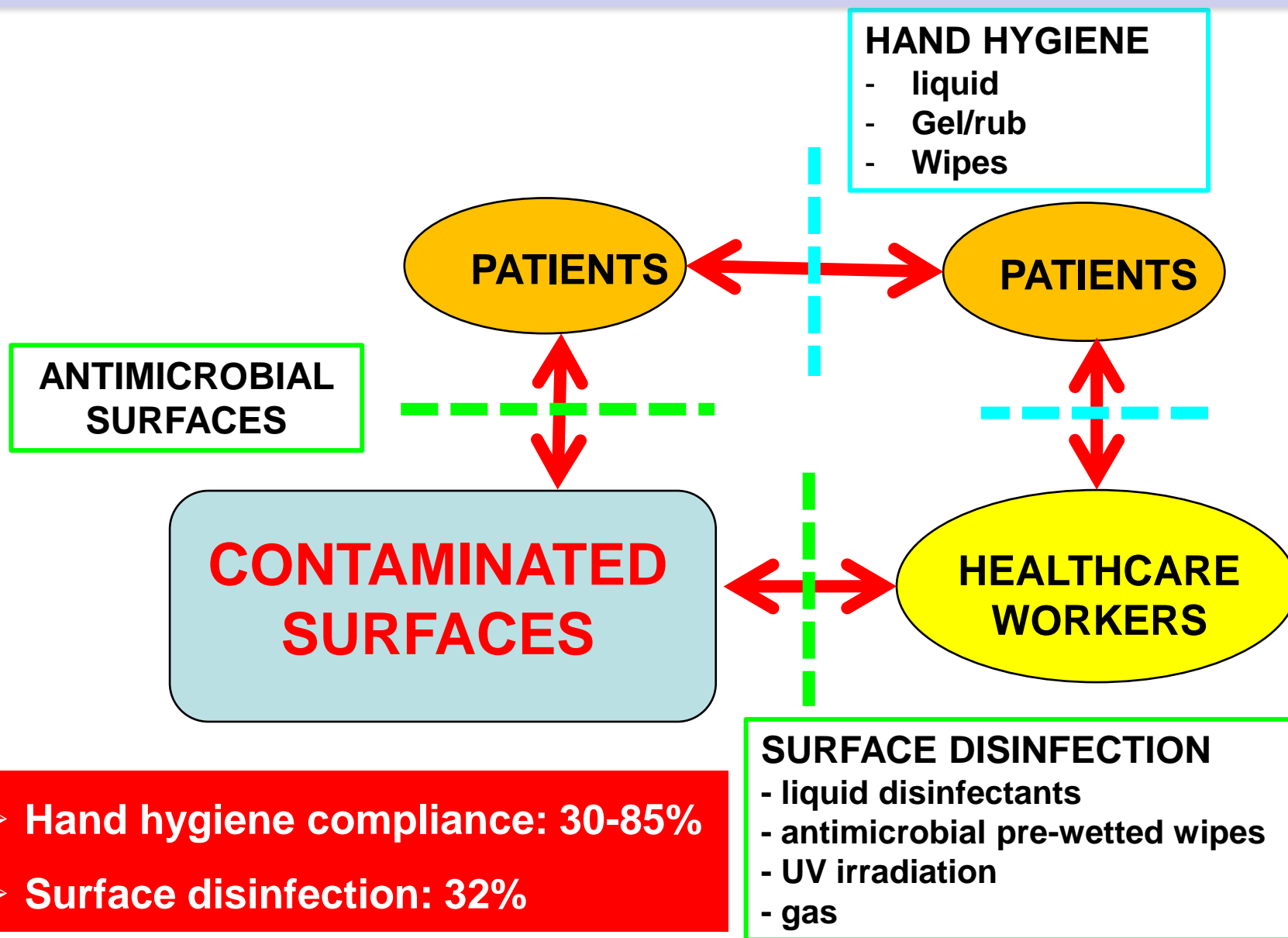
- Physical disinfection (UV-irradiation)
- Specific product efficacy
- Detailed standard efficacy test
- Product regulation



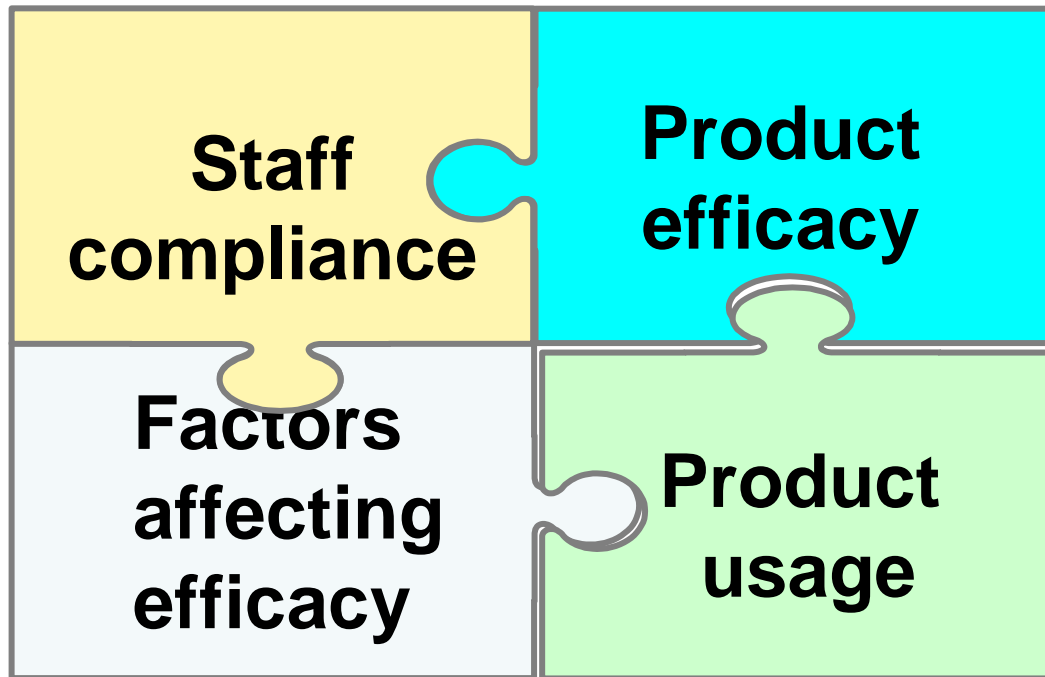
Breaking the chain of transmission

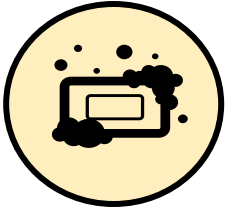


Breaking the chain of transmission



Breaking the chain of transmission



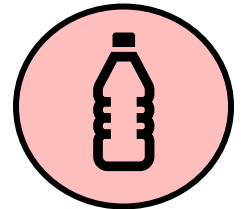


CLEANING

- Removal of dirt
- Might remove pathogens
- Not designed to kill pathogens

DISINFECTION

- Killing of pathogens
- Different levels of disinfection



DECONTAMINATION

- Decrease of microbial bioburden
- Render a surface safe?



- Make a surface safe?
- Decrease number of pathogens to a safe level?
- What is visibly clean?

Revised Spaulding Classification

Spaulding classification

CRITICAL DEVICES

In contact with sterile
tissue or vascular
system



Sterilisation required

- Physical sterilisation (heat)
- Radiation sterilisation
- Chemical disinfectant (ethylene oxide, hydrogen peroxide)

High-level disinfection

(may be acceptable)

- PAA, H₂O₂, ClO₂, GTA, OPA,

SEMI-CRITICAL DEVICES

In contact with
intact mucous
membrane



High-level disinfection required

- Heat
- Glutaraldehyde
- Peracetic acid
- Chlorine dioxide
- Hydrogen peroxide
- Chlorine releasing agents

NON-CRITICAL

Not in contact with
patient/ contact with
intact skin



Low-level disinfection required

- Alcohols
- QACs
- Biguanides

Cleaning or disinfection

- Cleaning and drying
- Disinfection in case of contaminated spillage

HARD SURFACE DISINFECTION

- Chlorine releasing agents
- Peroxygen-based products
- Quaternary ammonium compounds
- Biguanides
- Phenolics
- Organic acids
- Antimicrobial dyes
- Iodine
- Natural antimicrobials



FORMULATION

- Complex - multiple QACs
- Excipients - surfactants, wetting agents

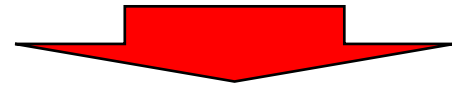
DELIVERY

- Spray
- foam
- Mist
- Pre-wetted wipes

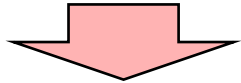
CLEANING

- Detergents: ionic, non-ionic, amphoteric surfactants, alcohols
- Cationic surfactants (QAC) – biocides

Factors affecting efficacy



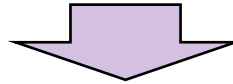
FACTORS INHERENT TO THE PRODUCT



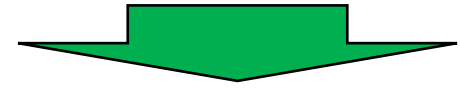
- Concentration
- Formulation
- pH
- Delivery



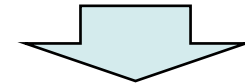
FACTORS INHERENT TO THE APPLICATION OF THE PRODUCT



- Surface
- Organic load (soiling)
- Temperature
- Contact time
- Humidity
- Material (fabric)
- *Delivery*



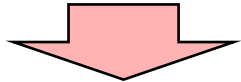
FACTORS INHERENT TO THE MICRO-ORGANISMS



- Type
- Number
- Phenotype
- Association (biofilms)



FACTORS INHERENT TO THE PRODUCT



- Concentration
- Formulation
- pH
- Delivery (material)

IMPORTANCE

+++

++

+*

++

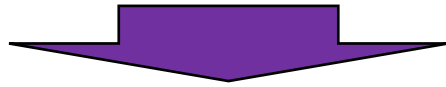
PREDICTABILITY

- Concentration exponent (η)
- Bioavailability
- Residual concentration

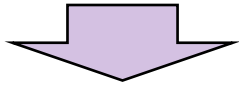
?

Organic acids

- Release or not of antimicrobial
- Bioavailability?



FACTORS INHERENT TO THE APPLICATION OF THE PRODUCT



IMPORTANCE

- Surface
- Organic load (soiling)
- Temperature
- Contact time
- Humidity
- Material (fabric)

++

++

+*

++

++

PREDICTABILITY

?

- Depend on the active
- Temperature coefficient (Q_{10})
- Continuous release (reservoir)?
- Short contact time
- Aerial disinfection
- Retention / compatibility

Factors affecting efficacy

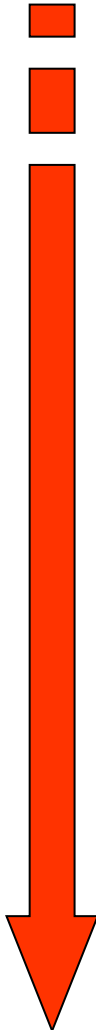
Biocide		Concentration Exponent	Soiling
Phenolics	Triclosan	4-9.9	+++
Alcohols	Benzyl alcohol Aliphatic alcohols	2.6-4.6 6.0-12.7	+
Cationics	Chlorhexidine Polymeric biguanides QACs Dyes (Crystal violet)	2 1.5-1.6 0.8-2.5 0.9	++
Aldehydes	Formaldehyde Glutaraldehyde	1 1	+
Peroxygens	Hydrogen peroxide Peracetic acid	0.5 0.5	+++
Metallic salts	Silver nitrate Mercurials	0.9-1.0 0.03-3.0	+++
Organic acid	Parabens Sorbic acid	2.5 2.6-3.2	++

➤ **Concentration exponent:** the higher it is the more loss of efficacy upon dilution

Factors affecting efficacy

LOW

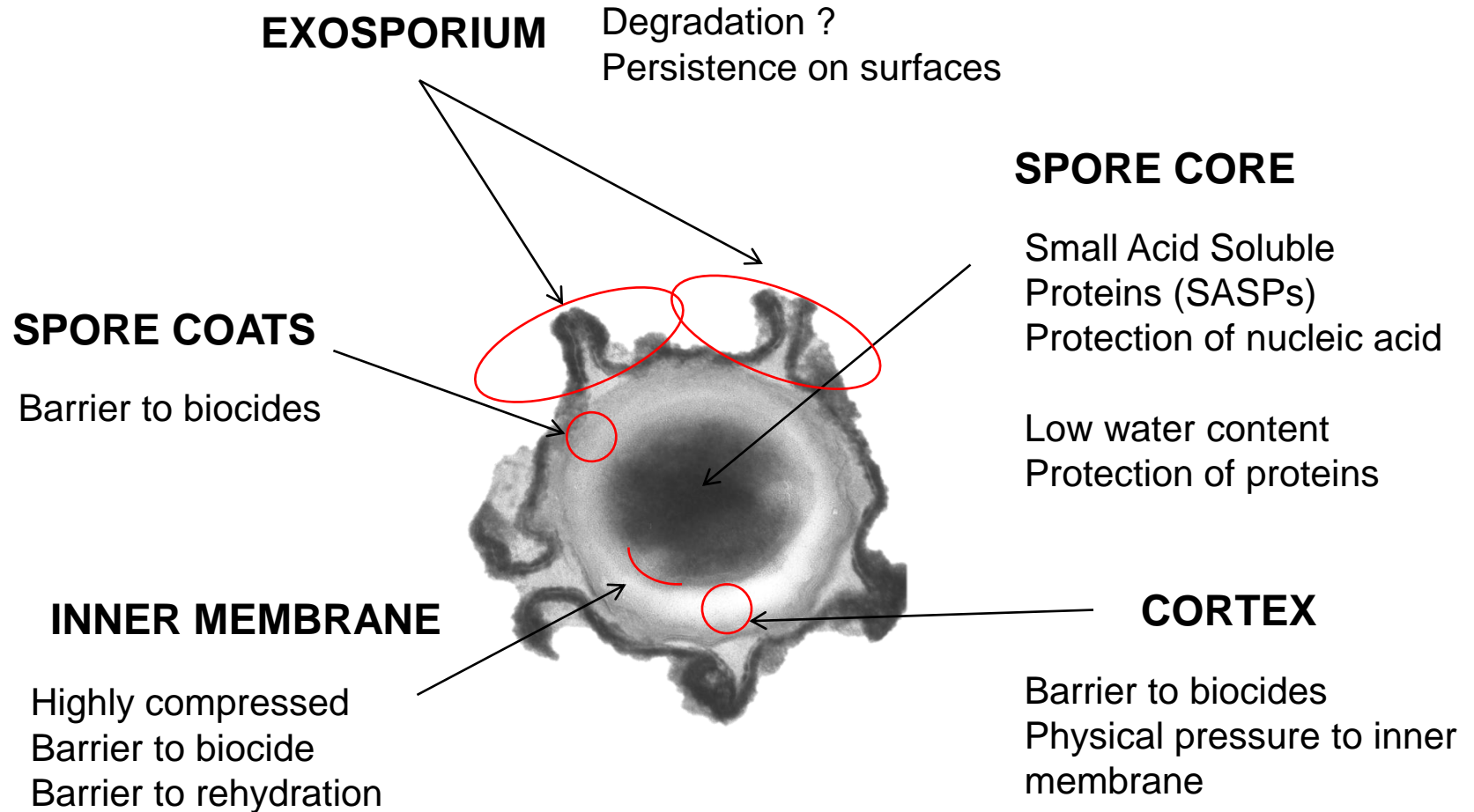
SUSCEPTIBILITY LEVEL TO
DISINFECTION



HIGH

Group	Examples
Prions	PrP ^{res} , a protein-only agent implicated in transmissible diseases such as scrapie (in sheep) and Creutzfeld-Jakob disease (in humans)
Bacteria Endospores	<i>Bacillus</i> , <i>Clostridioides</i> spores
Protozoal Oocysts ?	<i>Cryptosporidium</i> oocysts
Helminth Eggs or Cysts ?	<i>Ascaris</i> , <i>Enterobius</i>
Protozoal Cysts	<i>Giardia</i> , <i>Acanthamoeba</i>
Mycobacteria	<i>Mycobacterium tuberculosis</i>
Small, Non-Enveloped Viruses	Poliovirus, Parvoviruses
Fungal Spores	<i>Aspergillus</i> , <i>Candida</i>
Gram-negative bacteria	<i>Pseudomonas</i> , <i>Escherichia</i>
Vegetative Fungi	<i>Trichophyton</i> , <i>Candida</i>
Adult Helminths and Protozoa ?	<i>Ascaris</i> , <i>Cryptosporidium</i>
Large, non-enveloped viruses	Adenoviruses, Rotaviruses
Gram positive bacteria	<i>Staphylococcus</i> , <i>Streptococcus</i>
Enveloped viruses	HIV, Vaccinia

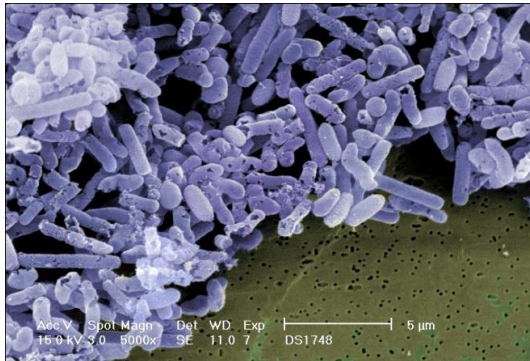
Challenge - endospores



Leggett et al. *J Appl Microbiol* 2012; 113: 485-98.

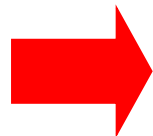
Product claim – sporicidal activity

**BACTERICIDAL
ACTIVITY**

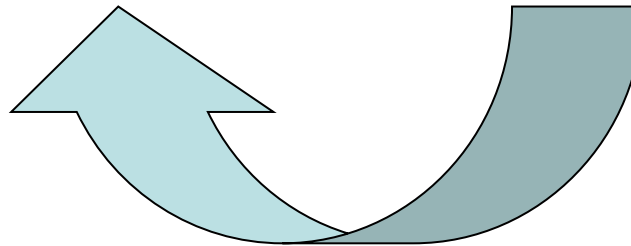
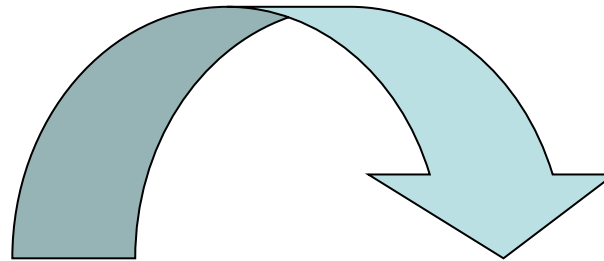


OUTGROWTH

**SPORISTATIC
ACTIVITY**

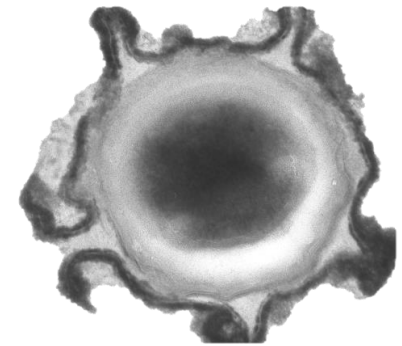


SPORULATION



GERMINATION

**SPORICIDAL
ACTIVITY**



SPORICIDAL ACTIVITY



- Hydrogen peroxide
- Peracetic acid
- Chlorine dioxide
- Ozone
- Ethylene oxide
- Glutaraldehyde
- Formaldehyde
- *ortho*-phthalaldehyde
- Sodium hypochlorite
- Sodium dichlororisocyanurate (?)
- Chloramine-T
- Calcium hypochlorite
- Iodine and iodophors

“SPORISTATIC” ACTIVITY

- Phenols and cresols
- Quaternary ammonium compounds
- Biguanides
- Organic acids and esters
- Alcohols





**“Sporicidal 70% Alcohol Gel & Moisturiser (250ml)
Kills 99.999% of bacteria**

Sporicidal 70% Alcohol Hand Gel with moisturiser, 250ml with flip top cap - the only alcohol hand gel effective against C. Difficile spores.”

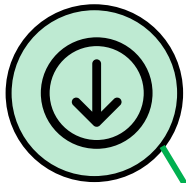


“Product C to date has killed every virus, spore and bacteria it has been tested on including MRSA, C-Diff, Norovirus and many more pathogens in 1/5 of the EN1276 required time making it 1000 time more effective than EN1276 requirements - EN1276 is the chemical disinfectants and antiseptics.”

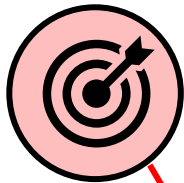
PRINCIPLE FOR MICROBICIDAL EFFICACY



➤ Need contact between the target micro-organism and the antimicrobial



➤ Need to penetrate to reach target site(s)



➤ Number of targets affected and overall damage to the target(s) produce a lethal or an inhibitory effect

Chlorhexidine-based products

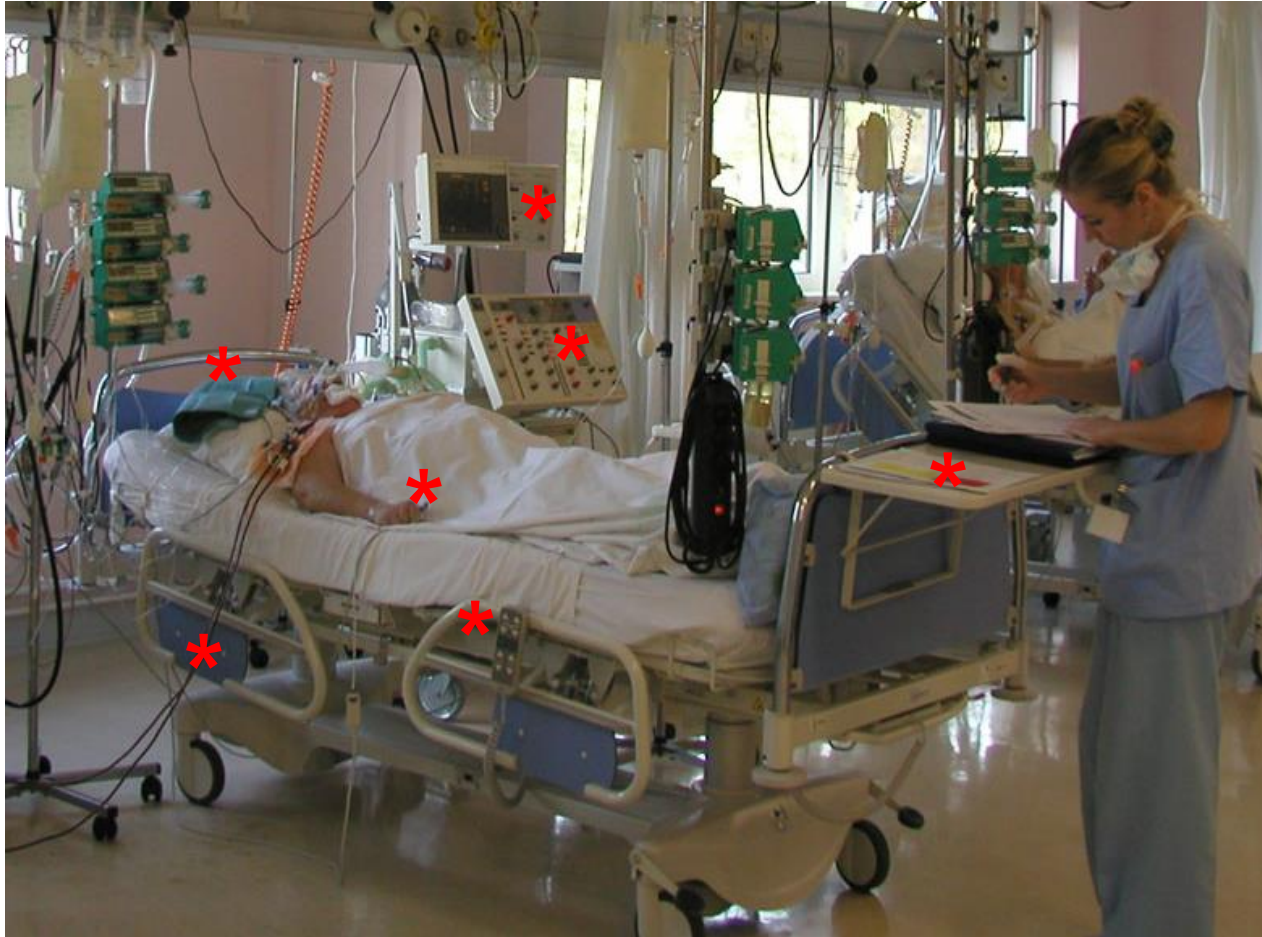
Contaminant(s)	Site(s) of microbes	Mechanism of contamination/source
<i>Pseudomonas</i> spp.	Not stated	Refilling contaminated bottles ; washing used bottles using cold tap water; contaminated washing apparatus; low concentration (0.05%)
<i>Pseudomonas</i> sp., <i>Serratia marcescens</i> , <i>Flavobacterium</i> sp.	Not stated	Not determined, but authors speculate due to overdilution or refilling of contaminated bottles
<i>Pseudomonas aeruginosa</i>	Wounds	Tap water used to dilute stock solutions ; low concentration (0.05%)
<i>Bulholderia cepacia</i>	Blood, wounds, urine, mouth, vagina	Metal pipe and rubber tubing in pharmacy through which deionized water passed during dilution of chlorhexidine; low concentration
<i>Ralstonia pickettii</i>	Blood	Contaminated bidistilled water used to dilute chlorhexidine ; low concentration (0.05%)
<i>Ralstonia pickettii</i>	Blood (pseudo-bacteremia)	Distilled water used to dilute chlorhexidine; low concentration (0.05%)
<i>Serratia marcescens</i>	Bood, urine, wounds, sputum, others	Not determined, but use of nonsterile water for dilution to 2% and distribution in reusable nonsterile containers
<i>Bulholderia cepacia</i>	Blood	Intrinsic contamination , Contaminated 0.5% chlorhexidine
<i>Serratia marcescens</i>	Blood	Intrinsic contamination , 2% aqueous chlorhexidine antiseptic

Failure to understand factors affecting efficacy

Benzalkonium chloride- based products

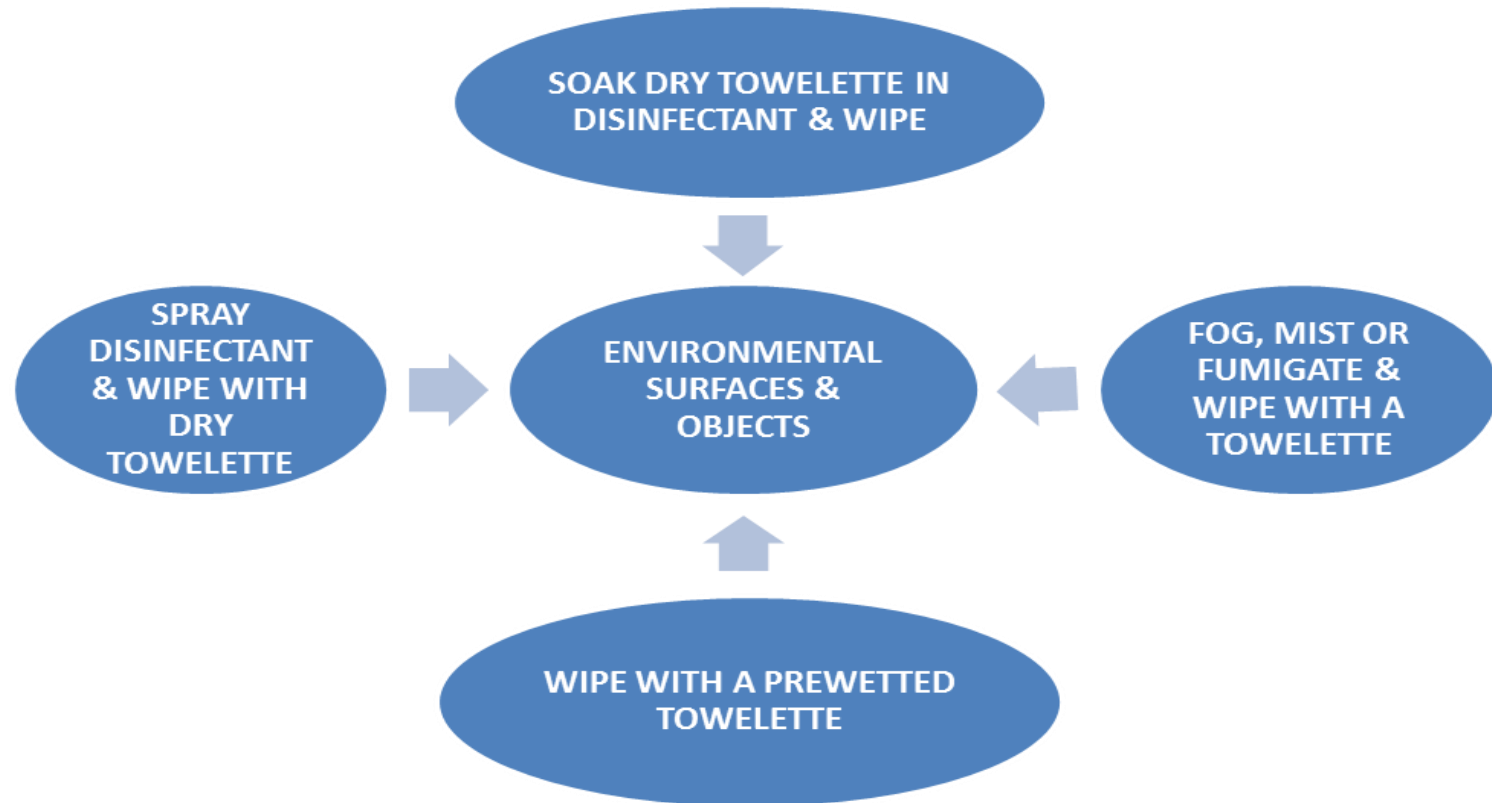
Contaminant(s)	Site(s) of microbes	Mechanism of contamination/source
<i>Enterobacter aerogenes</i>	Blood, sinus tract	Storage of benzalkonium chloride (0.13%) with cotton/gauze
<i>Pseudomonas-Achromobacteriaceae group</i>	Blood, urine	Storage of benzalkonium chloride (0.1%) with cotton/gauze; dilution with nonsterile water
<i>Enterobacter aerogenes</i>	Blood, sinus tract	Storage of benzalkonium chloride (0.1%) with cotton/gauze; dilution with nonsterile water
<i>Bulholderia cepacia, Enterobacter spp.</i>	Blood (pseudobacteremia)	Storage of benzalkonium chloride with cotton/gauze; improper dilution ; storage bottles infrequently sterilized
<i>Serratia marcescens</i>	Intravenous catheters (dogs and cats), other sites	Storage of benzalkonium chloride (0.025%) with cotton/gauze
<i>Mycobacterium chelonae</i>	Skin abscesses	Storage of benzalkonium chloride with cotton/gauze; improper dilution
<i>Pseudomonas aeruginosa</i>	Corticosteroid injection multidose vial	Inoculation with pseudomonads via needle puncture after vial septa were wiped with contaminated disinfectant
<i>Mycobacterium abscessus</i>	Joint	Storage of benzalkonium chloride with cotton/gauze; dilution with probable contaminated tap water
<i>Bulholderia cepacia</i>	Blood, catheter	1:1000 aqueous BAC solution

Surface decontamination



Possible scenarios for decontaminating environmental surfaces by wiping

Sattar and Maillard *AJIC* 2013;41:S97-S104.



Observation of usage in practice –cleaning staff in ITUs

- use of wipes – surface area
- contact
- rotation

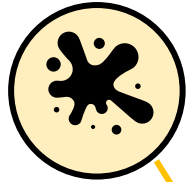
Wipe Number	Surface initially wiped	Time applied (seconds)	Number of consecutive surfaces wiped (other surfaces)
1	Bed Rail	4	5 (bedside table, monitor X2, monitor stand)
2	Steel Trolley	6	2 (both shelves on the trolley wiped)
1	Monitor	4	5 (monitors, two keypads, monitor stand)
2	Bed rail	7	4 (table, monitor, keypad)
3	Bedside table	10	4 (folder, two bed rails)

One wipe – one direction – one surface...bin it

Factors affecting efficacy of wipes



Sattar and Maillard *Am J Infect Control* 2013 41;S97-S104



Detergent wipes: to clean surfaces (removal dirt or organic matter)

- Essential to the cleaning process, acting to release dirt from a surface
- Not designed to remove microorganisms from surface, but they might



Disinfectant wipes

- Contain a biocide and may or may not contain an additional detergent
- Wipes that do not contain a detergent will have only limited cleaning properties
- Disinfectant wipes might have a claim to kill bacteria (bactericidal), fungi (fungicidal) viruses (virucidal) or bacterial endospores (sporicidal).



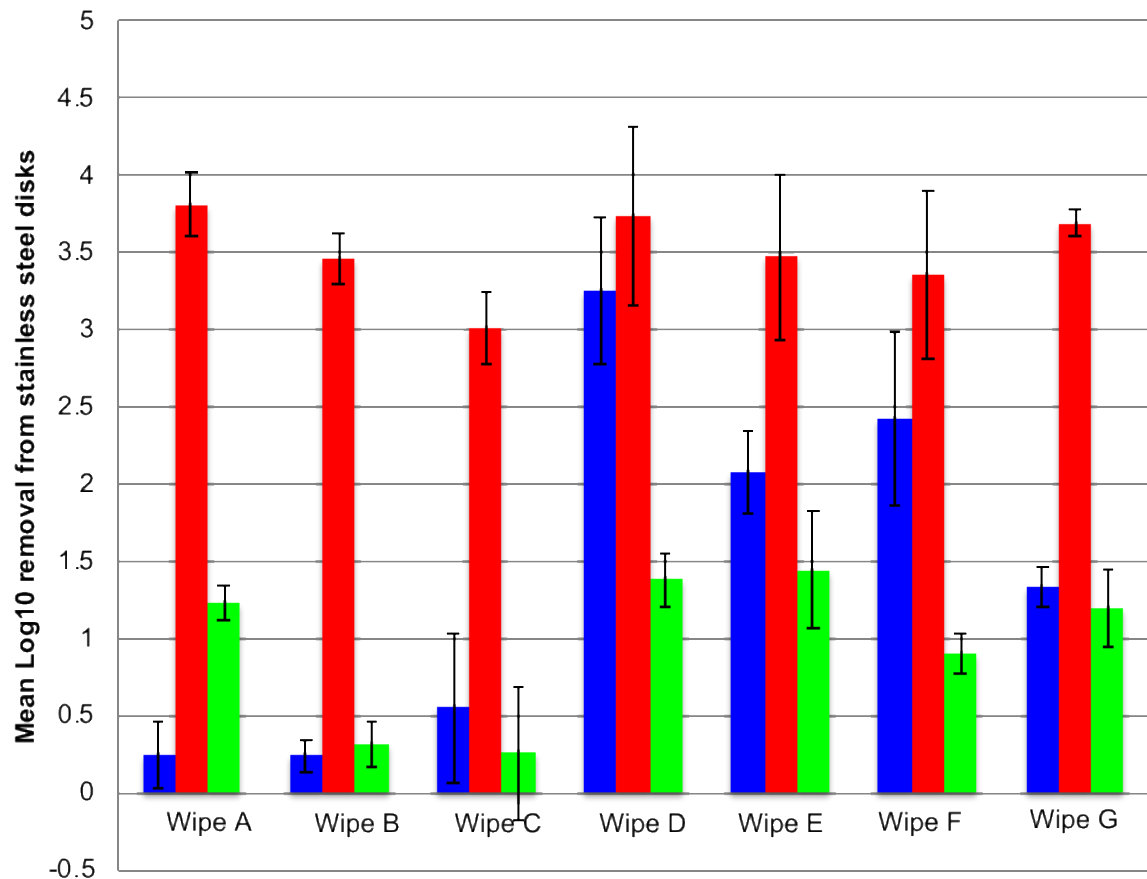
It is not possible to 'sterilise' surfaces of equipment/ patient environment using wipes alone; only to temporarily reduce the number of microbes present.

EFFICACY OF DETERGENT WIPES

Ramm *et al.* AJIC 2015;43:724-8.

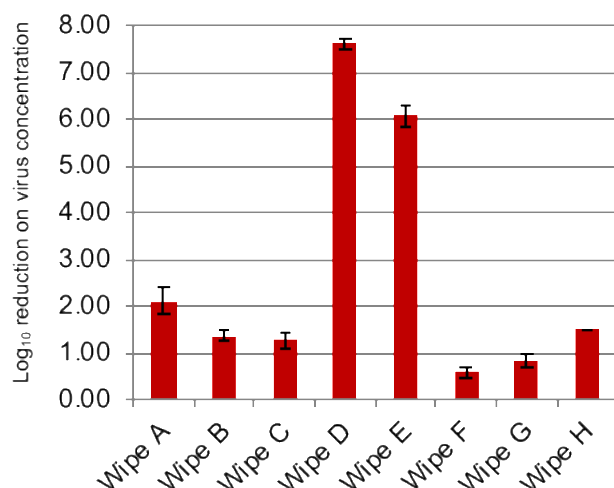
Bacterial/spore removal from surface

- *S. aureus*
- *A. baumannii*
- *C. difficile*

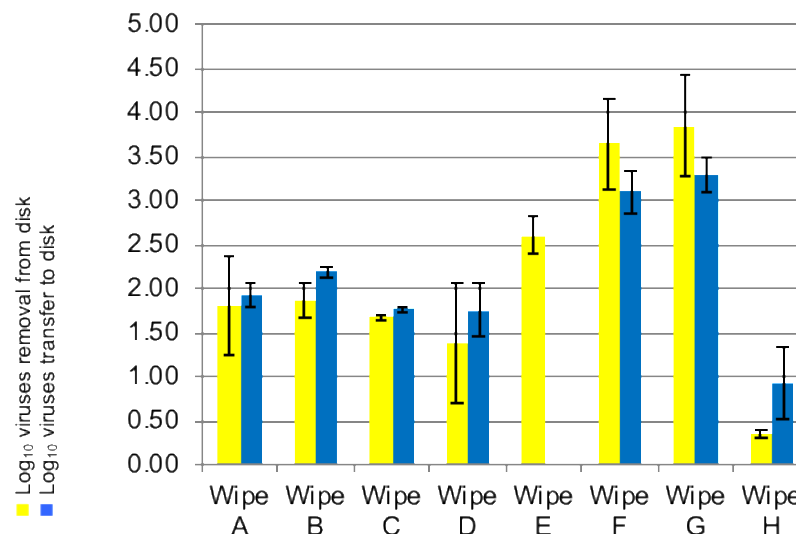


EFFICACY OF ANTIMICROBIAL WIPES AGAINST VIRUSES

Virucidal activity of expressed wipe solutions
(n=3)



Virus removal from disks and virus transfer
from wipes to disks. (n=3)



	Mean removal	Range
'Universal' wipes	1.68 log ₁₀	1.37-1.87 log ₁₀
Sporicidal wipes	3.13 log ₁₀	2.61- 3.65 log ₁₀
Detergent wipes	2.11 log ₁₀	0.36- 3.85 log ₁₀

Which wipes?

“SPORICIDAL” WIPES – efficacy testing against *C. difficile* NCTC12727

Siani *et al.* AJIC 2011; 39(3), 212-218

Wipes*	Bacterial Removal (log ₁₀ cfu/disk ± SD) 500 g surface pressure	Bacterial transfer following 10 s wiping time at 500 g surface pressure
Negative control	1.13 (± 0.36)	5 consecutive transfers. TNTC
NaOCl soaked wipe	2.02 (± 0.21)	5 consecutive transfers. TNTC
Wipe A	4.09 (± 0.79)	No spore transferred
Wipe B	0.22 (± 0.07)	5 consecutive transfers. From 0 to TNTC
Wipe C	1.30 (± 0.33)	5 consecutive transfers. From 0 to TNTC
Wipe D	0.57 (± 0.07)	5 consecutive transfers. From 1 to TNTC
Wipe E	+0.08 (± 0.08)	5 consecutive transfers. TNTC
Wipe F	1.14 (± 0.65)	5 consecutive transfers. From 83 to TNTC
Wipe G	0.67 (± 0.11)	5 consecutive transfers of ≤43 spores
Wipe H	0.88 (± 0.13)	5 consecutive transfers. From 2 to TNTC
Wipe I	0.84 (± 0.66)	5 consecutive transfers. From 40 to TNTC

* At the time of testing i.e. 2010-2011

Which wipes?

Study between 2006-2010; in 2008 chlorine based cleaning regimens and products changed to non-chlorine based sporicidal wipe

Average <i>C. difficile</i> rate per 1000 patients			
Financial year	Number of weeks	Mean <i>C. difficile</i> rate	Median <i>C. difficile</i> rate
2006-07	51	6.27	5.54
2007-08	52	6.99	5.95
Introduction of changes			
2008-09	51	2.05	1.74
2009-10	23	1.66	1.59

Carter Y and Barry D Nursing Time 2011, 107

- Changes included staff training on wipe usage
- Daily environmental cleaning (including surfaces at risks)
- Weekly ward visit to ensure good practice (e.g. supply of wipes)
- Weekly multidisciplinary ward round to monitor infection prevention and control measures
- Increase awareness campaign

Delivery – prewetted wipe vs. spray & cloth



Contents lists available at ScienceDirect

American Journal of Infection Control

Journal homepage: www.ajicjournal.org



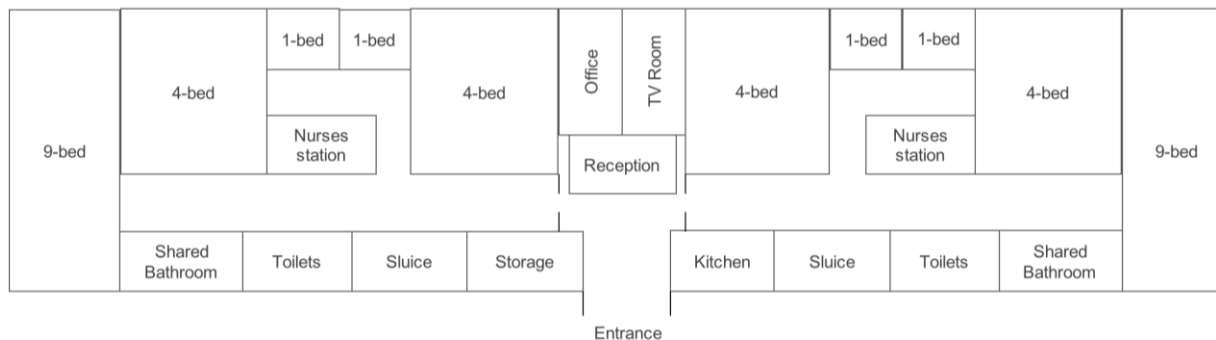
Major Article

Impact of antimicrobial wipes compared with hypochlorite solution on environmental surface contamination in a health care setting: A double-crossover study

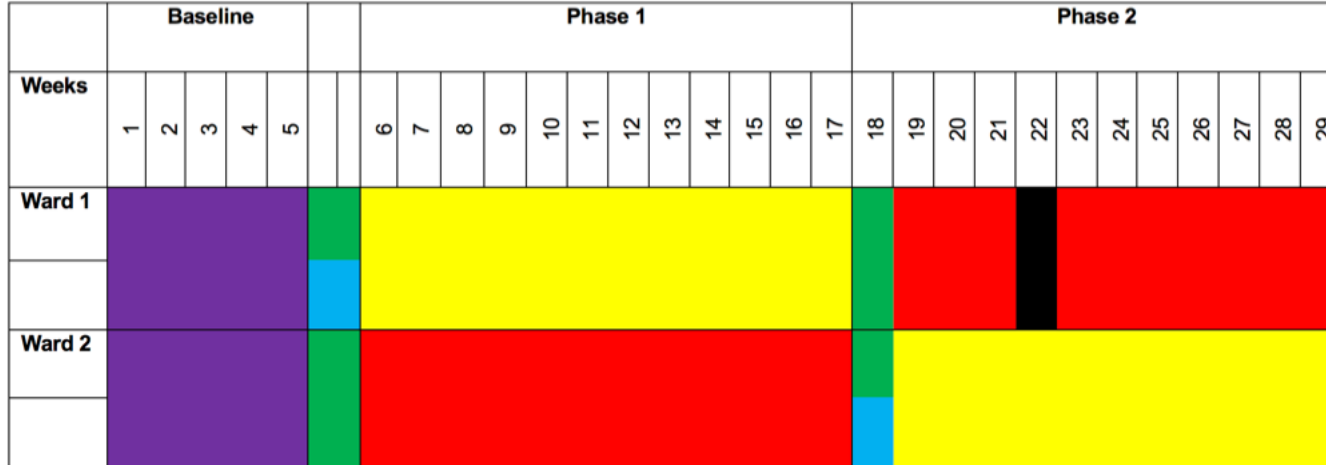
Harsha Siani BSc, Rebecca Wesgate BSc, Jean-Yves Maillard PhD *

School of Pharmacy and Pharmaceutical Sciences, Cardiff University, Cardiff, UK

- A double-crossover study was performed on 2 different surgical and cardiovascular wards in a 1,000-bed teaching hospital over 29 weeks.
- The intervention period that consisted of surface decontamination with pre-impregnated wipe or cloth soaked in chlorine



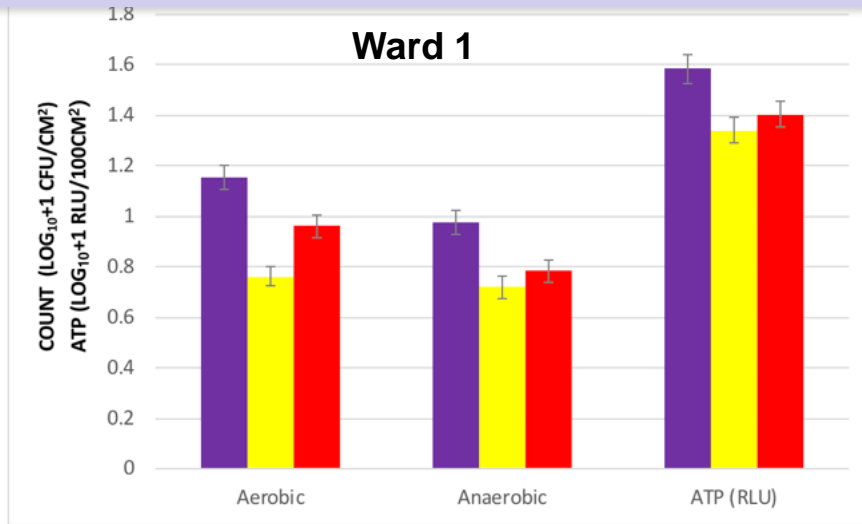
Delivery – preformulated wipe vs. spray & cloth



- Use of standard cleaning regimen.
- Use of detergent and chlorine 1,000 ppm.
- Use of pre-impregnated sporicidal wipes.
- Green shading indicates general training on disinfectant use, wiping, and infection prevention.
- Specific training on the use of preformulated wipes.
- Black shading indicates wards closure.

- Environmental samples from 11 surfaces were analysed weekly for their microbial content.
- In total, 1,566 environment samples and 1,591 ATP swabs were taken from the 2 wards

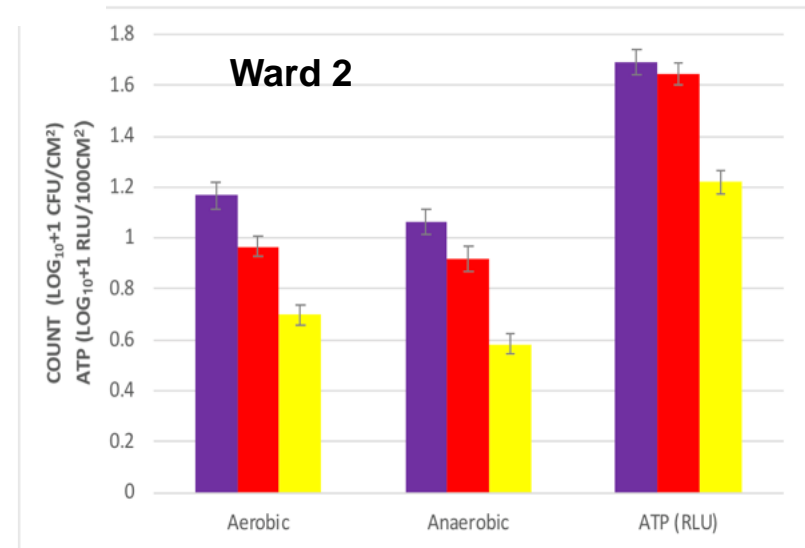
Delivery – preformulated wipe vs. spray & cloth



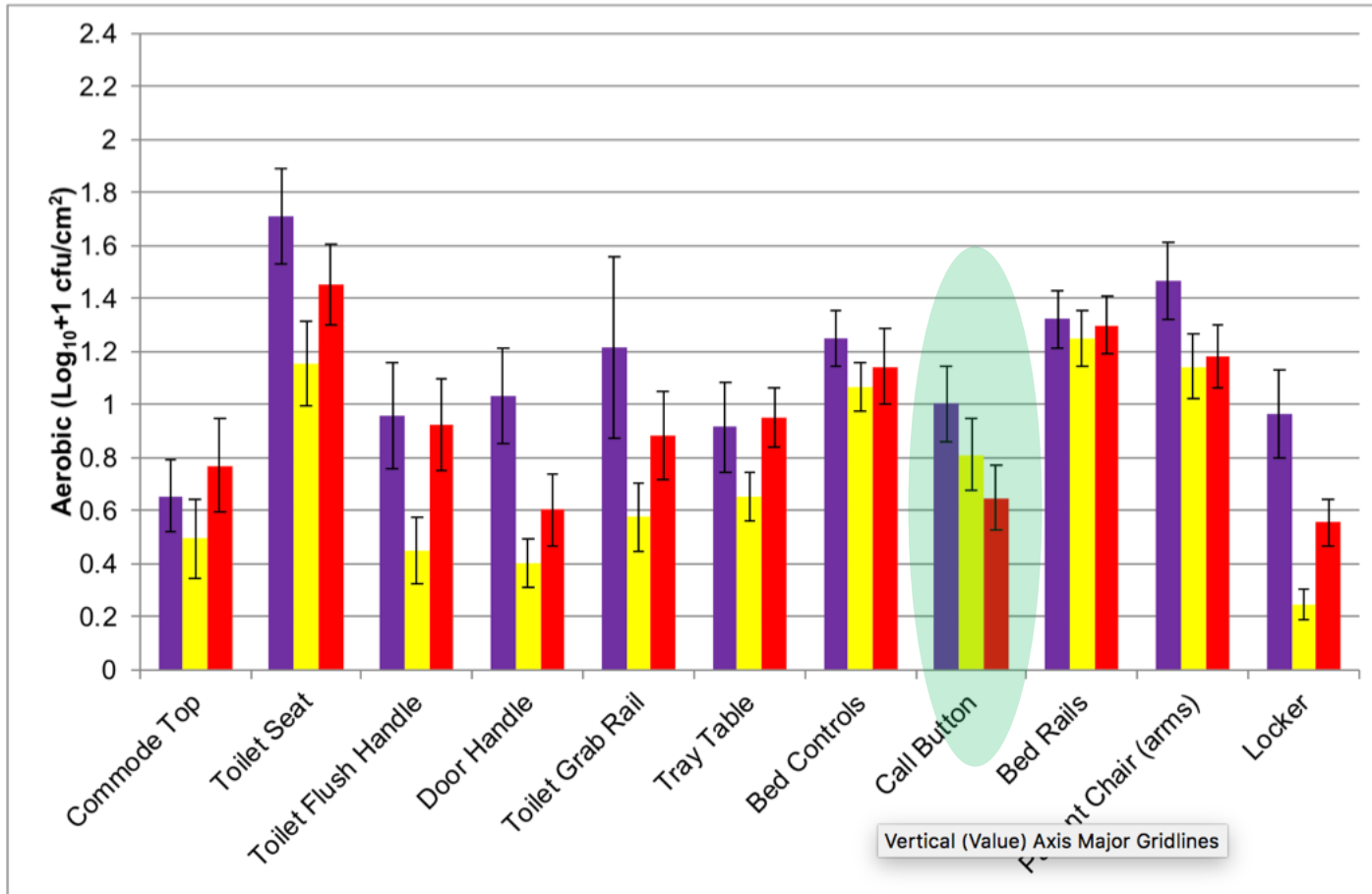
➤ Pre-formulated wipe first followed by cleaning + hypochlorite

- Baseline.
- Cleaning and use of chlorine 1,000 ppm
- Sporicidal wipe

➤ Cleaning + hypochlorite first followed by pre-formulated wipe



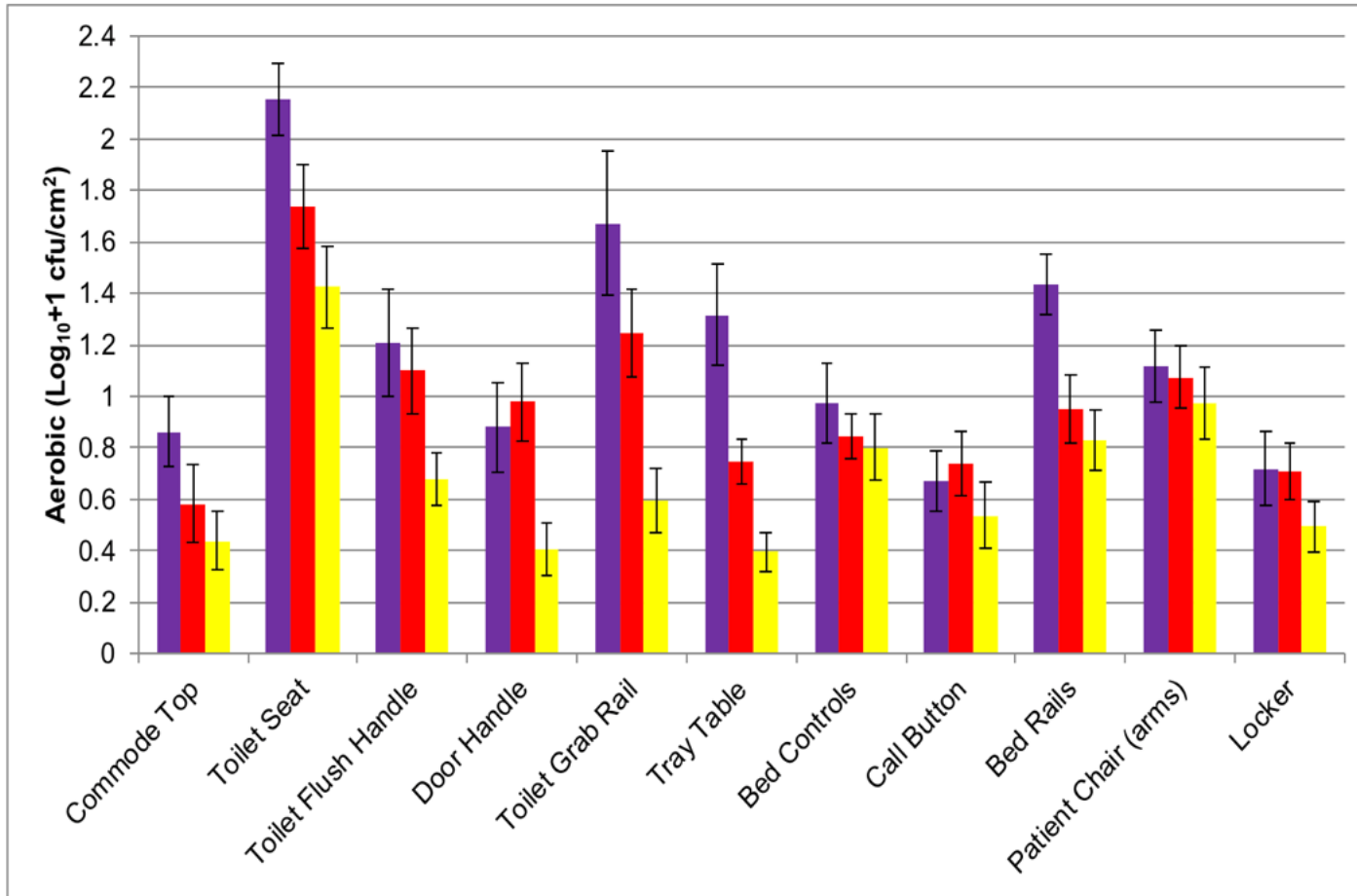
Delivery – preformulated wipe vs. spray & cloth



- Baseline.
- Cleaning and use of chlorine 1,000 ppm
- Sporicidal wipe

➤ Pre-formulated wipe first followed by cleaning + hypochlorite

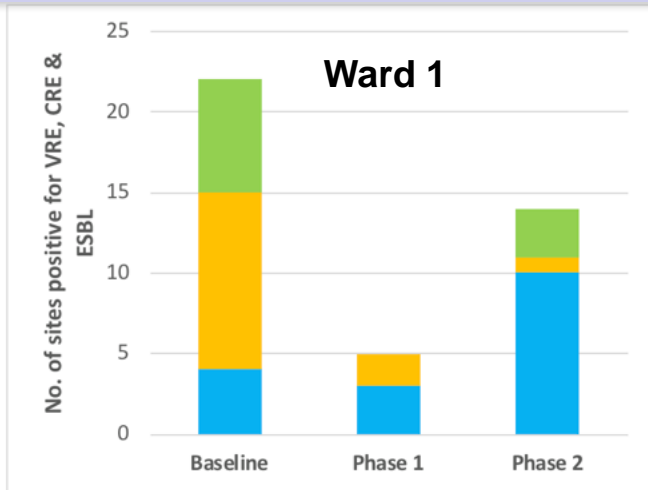
Delivery – preformulated wipe vs. spray & cloth



- Baseline.
- Cleaning and use of chlorine 1,000 ppm
- Sporicidal wipe

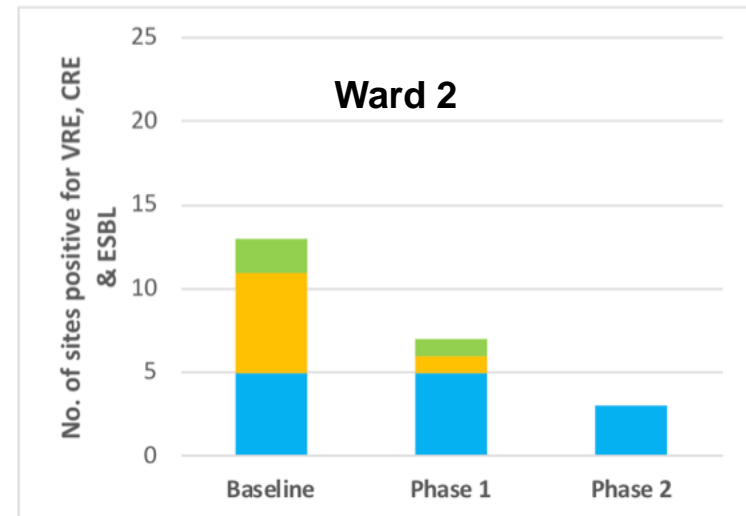
➤ Cleaning + hypochlorite first followed by pre-formulated wipe

Delivery – preformulated wipe vs. spray & cloth



➤ Pre-formulated wipe first followed by cleaning + hypochlorite

- Extended-spectrum beta lactamases (ESBL)
- Carbapenem-resistant Enterobacteriaceae (CRE)
- Vancomycin-resistant enterococci (VRE).



➤ Cleaning + hypochlorite first followed by pre-formulated wipe

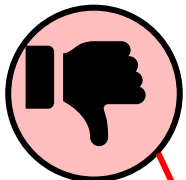
Delivery – preformulated wipe vs. spray & cloth



- Appropriate training & auditing was effective in reducing bacterial bioburden

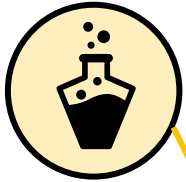


- The use of pre-formulated wipes significantly decreased microbial bioburden from high-touch surfaces compared to the use of cleaning and hypochlorite 1000ppm from a bucket

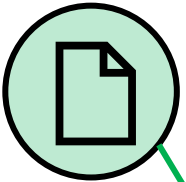


- Aerobic count / site & number of sites with MDRO increased again following the re-introduction of cleaning and hypochlorite 1000ppm from a bucket

Sattar & Maillard (2013) *Amer J Infect Control* 2013;41:S97-S104.

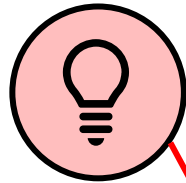


➤ Formulation tests

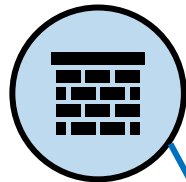


➤ Product test

- EN14776-15: “4-field test”

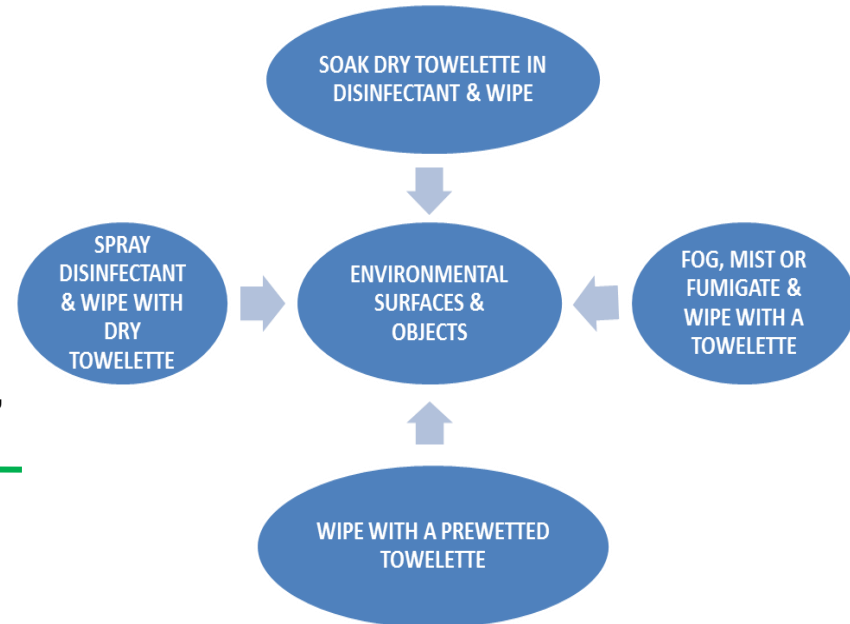


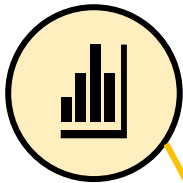
➤ New technology



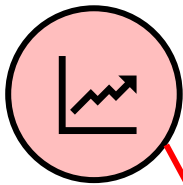
➤ Residual activity

- PAS2424





➤ What concentration left on surface?



➤ Risk associated with selection for/emerging bacterial resistance

TABLE 3 Clinically relevant changes from antibiotic sensitive to resistant according to EUCAST (2020) breakpoint values for *E. coli* before and after a 5-min initial exposure to, and passage in, CHX or broth only

Strain	Initially exposed to CHX at: ^a			
	0.0047 mg/ml, passaged in:		0.0075 mg/ml, passaged in:	
	CHX	Broth	CHX	Broth
<i>E. coli</i> 13P5				
Initial exposure	AMC	— ^b	AMC, FOX	—
Passage 1	AMC	—	AMC	—
Passage 5	AMC	AMC	AMC, IPM ^c	AMC
Passage 10	AMC	AMC	AMC	AMC
<i>E. coli</i> 1B2				
Initial exposure	—	—	AMC	—
Passage 1	—	AMC ^c	AMP ^c	AMC
Passage 5	—	AMP, AMC ^c , CIP, CPD, CF	AMP, AMC, CPD, CF	AMP, AMC, CIP, CPD, CF
Passage 10	CF	AMP, AMC, CPD, CF	AMP, AMC, CPD, CF	AMP, AMC, CPD, CF

^aAMP, ampicillin; AMC, amoxicillin/clavulanic acid; CPD, cefpodoxime; CF, cephalothin; CIP, ciprofloxacin; IMP, imipenem; FOX, ceftazidime.

^b—, no change in antibiotic susceptibility observed.

^cOnly observed in one-half of the repeats.

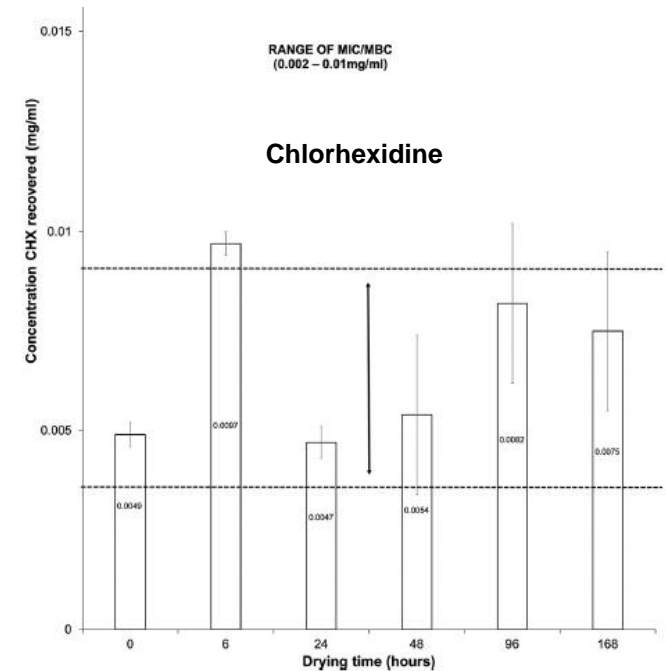


FIG 1 Concentration of CHX (initially set at 20 mg/m) recovered after drying this solution directly on a glass surface. Error bars are standard deviations from the means. Dashed lines depict the range of MIC and MBC values for all *E. coli* isolates tested (see Table S1 in the supplemental material). Note the abscissa distance for the histogram is not proportional to increasing drying time (data based on 3 biological repeats).

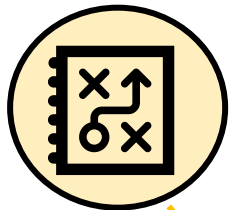


PRODUCT EFFICACY

- MDRO not an issue compared to non-MDRO
- SARS-CoV-2 (generally enveloped viruses) not an issue
- Environmental isolates more resilient

FACTORS AFFECTING EFFICACY

- Predictable for some
- Concentration is paramount
- Formulation & Formulation delivery key
- Cleaning (detergent) will remove but may contribute to pathogen spread



CHALLENGES

- Endospores
- Biofilms - Environmental dry surface biofilms
- Residual activity

THANK YOU



Prof Jean-Yves Maillard
MaillardJ@cardiff.ac.uk