

XXX CONGRESSO NAZIONALE

SIAAIC

Società Italiana di Allergologia,
Asma ed Immunologia Clinica



Scienze e Lettere - Accademia Nazionale dei Lincei



FIRENZE 6/9 APRILE 2017 | WWW.SIAAIC2017.ORG



I PROBIOTICI NELLA RESISTENZA AGLI ANTIBIOTICI:

DANNO O OPPORTUNITÀ

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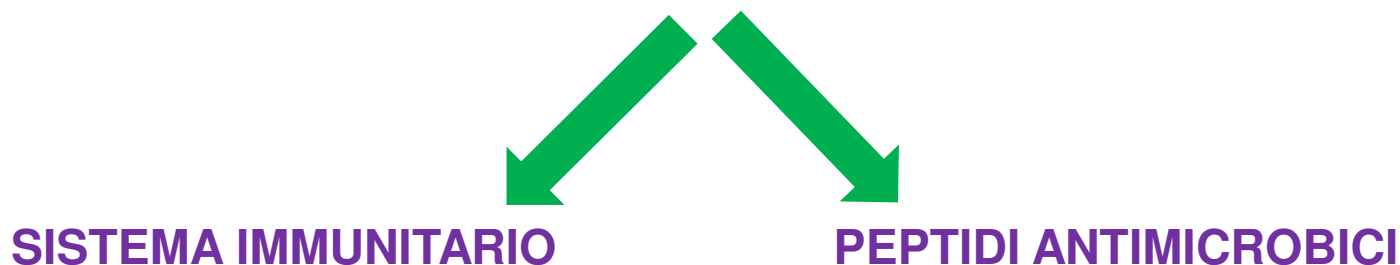


I PROBIOTICI

“ *Microorganismi vivi che se amministrati in adeguate quantità possono portare effetti benefici all’ospite* ”

...NON SOLO...

..anche componenti batteriche quali proteine, acidi nucleici e lipidi sembrerebbero possedere proprietà probiotiche



By FAO/WHO (2002) Expert Consultation <http://www.fao.org/es/ESN/Probio/probio.htm>



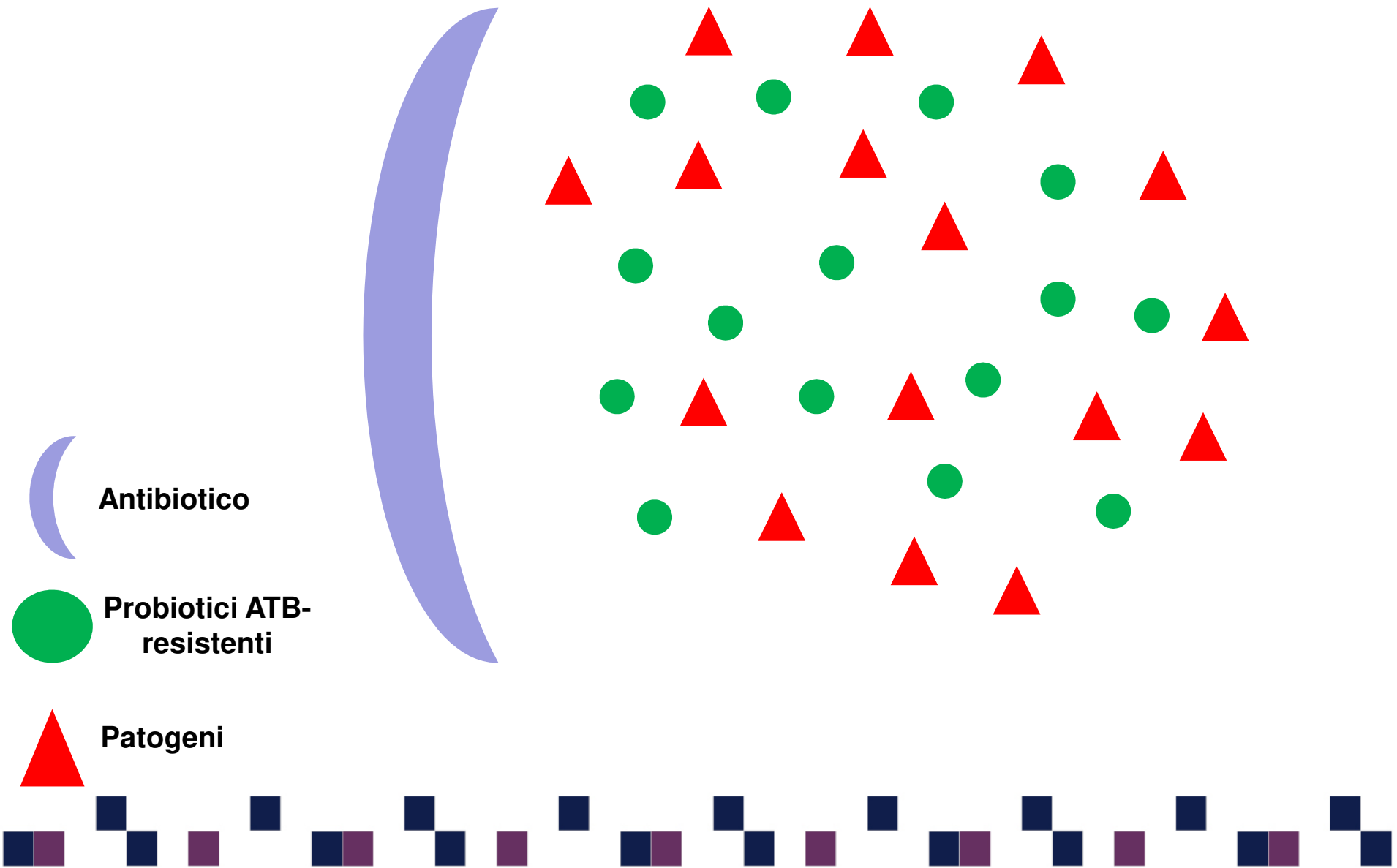


CARATTERISTICHE DEI PROBIOTICI

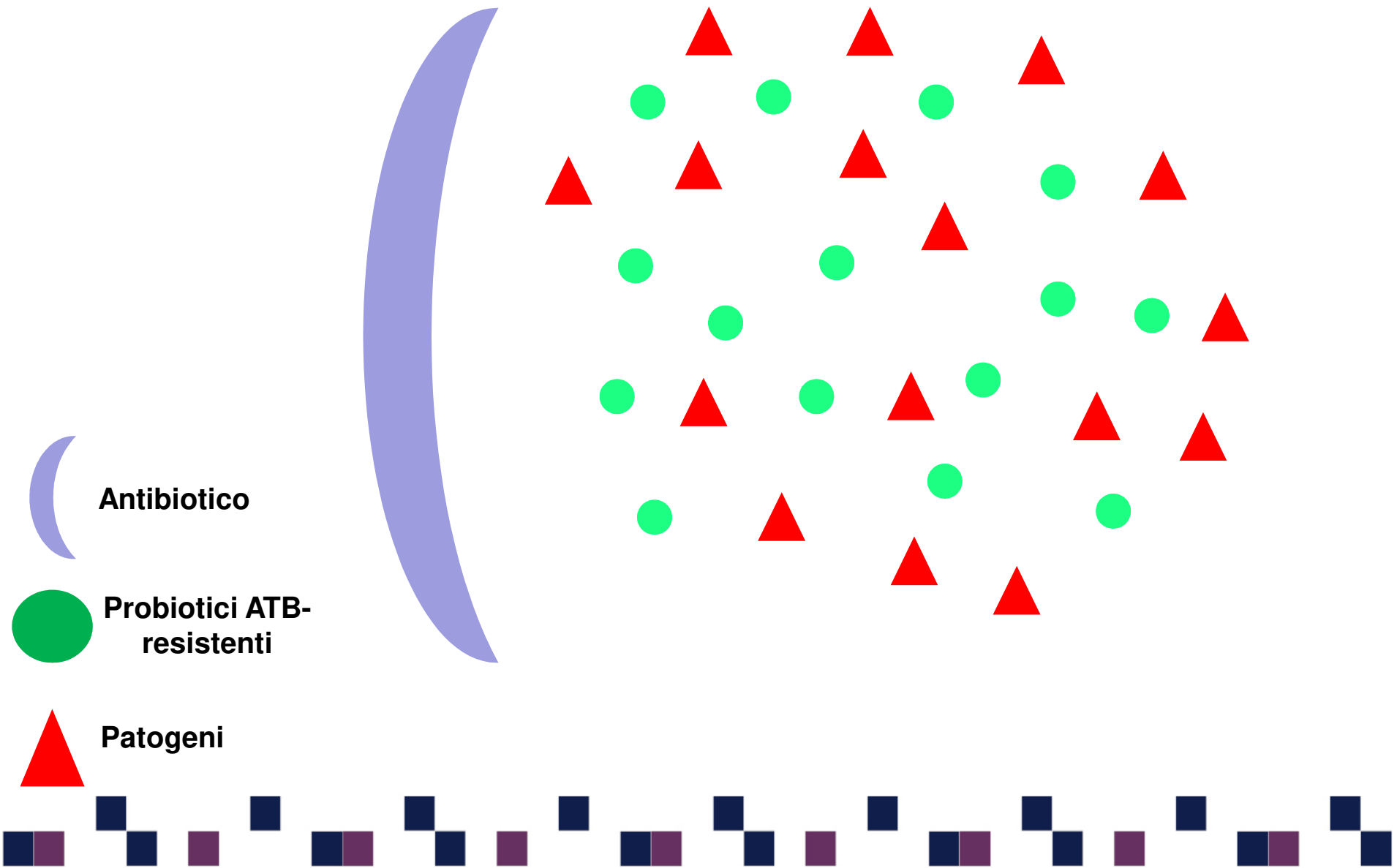
- **Resistenza al pH** gastrointestinale
- **Adesione alle cellule** intestinali
- **Colonizzazione dell'ambiente** intestinale
- **NO** fattori di virulenza
- **Effetti benefici** nell'ospite
- **NO** determinanti genetici di antibiotico-resistenza



PROBIOTICI e ANTIBIOTICO-RESISTENZA



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PROBIOTICI e ANTIBIOTICO-RESISTENZA



European Food Safety Authority

EFSA Journal 20YY;volume(issue):NNNN

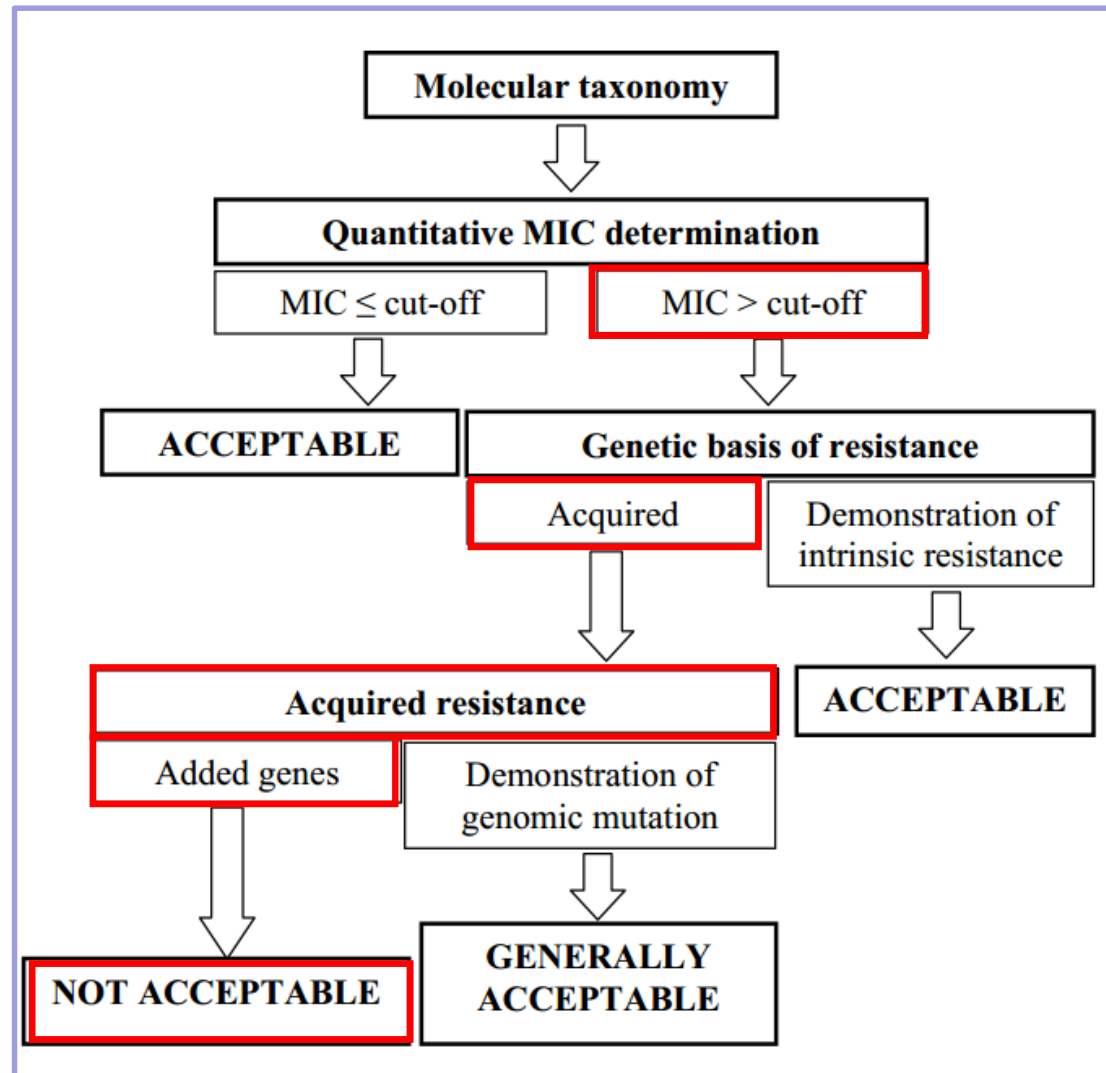
SCIENTIFIC OPINION¹

Guidance on the assessment of bacterial susceptibility to antimicrobials of human and veterinary importance²

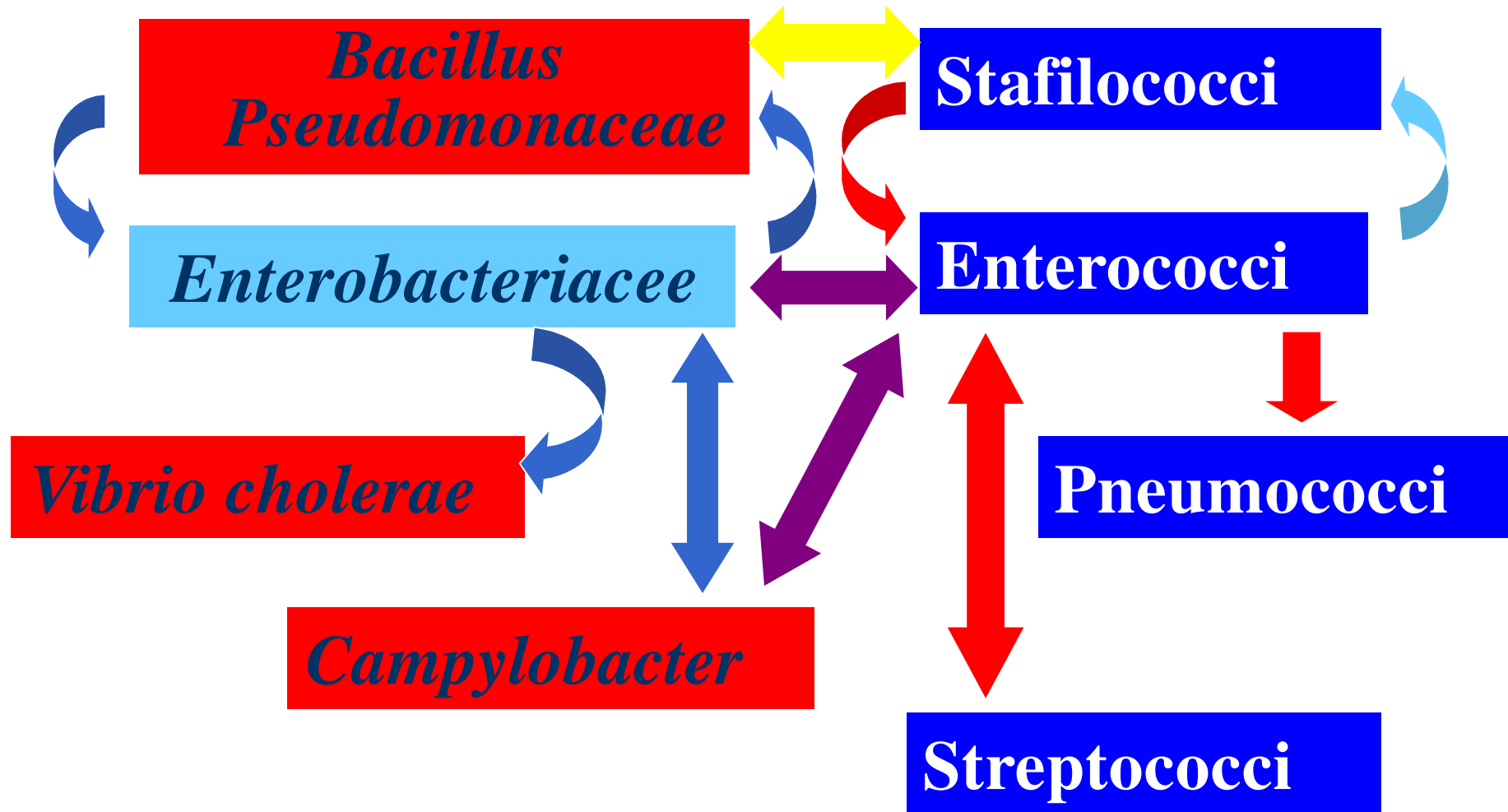
1. CRITERIA FOR IDENTIFYING BACTERIAL STRAINS WITH ACQUIRED RESISTANCE TO ANTIMICROBIALS

All bacterial products intended for use as feed additives must be examined to establish the susceptibility of the component strain(s) to a relevant range of antimicrobials of human and veterinary importance. It is essential that such tests are made in a consistent manner using internationally recognised and standardised methods. As a basic requirement, the minimum inhibitory concentration (MIC) of the antimicrobials expressed as mg/L or µg/mL should be determined for each of the following substances: ampicillin, vancomycin, gentamicin, kanamycin, streptomycin, erythromycin, clindamycin, tetracycline, chloramphenicol and, in specific cases, tylosin, apramycin, nalidixic acid, sulfonamide and trimethoprim (see Table 1). These antimicrobials are chosen to maximise the

PROBIOTICI e ANTIBIOTICO-RESISTENZA



PROBIOTICI e ANTIBIOTICO-RESISTENZA





ANTIBIOTICO-RESISTENZA in ITALIA

- **13** prodotti commerciali; **21** ceppi isolati;
- **12/13 (92%)** prodotti contenevano almeno 1 ceppo resistente ad almeno 1 antibiotico;
- **10/13 (77%)** prodotti presentavano almeno un ceppo contenente geni noti di antibiotico-resistenza.

Drago et al, *Int J Antimicrob Agents* (2013)





ANTIBIOTICO-RESISTENZA in ITALIA

- **21** ceppi isolati;
- **5/21 (24%)** ceppi resistenti all'eritromicina:
 - **2/5** *ermB*;
- **2/21 (9,5%)** ceppi resistenti alla tetraciclina:
 - **2/2** *tetW*; **1/2** *tetS*;
- **0/21 (0%)** ceppi resistenti alla penicillina.

Drago et al, *Int J Antimicrob Agents* (2013)



ANTIBIOTICO-RESISTENZA in ITALIA

| Product name | Species | Erythromycin | Tetracycline | Gentamicin | Penicillin |
|--------------|------------------------|--------------|--------------|---------------|------------|
| A | <i>B. clausii</i> | R (unknown) | S | S | S |
| B | <i>L. casei</i> DG | S | S | R (aph3-III) | S |
| C | <i>L. rhamnosus</i> GG | S | S | R (aadA) | S |
| D | <i>L. reuteri</i> | S | S | R (aac6-aph2) | S |
| E | <i>E. faecium</i> | R (ermB) | S | S | S |
| G | <i>L. paracasei</i> | S | S | R (unknown) | S |
| H | <i>L. delbrueckii</i> | S | S | S | S |
| | <i>B. bifidum</i> | S | S | S | S |
| I | <i>L. paracasei</i> | S | S | R (unknown) | S |
| J | <i>L. rhamnosus</i> | S | S | R (aac6-aph2) | S |
| | | | | (ant6-I) | |
| | <i>L. acidophilus</i> | R (ermB) | S | R (aadA) | S |

ANTIBIOTICO-RESISTENZA in ITALIA

| Product name | Species | Erythromycin | Tetracycline | Gentamicin | Penicillin |
|--------------|----------------------------|--------------|-----------------|------------------------|------------|
| K | <i>E. faecium</i> | S | S | S | S |
| | <i>L. rhamnosus</i> | R (unknown) | R (tetS) (tetW) | R (aadA) | S |
| | <i>L. acidophilus</i> | S | S | R (unknown) | S |
| | <i>Bifidobacterium</i> spp | S | S | R (aph3-III) | S |
| L | <i>L. paracasei</i> | S | S | R (aadA) | S |
| | <i>L. acidophilus</i> | S | S | R (ant6-I) | S |
| | <i>B. lactis</i> BB12 | S | S | R (aph3-III) | S |
| | <i>E. faecium</i> | R (ermB) | S | S | S |
| | <i>B. coagulans</i> | S | S | S | S |
| M | <i>L. paracasei</i> | S | S | R (aadA) | S |
| | <i>L. acidophilus</i> | S | S | R (ant6-I) | S |
| | <i>B. lactis</i> BB12 | S | R (tetW) | R (aph3-III) | S |
| | <i>B. coagulans</i> | S | S | S | S |
| N | <i>L. paracasei</i> | R (unknown) | S | S | S |
| | <i>L. plantarum</i> | R (unknown) | S | R (aph3-III) (aadA) | S |

COME COMPORTARSI nel FUTURO?



Contents lists available at [ScienceDirect](#)

Journal of Global Antimicrobial Resistance

journal homepage: www.elsevier.com/locate/jgar



Susceptibility to antibiotics and genes involved in antibiotic resistance.

| Strain | Antibiotic susceptibility [genes involved] | | | | | | |
|--|--|------------|------------|---------------|--------------|-------------|----------------------|
| | Erythromycin | Gentamicin | Penicillin | Ciprofloxacin | Tetracycline | Clindamycin | Ampicillin/sulbactam |
| <i>Lactobacillus kefir</i> ATCC 8007 (DSM20485) | S | R | S | R | R | S | S |
| <i>Lactobacillus kefir</i> ATCC 35411 (DSM20587) | S | S | S | R | R | S | S |
| <i>Lactobacillus kefir</i> LKF01 (DSM32079) LKEF | S | S | S | R | R | S | S |

S, sensitive; R, resistant.



Ceppo resistente ad alcuni antibiotici ma **NO** geni di ATB-
resistenza su elementi genetici mobili.

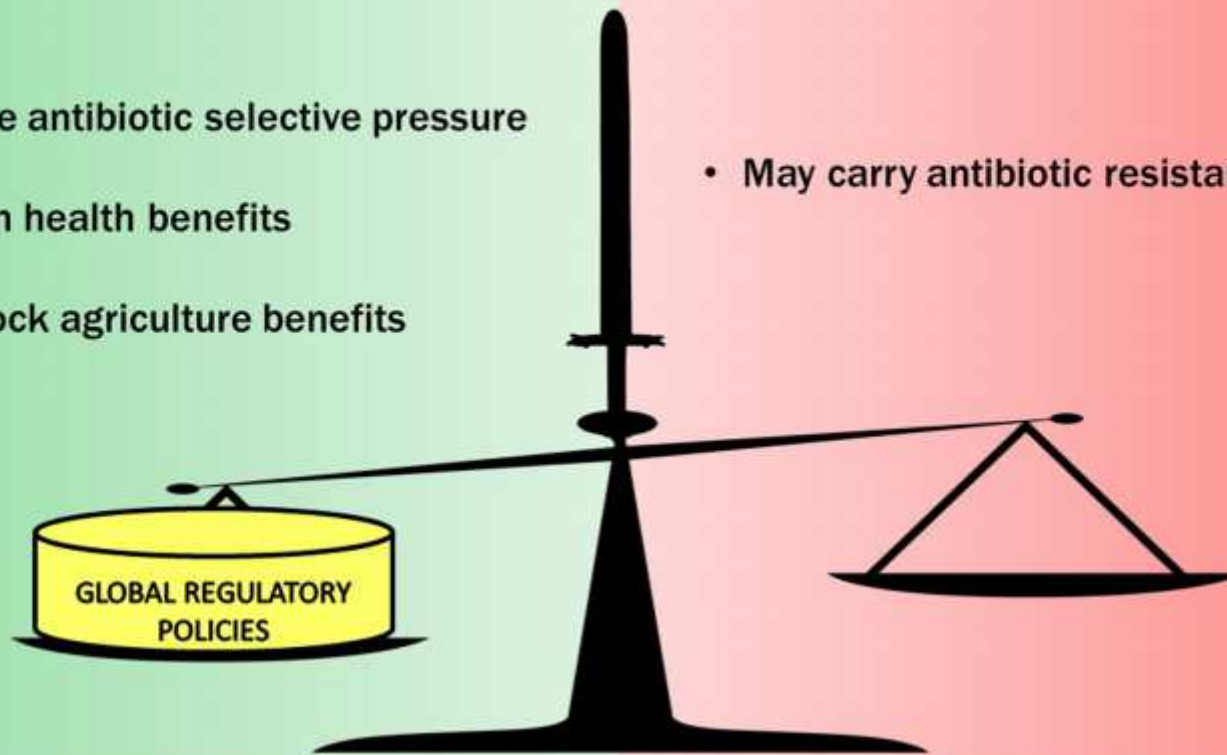
Drago et al., *J Global Antimicrob Res* (2015)

PROBIOTICI e ANTIBIOTICO-RESISTENZA

EFFECTS OF PROBIOTICS

- Reduce antibiotic selective pressure
- Human health benefits
- Livestock agriculture benefits

- May carry antibiotic resistant genes



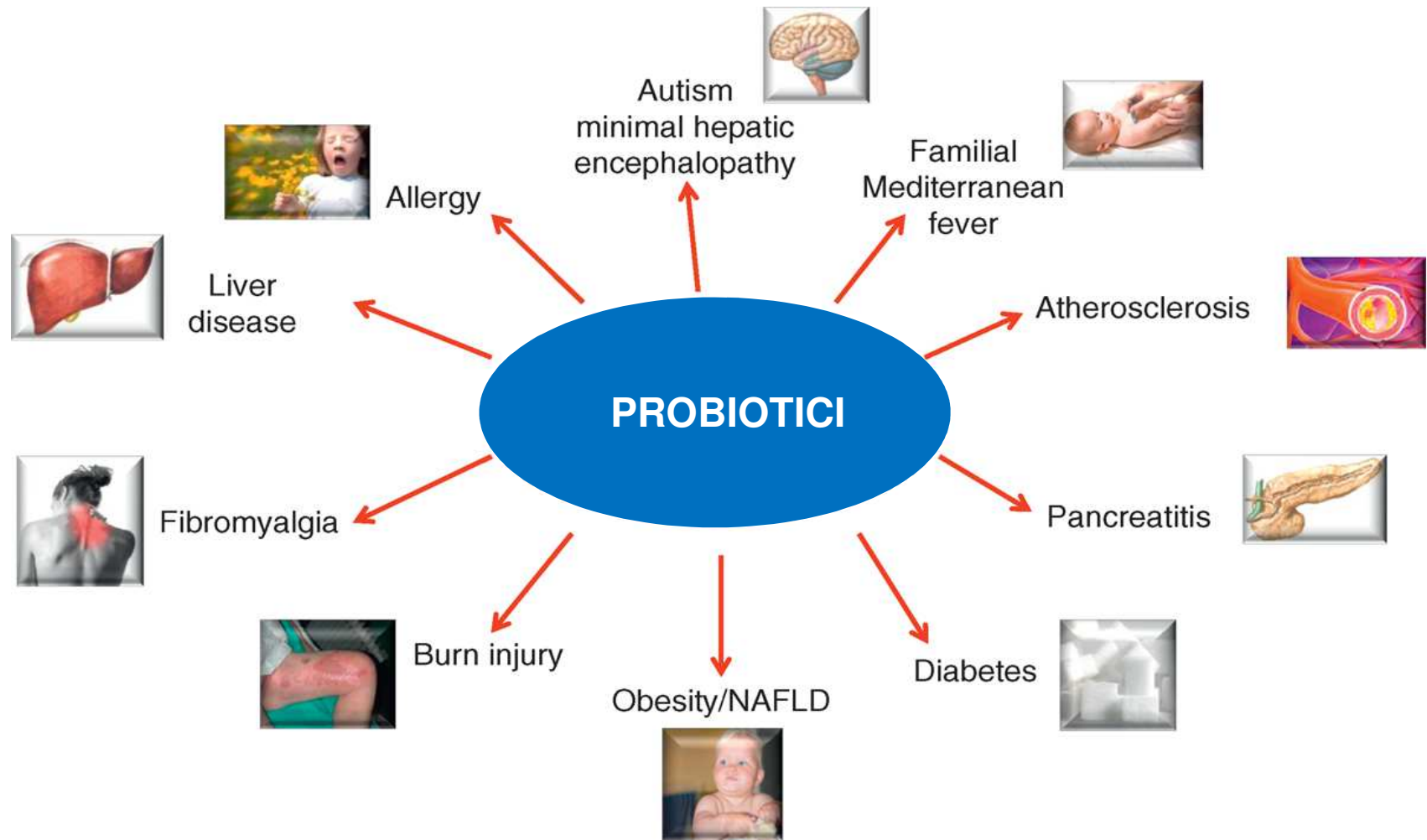
Imperial and Ibana, *Front Microbiol* (2016)

ANTIBIOTICO-RESISTENZA in ITALIA

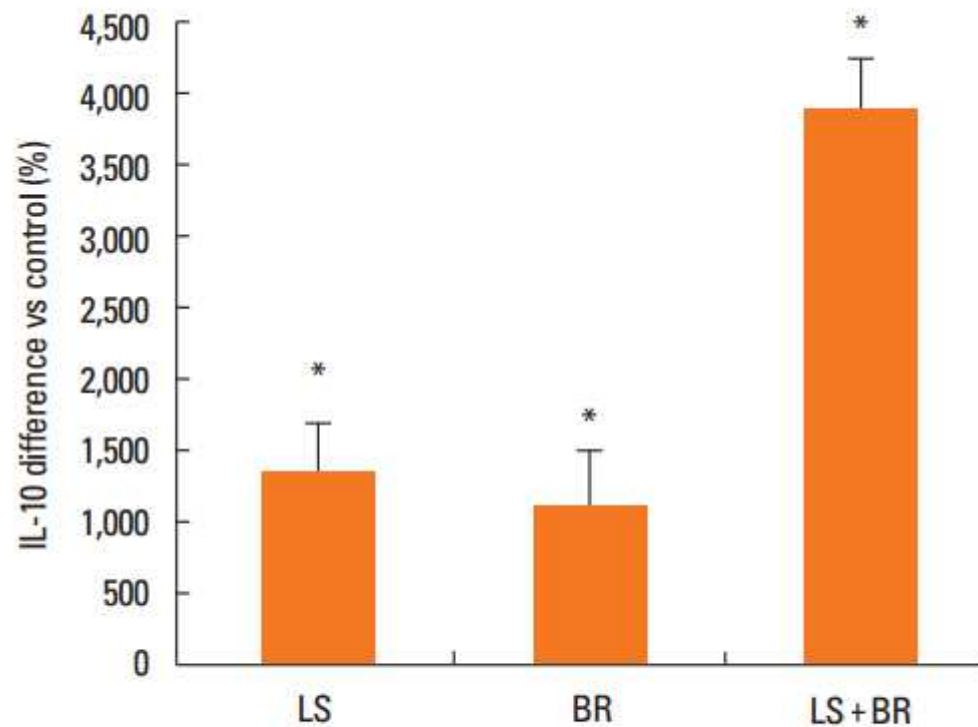
- Reduce antibiotic selective pressure
- Human health benefits
- Livestock agriculture benefits



VANTAGGI dei PROBIOTICI



PROBIOTICI e ALLERGIE



Stimolazione di citochine

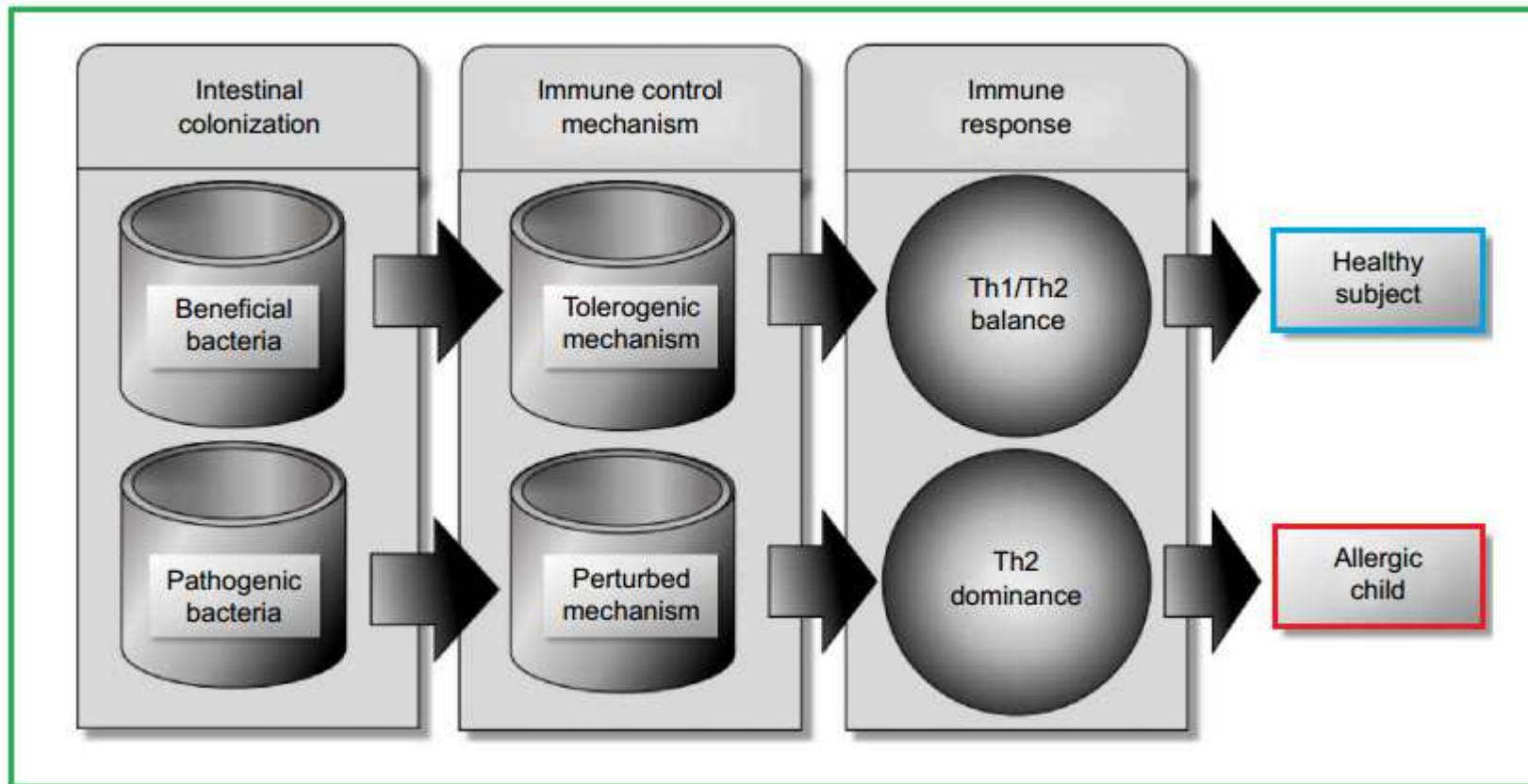
anti-infiammatorie,

riduzione asma allergica.

Release of IL-10 by PBMCs. * $P < 0.05$ vs control (non-stimulated cells).
Data are means \pm SD of the 3 experiments.

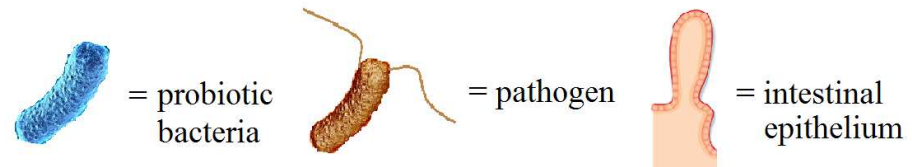
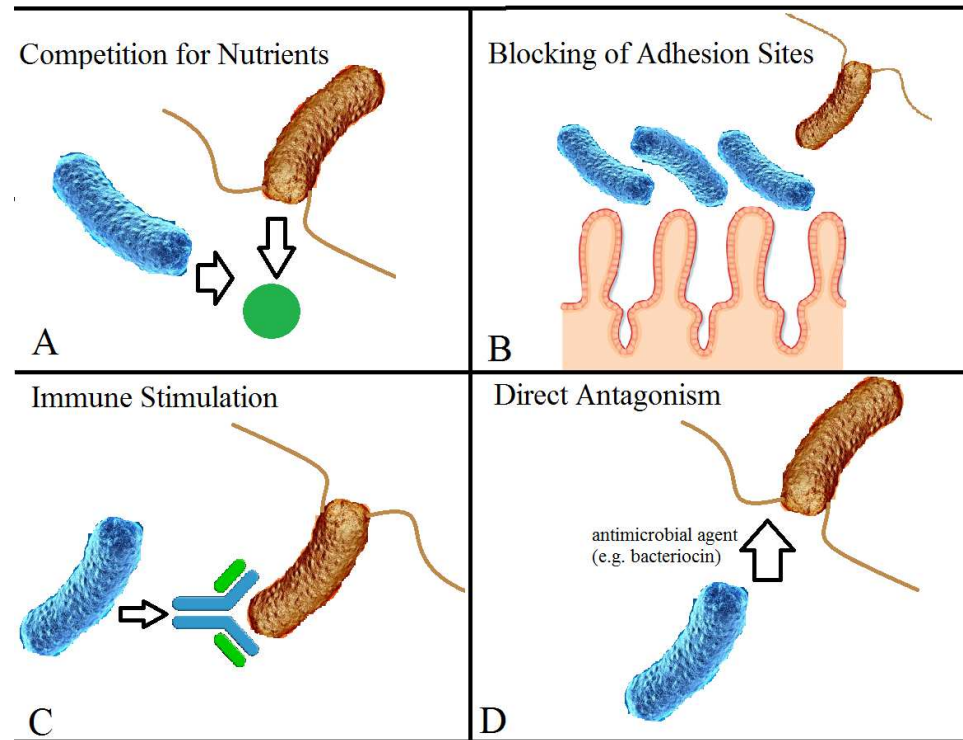
Drago et al, *Allergy Asthma Immunol Res* (2015)

PROBIOTICI e ALLERGIE



PROBIOTICI e INFEZIONI

How Probiotics Work





PROBIOTICI e INFEZIONI

- Benefici nelle infezioni da *Clostridium difficile*;
- Miglioramento delle gastroenteriti acute;
- Riduzione delle complicazioni infettive associate al ricovero in
Terapia Intensiva.
- Riduzione della sintomatologia infezioni gastriche nella popolazione
pediatrica.





PROBIOTICI e INFEZIONI

Ann Med. 2016;48(4):246-55. doi: 10.3109/07853890.2016.1161232. Epub 2016 Mar 26.

Probiotic approach to **prevent** antibiotic resistance.

*Ouwehand AC*¹, *Forssten S*¹, *Hibberd AA*², *Lyra A*¹, *Stahl B*³.

Abstract

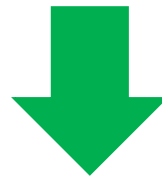
Probiotics are live microorganisms, mainly belonging to the genera *Lactobacillus* and *Bifidobacterium*, although also strain of other species are commercialized, that have a beneficial effect on the host. From the perspective of antibiotic use, probiotics have been observed to reduce the risk of certain infectious disease such as certain types of diarrhea and respiratory tract infection. This may be accompanied with a reduced need of antibiotics for secondary infections. Antibiotics tend to be effective against most common diseases, but increasingly resistance is being observed among pathogens. Probiotics are specifically selected to not contribute to the spread of antibiotic resistance and not carry transferable antibiotic resistance. Concomitant use of probiotics with antibiotics has been observed to reduce the incidence, duration and/or severity of antibiotic-associated diarrhea. This contributes to better adherence to the antibiotic prescription and thereby reduces the evolution of resistance. To what extent probiotics directly reduce the spread of antibiotic resistance is still much under investigation; but maintaining a balanced microbiota during antibiotic use may certainly provide opportunities for reducing the spread of resistances. Key messages Probiotics may reduce the risk for certain infectious diseases and thereby reduce the need for antibiotics. Probiotics may reduce the risk for antibiotic-associated diarrhea Probiotics do not contribute to the spread of antibiotic resistance and may even reduce it.



PROBIOTICI e INFEZIONI



RIDUZIONE INSORGENZA DI PATOGENI ANTIBIOTICO-RESISTENTI



La terapia probiotica riduce il rischio di infezioni secondarie e di conseguenza, il bisogno di antibiotici.

PROBIOTICI e INFEZIONI

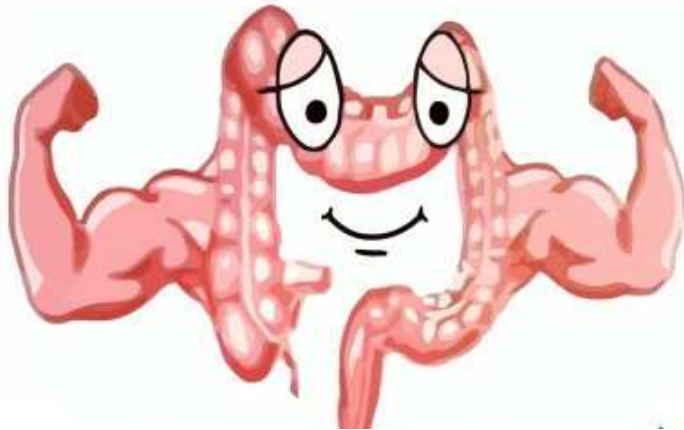


BARRIERA CONTRO I PATOGENI

RAFFORZAMENTO SISTEMA IMMUNITARIO

MIGLIORAMENTO GENERALE DELLA SALUTE

CONCLUSIONI



Grazie per l'Attenzione!

