

# Antibiotic Resistance (AR) Among Enteric Bacteria in Marine Environments of Marmara Sea



**Nüket SIVRI\***

*Istanbul University, Environmental Engineering Department, Turkey*

**Submission:** August 30, 2017; **Published:** October 20, 2017

**\*Corresponding author:** Nüket SIVRI, Istanbul University, Environmental Engineering Department, Engineering Faculty, 34320, Istanbul, Turkey, Tel: +90212737070; Fax: +902124737180; Email: [nuket@istanbul.edu.tr](mailto:nuket@istanbul.edu.tr)

## Abstract

Recently, antibiotic resistance has been identified as “the most important and the most urgent global danger” by United Nations. The subject of how and when the life-saver antibiotics have become the seven hundred thousand peoples’ cause of death has been on the United Nations’ agenda. Extraordinary meetings, political declarations and fast campaign decisions have shown the importance of the subject, the antibiotic resistance [1,2]. Each country had listed what they should do for their own and took action. The priority was the studies about determining the type of resistance and mechanisms about it. After that, the theme of raising awareness of community about rational usage of antibiotics became more important. The main topics aimed to be completed were also took shape for Turkey, an active attendee of the meetings. This mini review aims to address the need for a greater understanding of the role of the marine environment and host micro biome in influencing the evolution, acquisition and spread of antibacterial resistance, and acting as a reservoir for resistance, not only for Turkey but also for other countries.

**Keywords:** Antibiotic resistance; AR; Enteric bacteria; *Escherichia coli*; Marmara Sea; Turkey

## Introduction

Inappropriate and excessive consumption of one of the most important inventions in human history, antibiotics, with an increased rate, has resulted with the problem of antibiotic resistance (AR) [3]. Microorganisms are the oldest creatures of the Earth with a skill of fast adaptation for ever changing status of the environment. With that skill, sooner or later they develop an ability to combat with antimicrobial agents used for destroying them, the “resistance” [4]. This strategy, evolution of bacterial antibiotic resistances, had caused the resistance to become very important, life-threatening problem for all humanity [5]. Nowadays the rate of resistance is rapidly increasing not only for hospital-based cultures but also cultures taken from the environment, which enlarges the problem and makes it even more serious [6-8].

In International meetings, in order to solve the AR problem, the importance of supporting the projects is highlighted. Great deal of countries are supporting the cause and allocating the necessary budget in medical researches [7]. For this purpose, Research Councils like UCL, NERC, BBSRC and MRC, which help researchers willing to take part about the subject from other countries, are leading the projects with attendance of different countries with a budget of £6.5m. Four main topics have been defined to study about in these projects.

1. Understanding resistant bacteria.
2. Accelerating therapeutics and diagnostics development.
3. Understanding the real world interactions.
4. Behaviour within and beyond the health care setting.

As similar project calls, this programme also aims to address the need for a greater understanding of the role of the outdoor environment and host micro biome in influencing the evolution, acquisition and spread of antibacterial resistance and acting as a reservoir for resistance. The programme is restricted to antibacterial and resistant bacteria or bacterial resistance genes, of clinical and/or veterinary importance. Researches supported can be based in agriculture, aquaculture, wastewater and natural environments (and their interfaces), the human and animal host micro biome.

## Result and Discussion

Marine areas are special areas preferred for resting and recreational purposes. But unfortunately these special areas are also influenced by residential, industrial and ship wastes. Turkey is a rich country in marine areas, especially with Marmara Sea, which has a special attribute for being an inner-sea. However, in last 20 years, it gained a feature which itself feels and also shows

the pressure of increased infrastructure and intense pollution [9] 40-50 species that had been fished for commercial purposes have regressed to 20-25 species; annual fishing quotas have reduced with same rate, and most of the species can no longer be seen, yet hunted [10].

In addition, the Marmara Sea which suffered a lot in the subject of biodiversity has started to show changes in its micro flora, because of infrastructural problems and changes in coastal area usage awareness [11]. Even though in aquatic environments bacteria are also a natural part of the ecosystem [12] the enteric bacteria found in areas with residential wastes and heavy pollution represent dense fecal contamination. In recent years, just like the megacities of other countries, the most important city of Marmara Sea, İstanbul, is feeling the anthropogenic pressure amply. At the same time the other neighbours of Marmara Sea in İzmit and Bursa, which are the industrialization centres and in Yalova, which is the centre of shipyards are under a similar pressure. The reason for changes we face in flora of Marmara Sea can be explained with anthropogenic sourced wastewaters reaching the sea and excellent adaptation of bacteria to the environment. In Turkey, maximum and minimum limits for all the parameters necessary and recommended values for all water sources (excluding open sea water) are regulated in Surface Water Quality Regulation [13]. For "microbiologic water quality", accepted bacteriologic parameters are defined as "Total coliform" and "Fecal coliform", by Environmental Protection Agency (EPA) and World Health Organization (WHO). Maximum limits calculated by "Membrane Filtration (MF)" method are in limitations of >10000/100ml and >200/100ml respectively, in Turkey. But according to WHO, USEPA and WRC especially in aquatic areas used for recreational purposes, fecal coliform bacteria value should not exceed 200/100ml [14]. Unfortunately, in some coastal areas, particularly in ones without proper wastewater treatment systems and wastes reaching the sea, these values may reach up to 10<sup>6</sup>.

In the list of megacities of the world, İstanbul is in the 25<sup>th</sup> place with approximately 18% of Turkey population. In İstanbul, "problems based on environmental pollution and inadequacy of recreational areas and public social life areas" is the 5<sup>th</sup> most important problem being faced. To prevent the problems in place, the İstanbul coastal area is being monitored via testing the samples taken from at least 40 different stations monthly and for longer than 15 years [15,16]. Miniaturization is not being done only by universities, but also by authorized government units with care. In studies of determination of enteric bacteria in İstanbul's southwest shore surface water, it has been found that present bacterial pollution levels for last 8 years are exceeding aquaculture, fishing and recreational levels designated, frequently [17,18]. In molecular studies for determination of pathogenic existence, especially Gram negative pathogen species, *Escherichia coli*, *Enterobacter sp.*, *Klebsiella oxytoca*, *Pseudomonas aeruginosa* had been found dominant in Marmara Sea [19,20]. This prominent situation threatening

both the ecosystem and public health may cause a different risk with addition of antibiotic resistant bacteria existence in place [21]. These bacteria with different antibiotic resistant genes are foreseen to spread on surface waters, deep waters, in sediment, in other words across all marine ecosystem [22,23].

In studies for molecular characterization and antibiotic resistance for same areas in residential wastes, bacteria from human intestinal flora with R-plasmid found present. R-plasmid in these bacteria, which makes them resistant to antibiotics, can easily spread to environment because of wastewaters mixing with environmental waters [24,25]. In studies of Marmara Sea coastal areas, 10 chosen antibiotics (ampicillin, amoxicillin, amikacin, streptomycin, nalidixic acid, trimethoprim-sulfamethoxazole, tetracycline, chloramphenicol, imipenem, ceftazidime) resistance percentages were investigated. The results point out the high density of especially beta-lactam type antibiotic resistant bacteria. With changes yearly, the highest resistance percentages were identified as, in order; ampicillin (%83.1), amoxicillin (%57.8), tetracycline (%50.7), trimethoprim-sulfamethoxazole (%42.2), nalidixic acid (%38), ceftazidime and streptomycin (%16.9), chloramphenicol (%14.1) [26,27]. In sea water samples of İzmit Gulf, these ratios were found as tetracycline %50, sulbactam/ampicillin %62.5, penicillin %62.5, gentamicin %50, amikacin %12.5, chloramphenicol %37.5, cephoperazone %25, kanamycin %37.5, trimethoprim/sulfamethoxazole %62.5 [28].

For studies accomplished in Turkish seas, it is possible to find similar percentages to Marmara Sea. In a study of sea water sourced coliform class 1 and class 2 integron gene cassettes and antibiotic resistance characterization in East Coast of Black Sea, resistance percentages were tetracycline (%23.2), ampicillin (%20), sulfamethoxazole (%11.6), streptomycin (%9.3), chloramphenicol (%4.6), trimethoprim (%2.3), respectively [29]. In a study of Iskenderun Gulf area, in isolated Gram negative bacteria strains, resistance percentages were, in order; ampicillin (%93.2), streptomycin (%90.2), cefazolin (%81.3), imipenem (%16.5), meropenem (%13.9), cefepime (%8), respectively [27].

Similar to seas of other countries, high resistance values and resistance genes which can be transferred by mobile elements like plasmid, transposon, integron have been detected in Turkey [28]. But intriguingly, in isolated bacteria strains coming from the samples of ship ballast tanks, cruising in most of our seas and going into and coming from Marmara Sea, high resistance to antibiotics were also identified [29,30]. This shows that the studies about the antibiotic resistance are necessary not only in medical practice but also in surface and sea waters and even in international transit ships.

### Conclusion

Apparently, common usage of antibiotics for human and animal health causes a specific pressure on bacteria, which

results with emergence and spread of antibiotic resistance. Antibiotic resistant bacteria, which are the food source for some of the organisms, can easily reach seas via rivers, streams and/or other kind of aquatic systems and spread their resistance genes to aquatic ecosystems.

Antibiotics indispensable for treatment, coming against us as “the most important and most urgent global danger: antibiotic resistance problem” in recent times. In order to solve this issue, preventing irrational use of medications, raising national awareness campaigns and creating “Rational Antibiotic Usage” educations are necessary. But in the same time, it should be known that antibiotic resistant bacteria may reach natural waters and seas, and can spread their resistance genes. In addition to successful national campaigns, inclusion of national and international environmental projects and studies are necessary for solution aims of this global problem. In this subject, Turkey continues on awareness campaigns and research projects as a strong leader in its region.

### Acknowledgement

The author is grateful to Dr. M. Kutay SIVRI for providing the scientific infrastructure required to carry out the paper. Author would like to thank Mr. Oğuz Kuzu for his critical reading and quick editing. The results part of this work is supported by the Research Fund of the Istanbul University (Project number : FYO-2016-3735)”. The author has declared no conflict of interest.

### References

1. Mc Kenna M (2016) In First, UN will consider antibiotic resistance. National Geographic, USA.
2. Mc Kenna M (2016) A third of people given antibiotics don't need them. National Geographic, USA.
3. Baggs J, Fridkin SK, Pollack LA, Srinivasan A, Jernigan JA, et al. (2016) Estimating national trends in inpatient antibiotic use among US hospitals from 2006 to 2012. *JAMA Intern Med* 176(11): 1639-1648.
4. Quintiliani R, Sahm DF, Courvalin P, Murray PR, Jo Baron E, et al. (1999) Mechanisms of resistance to antimicrobial agents. *Manual of Clinical Microbiology*, (7<sup>th</sup> edn), ASM Press, Washington, DC, USA, pp. 1505-1525.
5. Hawkey PM (2008) The growing burden of antimicrobial resistance. *J Antimicrob Chemother* 62(Suppl 1): i1-i9.
6. Kim TW, Joung Y, Han JH, Jung W, Kim SB, et al. (2015) Antibiotic resistance among aquatic bacteria in natural freshwater environments of Korea. *J Water Health* 13(4): 1085-1097.
7. WHO (2016) Antimicrobial resistance.
8. Jozić S, Šolić M (2017) Effect of environmental conditions on *Escherichia coli* survival in seawater, Chapter 9, *Escherichia coli* recent advances on physiology, Pathogenesis and Biotechnological Applications, pp. 169-186.
9. Sivri N, Akbulut V (2016) Antimicrobial susceptibility of *Escherichia coli* strains collected from the south western coast of Istanbul. *Biosciences Biotechnology Research Asia* 13(2): 785-793.
10. Çelikkale M.S. & Sivri N. (2016) «Değişen Marmara Denizi ve Değişen Su Ürünleri», Şehir ve Toplum, (4) 19-27.
11. Öztürk R (2002) Antimikrobik İlaçlara Karşı Direnç Gelişme Mekanizmaları ve Günümüzde Direnç Durumu, Akılcı Antibiyotik Kullanımı ve Erişkinde Toplumdan Edinilmiş Enfeksiyonlar Sempozyum Dizisi, İÜ. Cerrahpaşa Tıp Fakültesi Sürekli Tıp Eğitimi Etkinlikleri 31: 83-100.
12. Robinson JB, Tuovinen OH (1984) Mechanism of microbial resistance and detoxification of mercury and organomercury compounds: Physiological, Biochemical, and Genetic Analyses. *Microbiol Rev* 48(2): 95-124.
13. (2016) Turkish Surface Water Quality Management Regulation (TSWQMR).
14. Oram B (2014) Bacteria in the environment and drinking water. Water Research Center.
15. Sivri N, Seker DZ (2010) Investigation of Enteric Bacteria of surface waters in the southwestern istanbul coast by means of GIS Turkish. *Journal of Fisheries and Aquatic Sciences (TrJFAS)* 10(4): 505-511.
16. Sivri N, Sislı NS, Algur D, Akbulut V (2011) Multiple antibiotic resistance indexing of *Escherichia coli* in the southwest coast of Istanbul. marine ecosystems & the issue of marine natural products and their bioactive metabolites, Bogor, Indonesia.
17. Sivri N, Şeker DZ, Şapçı Z, Üstün B (2011) Monitoring of the enteric bacteria of the Küçükçekmece Lagoon (Istanbul) by GIS. marine ecosystems& the issue of marine natural products and their bioactive metabolites, Bogor, Indonesia.
18. Sivri N, Allen MJ, Jones M, Akbulut V (2014) Potential public health significance of faecal contamination in south-western coastal area in Istanbul , Turkey. *Journal of Pure and Applied Microbiology (JPAM)* 8(5): 3789-3796.
19. Sivri N, Balcı M, Balkis N, Jones M, Allen MJ, et al. (2013) Detection of *Escherichia coli* with UidA Gene in marine environment of Kapıdağ Peninsula (Marmara Sea)”, 40<sup>th</sup> CIESM Congress, Marseille, France.
20. Sivri N, Jones M, Allen M (2014) *Pseudomonas aeruginosa* isolated from seawater the marine environments in the Istanbul coastal area. *Fresenius Environmental Bulletin (FEB)* 23(12): 3340-3344.
21. Çelebi H, Sponza D (2007) Antibiyotiklerin çevresel etkileri, toksisiteleri ve anaerobik aritilabilirlikleri 7. Ulusal Çevre Mühendisliği Kongresi, Ekim, İzmir, pp. 24-27.
22. Sivri N, Sandallı C, Özgumus OB, Colakoglu F, Dogan D, et al. (2012) Antibiotic resistance profiles of enteric bacteria isolated from Küçükçekmece lagoon (Istanbul-Turkey). *Turkish Journal of Fisheries and Aquatic Sciences (TRJFAS)* 12: 699-707.
23. Sandallı C, Özgümüş OB, Sevim A (2010) Characterization of tetracycline resistance genes in tetracycline-resistant Enterobacteriaceae obtained from a coliform collection. *World J Microbiol Biotechnol* 26(11): 2099-2103.
24. Çolakoğlu F, Özgümüş OB, Sandallı C, Çelik Sevim E, Alpay Karaoğlu Ş, et al. (2010) Deniz suyu kökenli koliformda sınıf 1 ve sınıf 2 integron gen kasetleri ve antibiyotik direncinin karakterizasyonu. *Türk Mikrobiyoloji Cemiyeti Dergisi* 40(2): 97-108.
25. Sivri N, Balcı M, Durmus T, Seker DZ, Balkis N, et al. (2012) Analysis of enteric bacteria distribution in the gulf of Gemlik By Means of GIS. *fresenius environmental bulletin (FEB)* 21(11): 3224-3232.
26. Cingilli Vural H, Akçin A (2011) Investigation of “Contagious Type Antibiotic Resistance Properties” related with R plasmids in *Escherichia coli* strains isolated from Izmit Gulfs (Turkey). *Journal of Kafkas Univ Vet Fak* 17(Suppl A): 23-30.
27. Matyar F, Dinçer S (2010) Antibiotic and heavy metal resistance of *Enterococcus faecalis* isolated from Eastern Mediterranean Sea. *SDU Journal of Science (E Journal)* 5(2): 172-178.

28. Karayakar F, Ay Ö, Cıck B (2004) The identification of plasmide dependent resistancy of *Escherichia coli* against some antibiotics isolated from stations on Mersin shore line. Ecology Cev Kor 13(52): 28-32.
29. Çardak M, Altuğ G (2010) Distribution of members of the family Enterobacteriaceae in the Istanbul Strait. J Black Sea Mediterranean Environment 16: 295-310.
30. Altuğ G, Gürün S, Çardak M, Çiftçi PS, Kalkan S, et al. (2012) Occurrence of pathogenic bacteria in some ships' ballast water incoming from various marine regions to the Sea of Marmara, Turkey. Marine Environmental Research 81: 35-42.



This work is licensed under Creative Commons Attribution 4.0 License  
DOI: [10.19080/AIBM.2017.06.555694](https://doi.org/10.19080/AIBM.2017.06.555694)

### Your next submission with Juniper Publishers will reach you the below assets

- Quality Editorial service
- Swift Peer Review
- Reprints availability
- E-prints Service
- Manuscript Podcast for convenient understanding
- Global attainment for your research
- Manuscript accessibility in different formats  
( Pdf, E-pub, Full Text, Audio)
- Unceasing customer service

Track the below URL for one-step submission

<https://juniperpublishers.com/online-submission.php>