

CREATING A WWW COLLECTION OF THERAPEUTIC BACTERIOPHAGES ACTIVE AGAINST GLOBAL BACTERIAL INFECTIONS

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Bacteriophages are natural predators of bacteria also those pathogenic to humans. Phages outnumber bacteria at least tenfold and are the most abundant life form on the Earth. Typically phages are highly specific and demonstrate narrow host ranges of their lytic activity restricted to single bacterial clones. Phages together with the sensitive bacteria form predator-prey pairs in evolutionary arms race that tends to keep the populations of both species in unbalanced equilibrium. Thus phages and bacteria must have a history of previous contacts and interactions and physically close to each other. Accordingly phages isolated in certain places may not be active against bacterial strains isolated from the other locations simply due to the fact that they do not have such a history of previous interactions. On the other hand, if bacteria from one part of the globe are successfully attacked by the phages isolated from the other part, this means either a global dissemination of the bacterial-phage pair or an extremely wide phage host range due to commonly used receptors.

We have tested the sensitivity of 65 clinical bacterial pathogens including MRSA, multiple resistant *Pseudomonas aeruginosa*, *Acinetobacter baumannii* and *Klebsiella pneumoniae*, carbapenem resistant *Escherichia coli* and *Enterobacter cloacae*, as well as vancomycin resistant *Enterococcus faecium* isolated in Singapore hospitals against a collection of the corresponding resident bacteriophages isolated from the waste waters of Munich. Although some species like MRSA and *Pseudomonas* demonstrated 80% and 90% sensitivity to the locally isolated phage preparations, respectively; 80% *Kl. pneumoniae* and 80% *E. coli* were resistant to the corresponding phages. Moreover all tested *Acinetobacter* and *E. cloacae* strains were resistant to the phages previously isolated in Germany.

We have used different water samples from Munich waste water collectors to isolate phages active against the phage-resistant bacteria isolated in Singapore. We have successfully isolated lytic phages to the representatives of each group of the pathogenic bacteria. The phages demonstrated different host ranges and activities. The DNAs of the collected lytic bacteriophages were NGS sequenced, their genomes assembled and compared to the phage DNAs presented in the available public databases. Such detailed characterization of the therapeutic bacteriophages allow us covering both potential threats present in the phage genomes and epidemiological and evolutionary linkages of these abundant life forms.

Thus, Munich waste waters already contain phages active against bacterial cultures isolated in Singapore hospitals that make it possible to establish a representative well characterized “world-wide” collection of therapeutic bacteriophages active against both locally and globally disseminated pathogenic microorganisms.