| Diversification of vibrio anguillarum driven by the bacteriophage CHOED                                 |
|---------------------------------------------------------------------------------------------------------|
| León M.                                                                                                 |
| Kokkari C.                                                                                              |
| García K.                                                                                               |
| Castillo D.                                                                                             |
| Katharios P.                                                                                            |
| Bastías R.                                                                                              |
| Bacteriophages are an important factor in bacterial evolution. Some reports suggest that lytic          |
| bacteriophages can select for resistant mutant strains with reduced virulence. The present study        |
| explores the role of the CHOED bacteriophage in the diversification and virulence of its host Vibrio    |
| anguillarum. Nine phage-resistant strains were analyzed for their phenotype and different virulence     |
| factors, showing alterations in their fitness, motility, biofilm formation, lipopolysaccharide profiles |
| and/or protease activity. Seven of the nine phage-resistant strains showed virulence reduction in a     |
| Sparus aurata larvae model. However, this is not generalized since two of the resistant strains show    |
| equal virulence compared with the parental strain. The genomic analysis of representative resistant     |
| strains displayed that the majority of the mutations are specific for each isolate, affecting genes     |
| related to lipopolysaccharide biosynthesis, quorum sensing, motility, toxin and membrane transport.     |
| The observed mutations were coherent with the phenotypic and virulence differences observed.            |
| These results suggest that the CHOED phage acts as a selective pressure on V. anguillarum,              |
| allowing proliferation of resistant strains with different genotypes, phenotypes and degrees of         |
| virulence, contributing to bacterial diversification. © 2007 - 2019 Frontiers Media S.A. All Rights     |
| Reserved.                                                                                               |
| Bacterial evolution                                                                                     |
| Bacteriophage                                                                                           |
| Bacteriophage resistance                                                                                |
| Diversification                                                                                         |

| Vibrio anguillarum       |
|--------------------------|
| Virulence                |
| bacterial DNA            |
| bacterial toxin          |
| lipopolysaccharide       |
| proteinase               |
| siderophore              |
| virulence factor         |
| Article                  |
| bacterial growth         |
| bacterial strain         |
| bacterial virulence      |
| bacteriophage            |
| bacterium identification |
| bacterium isolation      |
| cell motility            |
| cell proliferation       |
| DNA extraction           |
| DNA sequence             |
| electron microscopy      |
| fitness                  |
| genetic identification   |
| genotype                 |
| growth curve             |
| hemolysis                |
| ee.ye.e                  |

nonhuman

phenotype

quorum sensing

Sparus aurata

Vibrio anguillarum