

Less Contact Isolation Is More in the ICU: Not Sure

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Less Contact Isolation is more in the ICU: not Sure. Lucet JC, MD, ¹, Harris AD, MD², Guidet B, MD³ 1Université de Paris, IAME, INSERM, F-75018 Paris, France ; AP-HP, Hôpital Bichat, Infection Control Unit, F-75018 Paris, France 2 University of Maryland. School of Medicine, Baltimore, Maryland 21201 3: Sorbonne Université, INSERM, Institut Pierre Louis d'Epidémiologie et de Santé Publique, AP-HP, Hôpital Saint-Antoine, service de réanimation, F75012, Paris, France Corresponding author: Pr B Guidet Medical ICU Hôpital Saint Antoine 184 rue du Faubourg Saint Antoine 75012 Paris, France bertrand.guidet@aphp.fr Word count: 1080 Ref: 17 Table: 1

The debate between the use of standard (SP) or contact precautions (CP) for stopping the spread of multidrug-resistant organism (MDRO) has been controversial for years (1-5) and still persists despite recent high quality cluster-randomized studies (6-9). SP are based on a universal (also called "horizontal") approach for all patients, whatever they are known as MDRO carriers or not; SP include compliance with hand hygiene and cleaning the environment. CP with a so-called "vertical" approach still include compliance with SP for all patients, additional control barriers for colonized patients, i.e. gloves and gown, and placement in single room if possible. Identification of MDRO carriage through screening is frequently associated with CP.

 The introduction of alcoholic handrub (AHR) in the early 2000's has been a major step for improving compliance with hand hygiene in healthcare setting. Many studies demonstrated that including AHR in a multifaceted strategy, based on education, observation and feedback and other bundled measures are necessary for effective AHR implementation. Since CP and SP both aim at interrupting transmission, SP now including AHR (as compared to handwashing) has higher efficacy, likely closer to that of CP, thus fueling the debate between CP and SP.

 The epidemiology of MDRO is rapidly changing. MDRO comprise Methicillin-resistant *Staphylococcus aureus* (MRSA), extended-spectrum betalactamase-producing enterobacteriacae (ESBL-PE), vancomycin-resistant enterococci (VRE), carbapenemase-producing enterobacteriacae (CPE), and carbapenemase-producing Gram negative bacilli, *Pseudomonas aeruginosa* and *Acinetobacter baumannii* (CRAB). The latter, i.e. CPE, carbapenemase-producing Gram negative bacilli, and VRE in some countries, are classified as extensively-resistant in the late 2010's.

In ICUs facing multiple endemic MDROs, placing a large proportion of patients into CP may result in lower compliance for interrupting cross-transmission from each of these patients (10). Priorities must be defined for selecting the most threatening MDRO, in terms of the individual consequence of infection and the collective risk of dissemination, so relevant infection control practices can be selected.

This complexification of epidemiology and control measures, together with legal mandates issued in several countries may darken the central question for controlling MDRO spread: what is the most effective method to interrupt MDRO cross-transmission? Hospital epidemiologists are often facing difficult choices with CP having some benefit for many preventing MDRO transmission and subsequent infection but also having negatives of cost, environmental waste and healthcare worker dissatisfaction.

 Many factors contribute to the dynamic of MDRO dissemination in the ICU, which should be taken into consideration when designing a policy (Table). Some are of key importance:

- The MDRO under consideration and its local/regional and national epidemiology. Several MDROs can be more easily transmitted suggesting that CP might be more relevant for these bacteria: for example non-E. coli ESBL as compared to ESBL E. coli (4, 11), or ESBL-PE (taken globally) as compared to MRSA (8). Other may have rapid spread, i.e. VRE and CRAB, partially owed to their environmental reservoir; therefore requiring enhanced environmental cleaning;
- Several success stories in controlling MDRO derived from a national policy, strictly enforced by all healthcare facilities, such as CPE in Israel (12) or MRSA in the Northern European countries and recently in the United Kingdom and France (13). Statistical modelling and regional surveillance of VRE and CPE showed that some HCFs may be hotspots for acquisition with subsequent dissemination in many other facilities, thus demonstrating the role of a policy enforced in all healthcare structures (14); Many of these national policies involved the use of active surveillance and CP in their control measures;

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AH is editor for "up-to-date"

Conflict of interest

- This is especially true at the beginning of an epidemic, where extended active surveillance cultures and strict control measures including CP have the highest chance to be effective. By contrast, an endemic situation with high prevalence at ICU admission can only be curtailed, and SP may be preferred in this situation. Importantly, any strategy ideally should be evaluated, by performing admission and discharge screening;

- Compliance with hand hygiene for SP and CP actually is lower than that measured through auditing, due to a Hawthorne effect (15). The first objective in low compliance/low AHR consumption ICUs should be to improve hand hygiene practice, before implementing CP;
- Resources are critical for effective implementation of any precaution, including the availability of single room for CP. In ICU from developing countries, it may be decided first to improve compliance with hand hygiene, while limiting CP to the most aggressive MDROs.

Looking at the parameters to consider from the Table before deciding a control strategy, we suggest that each ICU could implement its own policy, tailored to local epidemiology and resources. Leadership and effective implementation of recommended measures are key for success. For example, two multicenter ICU studies aiming at controlling MRSA were published simultaneously in 2011 (6, 7). They used essentially the same control measures, but resulted in different impact in controlling MDRO. Although both have methodological concerns, the successful intervention was conducted using a behavioral approach, with performance feedback and resolution of local challenges, in addition to technical measures of screening and CP. This illustrates the importance of leadership and HCWs involvement in conducting such intervention, possibly as important as the recommended precautions themselves.

CP and SP do not include measures for cleaning the environment. There is growing evidence that dry surfaces and humid areas may be reservoirs of MDRO. This has been evidenced for a long time for VRE and CRAB, and in a lesser extent MRSA. Recent publications showed that MDR-GNB, including CPE, may persist in humid reservoirs, responsible for outbreak (16). Any strategy, SP or CP, to control MDRO spread should include thorough environmental cleaning.

Although the MDRO epidemiology varies across ICUs, it is of critical importance to adhere to a regional and national strategy. Local situation however, may help to select targeted MDRO where CP has the highest chance to be effective, and other MDROs being tackled with SP. taken into account a maximum number of patients to be placed into CP for higher efficacy.

In this very complex field, there are good arguments from both sides, SP or CP, for

controlling MDRO spread (1, 17). There is however some solid evidence. CP should be

implementing CP may be futile with very high compliance with hand hygiene. Moreover,

enforced only if compliance with HH is high enough, e.g. > 40-50%, otherwise the first

objective should be to improve it. In case of already intermediate or high compliance.

given its potential drawback, CP is probably less relevant for endemic situation. Local

resources should be devoted to the most cost-effective measures based on local

JCL and BG have no COI.

epidemiology, whilst respecting national guidelines

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Table: Circumstances with likelihood of effective standard or contact precautions

Table: Circumstances with I	In favor of standard	In favor of contact	Rationale
	precautions	precautions	
Patient			
Bacterial burden in the source patient	Asymptomatic	Diarrhea, UTI, wounds	The risk of cross-transmission and environmental contamination increases with the bacterial burden of the source patient, making CP and single room potentially more effective than SP
Care dependency, workload	Low	High	Higher workload and care dependency increase the number of contacts with the source patient, and the risk of cross-transmission
Patient's risk for infection	Healthy	Vulnerable	In an ICU with patients at high risk for infection, e.g. transplant ICU or burn ICU, placing colonized patients into CP has a higher chance be effective and reduce the risk for (severe) infection
MDRO Epidemiology			
Local epidemiology	Endemic	Sporadic, outbreaks	It is shown that the higher the number of patients placed into CP, the lower the adherence to control measures
Type of MDRO	Multiply-resistant	Extensively-resistant	CP has higher chance to be effective for some extensively-resistant MDRO in patients with difficult-to treat infections than for patients at risk for infection with less resistant MO. In addition, adherence to a national policy, usually targeting more resistant MDRO, is key for success at the national level
Ease of transmission ("transmissibility")	Lower (ex, ESBL E. coli)	Higher (ex, ESBL non <i>E. coli</i> ; CRAB)	The frequency of transmission depends on the type of MDRO
Route of dissemination	Participation of the antibiotic selective pressure	Mostly by cross- transmission	Emergence of several MDRO variably depends on selective antibiotic pressure, e.g. ampC producing <i>Enterobacteriacae</i> and exposure to 3GC or several resistance mechanisms in <i>P. aeruginosa</i> being induced due to antibiotics. CP may be less useful for patients with these MDRO
HCW practices			
Compliance with hand	High (> 70%) or low (<	Intermediate (40-70%)	High compliance with hand hygiene forms the basis for

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hygiene	40%)		efficacy of SP. But in a situation with a low compliance with hand hygiene, most efforts must target improving compliance before instituting CP. The benefit of CP may be higher in intermediate compliance
AHR consumption of the ICU	High (> 150 mL/Pt.d)	Low (< 100 mL/Pt.d)	AHR consumption is a useful surrogate of hand hygiene in an ICU, additionally to compliance with hand hygiene
Resources			
Environment	Clean, spacious rooms	Crowded	Architectural, human and financial resources are critical to improve compliance with CP
Single room	Limited number	Available	Using single room for all patients in the ICU improves compliance with hand hygiene at room entry and exit, therefore increasing compliance with CP
Screening of patients	Limited	Available	To be effective, active surveillance cultures are required for identifying the whole reservoir of MDRO in patients to be placed into CP
Human workforce	Limited	Available	CP require time for healthcare workers to comply with donning and doffing protective equipment
Financial resources	Limited	Large	CP require financial resources to purchase protective equipment

Adapted from Kirkland K [1]

Abbreviations: UTI, urinary tract infection; CP, contact precautions; SP, standard precautions; ICU, intensive care unit, MDRO, multidrugresistant organism; ESBL, extended-spectrum betalactamase; CRAB, carbapenem-resistant *Acinetobacter baumannii*;