

# **Current Research in Nutrition and Food Science**

Journal Website:www.foodandnutritionjournal.org

# The Use of Electrochemical Biosensors in Food Analysis

# JOHN BUNNEY, SHAE WILLIAMSON, DIANNE ATKIN, MARYN JEANNERET, DANIEL COZZOLINO,\* JAMES CHAPMAN, AOIFE POWER and SHANEEL CHANDRA

Agri-Chemistry Group, School of Health, Medical and Applied Sciences Central Queensland University, Rockhampton North, QLD 4702, Australia.

# Abstract

Rapid and accurate analysis of food produce is essential to screen for species that may cause significant health risks like bacteria, pesticides and other toxins. Considerable developments in analytical techniques and instrumentation, for example chromatography, have enabled the analyses and guantitation of these contaminants. However, these traditional technologies are constrained by high cost, delayed analysis times, expensive and laborious sample preparation stages and the need for highly-trained personnel. Therefore, emerging, alternative technologies, for example biosensors may provide viable alternatives. Rapid advances in electrochemical biosensors have enabled significant gains in quantitative detection and screening and show incredible potential as a means of countering such limitations. Apart from demonstrating high specificity towards the analyte, these biosensors also address the challenge of the multifactorial food industry of providing high analytical accuracy amidst complex food matrices, while also overcoming differing densities, pH and temperatures. This (public and Industry) demand for faster, reliable and cost-efficient analysis of food samples, has driven investment into biosensor design. Here, we discuss some of the recent work in this area and critique the role and contributions biosensors play in the food industry. We also appraise the challenges we believe biosensors need to overcome to become the industry standard.



# **Article History**

Received: 22 September 2017 Accepted: 23 October 2017

#### Keywords

Biosensors, Food analysis, Selectivity, Sensitivity, Rapid analysis.

# Introduction

Food Safety

The issue of food safety has emerged as increasingly significant public concerns worldwide due to sub-

quality foods being linked to increased morbidity, mortality, human suffering, and economic burden<sup>1</sup>. Accordingly, in an information-age society where consumer awareness and expectations of safety

**CONTACT** Daniel Cozzolino d.cozzolino@cqu.edu.au QAgri-Chemistry Group, School of Health, Medical and Applied Sciences Central Queensland University, Rockhampton North, QLD 4702, Australia.

© 2017 The Author(s). Published by Enviro Research Publishers

This is an **b** Open Access article licensed under a Creative Commons Attribution-NonCommercial-ShareAlike 4.0 International License (https://creativecommons.org/licenses/by-nc-sa/4.0/), which permits unrestricted NonCommercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

To link to this article: http://dx.doi.org/10.12944/CRNFSJ.5.3.02

are high, food manufacturers have to meet the need of modern consumers to make informed purchase decisions and their preference for food products with high quality and affordable price, and at the same time, must maintain high-quality standards and assurance of product safety<sup>2</sup>.

Matching the end-user compliance with regulatory guidelines on food quality, the instrumentation and scientific industries have responded with continuous improvement and development of analytical methodologies of many analytical methods, liquid chromatography has acquired a role of great importance in a majority of food analysis, as witnessed by the wide range of applications that can be found throughout the whole literature<sup>3-5</sup>. However, choromatographic analysis is constrained by the rigors of often elaborate sample preparation, and homogenization, clean up and then the analytical component of the test to determine a viable concentration<sup>4</sup>. Consequently, the process often must be repeated multiple times, as many samples are needed to give an accurate result due to the number of interferences in the matrix extracts, which can result in inaccurate identification and false positives<sup>6</sup>. Additionally, the extensive set-up and extraction / clean-up processes required for HPLC analysis can cause prolonged delays in contaminant identification<sup>7</sup>. This hefty time requirement makes HPLC methods unsuitable for "fresh foods" which are typically consumed in a short period, given short shelf lives. It is possible that by the time a contaminant is detected, multiple individuals may have been exposed to it, increasing likelihood of contamination<sup>8,9</sup>.

The need for selective measurement of analytes in food is paramount<sup>10</sup>. Here, "younger" technologies like those on an electrochemical platform may present viable alternatives. Biosensors are an examples of new, innovative methods to tacking old but important problems in a quality-conscious society and have become powerful instruments in clinical, environmental and especially, food analyses<sup>11</sup>.

Therefore, in this review, we appraise biosensors applied to food analysis. We will examine the attributes of biosensors that present attractive alternatives to traditional technologies and instrumentation, briefly explore recent advances in biosensor technologies and also critique their limitations. We conclude the review by proposing future directions and challenges that the biosensor research arena has to overcome to establish as the new order in food analysis and safety.

## **Biosensor Attributes**

A biosensor can be defined as an analytical device that combines a biologically sensitive recognition element (such as antibodies, nucleic acids, enzymes, organelles, whole cells and aptamers) immobilized on a physicochemical transducer, and connected to

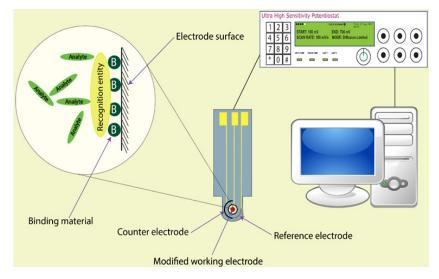


Fig. 1: A simplified, general scheme of a biosensor depicating the three electrode system, direction of electron transfer on the working electrode and a close-up of the working electrode interface with the recognition entity. Reproduced with permission from Ref.<sup>14</sup>

a detector to identify the presence of one or more specific analytes, their concentrations, and kinetics in a sample<sup>12</sup>. Electrochemical biosensors use an electrode transducer to detect electrons released by the reaction of the bioreceptor and analyte to obtain a measurable analysis of the contaminant<sup>13</sup>. Figure 1 shows the general scheme of a biosensor.

Key to the operative success of biosensors is their biological recognition elements which imparts a superior level of specificity and binding affinity with the target molecule. Such binding is termed specific binding or coupling and determines if an interaction occurs, which creates the electrical signal that is recorded and amplified<sup>14</sup>. Because of the particularity of the recognition entity toward the analyte, a high level of selectivity is achieved which results in signals generated solely from such precise interactions, irrespective of the matrix complexity<sup>15</sup>. This is terrifically illustrated by, commercially available, glucose meters that exploit a working electrode modified with glucose oxidase ensuring that a response is solely derived by glucose<sup>16</sup>.

The demand for high-speed, accurate and selective identification of analytes present within food produce, such as pathogenic bacteria<sup>17</sup>, pesticides<sup>18</sup> and toxins<sup>19</sup>, has facilitated rapid advances in biosensor development and enabled quantitative detection and screening. Apart from their inherent specificity, these biosensors help address the multifactorial food industry challenge of high analytical accuracy in the midst of the complex food matrices, overcoming differing densities, pH and temperature<sup>20</sup>. It has been argued that for biosensor use to become widespread, they need to offer further substantial benefits over existing methods<sup>21</sup>. One such advance is the potential for sensor miniaturization, which results in the sensor requiring greatly reduced sample sizes or volumes. However, sensor miniaturization has only partially been achieved to date<sup>22</sup> due to limitations in the structural integrity of very fine electrode tips associated with microelectrodes<sup>23,24</sup>.

The recent development of novel bio-recognition molecules, such as synthetic aptamers, DNA, proteins and viruses has enabled considerable selectivity in analysis<sup>25,26</sup>. Furthermore, parallel improvements in the immobilization of bio-recognition

molecules<sup>27</sup> through robust attachment methods like electrodeposition<sup>28,29</sup> and nanoparticle-bound entities at the working electrode interface is a significant step in the increased application of biosensors in food analysis. This is due in no small part to their greater specificity, selectivity and affinity for their target analytes<sup>25,26</sup>.

#### New Leaps in Biosensing

The field of biosensors has certainly witnessed astonishing growth in recent times. Over the last three decades, the number of papers published on the subject each year has increased/risen approximately 4000%<sup>30</sup>. Biosensor development and construction has focused predominantly on clinical research, continuing on the pioneering efforts of Clarke in 1950's and 60's<sup>31,32</sup>. Yet, a shift in biosensor focus towards food analysis has grown in the last decade due to the improved accuracy in target pursuit, intensifying demands about quality from stakeholders, such as safety regulators, traders and consumers as well as significant reduction in analysis times associated with electrochemical detection<sup>33</sup>.

In the agriculture and food industries, early detection and sensitive analysis of potential contaminants and toxins is crucial<sup>34</sup> and driven by a multiplicity of factors, such as the short shelf life of many fresh food products<sup>35</sup>, increasing consumer preferences for chemical free and unprocessed foods<sup>36</sup>, minimization of waste/reduction of costs in processing operations<sup>37</sup>, and the need for detection in very low quantities, and removal of pathogens from the supply chain that may cause serious illness to the consumer<sup>38</sup>.

These pressures are exacerbated by the inherent limitations of traditional food analysis methods that involve expensive, cumbersome instrumentation<sup>39</sup> and as a result, have helped shape biosensor development relating to food analysis<sup>40</sup>. Naturally, researchers have recognized the need for inexpensive, portable sensors that can perform rapidly and accurately with great sensitivity<sup>41</sup>.

# **Food Analysis Challenges**

Biosensors address three broad categories of food analysis expectations: safety, quality and authenticity<sup>42</sup>. Food safety screening focuses on the detection of undesirable contaminants in food, such as pesticide and antibiotic residues<sup>43,44</sup>, allergens<sup>45</sup>, biological toxins<sup>46</sup> and pathogenic microbes<sup>47</sup>. Similar analysis is also used to establish or confirm the nutritional value of a food product<sup>48</sup>. Authenticity analysis seeks to confirm the origin and/or production process of a food stuff, while also providing information about the adulteration or counterfeiting of food<sup>48,49</sup>. The literature indicates that presently, electrochemical biosensors are primarily being utilized in food safety rather than quality and authenticity analysis<sup>50,51</sup>.

Traditional analysis methods for detecting harmful microorganisms, such as pathogenic *Escherichia* and *Salmonella*<sup>52</sup>, aflatoxins<sup>53</sup> and pesticides such as organophosphates and carbamates<sup>54</sup> could only be conducted post-production. This limitation is easily overcome by the use of biosensors which allows food items to be tested at all phases of production<sup>52</sup> from raw materials screening to the product on shelf,

resulting in more efficient means of ensuring of food safety and outbreak prevention<sup>55</sup>.

# **Timeliness and Costs**

Improved analysis times is another benefit to biosensor application in food analysis. Using an array of biosensors on a microfluidic or lab-on-achip platform, low volume samples can be analyzed directly, thus eliminating the need for laborious and costly sample preparation stages<sup>56</sup>. This is a particularly attractive feature of biosensors, where toxin accumulation often correlates with time<sup>57</sup>, for example, mycotoxins are harmful carcinogenic metabolites produced by mold which affects many food products, including but not limited to; bread, cereals, dried fruit, wine and meat products<sup>7</sup>.

Biosensors present an attractive alternative as their capability of being used *in situ* allows reduced detection times, from several days to hours, or

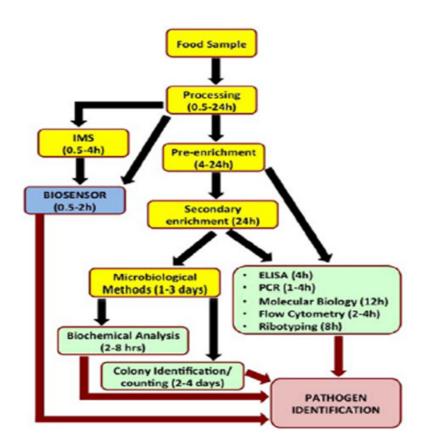


Fig. 2: A flowchart elucidating the processing steps involved and relative time taken in detecting a pathogen in a food sample. reproduced under license from Ref. <sup>67</sup>

even minutes<sup>58-60</sup> as illustrated in Figure 2. Other advantages stemming from in-situ determination capabilities include minimized sampling protocols, reduced storage requirements and the removal of often elaborate sample preparation procedures<sup>61</sup>. Furthermore, in-situ detection capabilities allow for the improved portability of analysis tools such as handheld detection devices which generally require minimal training to operate<sup>62</sup> and can facilitate the integration of real time analysis in food processing work centers/systems63. Improved timeliness within food processing systems can also reduce spoilage, particularly in fresh produce, such as seafood, this was illustrated by the development of for the rapid detection of Vibrio parahaemolyticus<sup>64</sup> - a leading global cause of bacterial gastroenteritis. Whilst bacteriophages have been successfully used to remove antibiotic strains of V. parahaemolyticus from seafood<sup>65</sup>, the method lengthens the time between catch and plate, thus reducing the seafood's freshness and ultimately its value<sup>66</sup>.

The cost effectiveness of biosensors cannot be overstated. The rapid analysis rendered from biosensing allows significant gains through cost mitigation normally reserved for sample preparation methods and the need for expensive laboratories with highly trained staff<sup>68</sup>, and the additional possibility of automated on-line analysis in food processing plants69 which will further reduce cost. Moreover, the ability of biosensors to detect contaminants in raw foods in real-time with high specificity and very low concentrations reduces waste<sup>55</sup> and the economic costs associated with health issues and product recalls<sup>67</sup>.

#### Losses Due to Sample Preparation

A fundamental prerequisite to using traditional methodologies in food analysis is the sample homogenization process, which can be problematic because of the organic acids and antimicrobial compounds present in many fruits and vegetables<sup>70</sup>. The release of these compounds during sample preparation can inhibit the detection of certain contaminants, potentially having detrimental impacts on product consumers, a problem not encountered by biosensors as they require little or no sample preparation<sup>8</sup>. Methods commonly used for detecting pathogenic bacteria detection in foods, such as enzyme-linked immunosorbent assay (ELISA),

polymerase chain reaction (PCR) or cell culture<sup>71</sup> are incredibly time consuming. The identification of certain pathogens may take days as they have lengthy sample preparation times coupled with low sensitivity, which can often result in false positives: ELISA requires a 24-48 hr period to successfully detect harmful pathogens such as *Escherichia coli*, a leading cause of death in children under five<sup>72</sup>. Target-induced aptamer displacement strategies can overcome the time and sensitivity barriers by completing the test within 3.5 hours at a sensitivity of 112 CFUmL-1 in a phosphate buffer saline and 305 CFUmL-1 in a milk solution<sup>73</sup>. This far exceeds the sensitivity of ELISA for *E. coli* detection.

## **Biosensor Detection Process**

Despite the ubiquity of microbes, their detection in food is difficult<sup>74</sup> and further complicated by the fact that only some strains are pathogenic<sup>75</sup>. Therefore, screening for the presence of bacteria alone is insufficient for food safety analysis and ideally, only the pathogenic strains, such as *E. coli* is one of two pathogenic strains responsible for 5 food poisoning deaths in Japan in 2011, should be identified<sup>76</sup>. Here, biosensors present notable advances compared to traditional analysis methods in targeting only the analyte, such as the enterohaemorrhagic *Escherichia coli* strain O111<sup>73</sup>.

It should be noted that the specificity of biosensors is not limited to the detection of a singular analyte. Several biosensors have been developed to detect minute levels of multiple pesticide residues in foods based on the biochemical pathways the pesticides act upon, such as acetylcholinesterase (AChE) inhibitors. This means biosensor usefulness can extend to an entire class of pesticides77. Similarly biosensors have been designed to detect certain compounds or toxin vectors, because of their inherent potential for inducing acute toxicity. Screening for these is critically important to food safety as such contaminants may have devastating effects even in very low concentrations. For example as few as ~10 bacteria can cause infection<sup>67</sup>; carbamate pesticides which, despite having a low bioaccumulation potential are considered carcinogenic<sup>78</sup>, and antibiotic residues in animal-derived foods can cause allergic reactions and even secondary infections<sup>79</sup>. Biosensors can also detect traditionally challenging 'viable but not culturable' (VBNC) bacteria, differentiating dangerous pathogens that are in a state of dormancy from non-living, non-threatening bacteria<sup>80</sup>.

#### **Current Innovations in Biosensor Design**

While enzymatic biosensors were recognized as a leap into elevated or ultimate selectivity, the next stage in biosensor design includes gene based sensors involving DNA; as the recognition or coupling entity (via hybridization), antibody or antigen based biosensors; and whole cell sensors<sup>81</sup>. Within the agri-food industry, pathogen detection trends have focused on the utilization of single sensor platform for detection of multiple pathogens/toxins<sup>82</sup>. More recently, biotechnology has shifted into ever smaller systems to allow for portability, cost reduction, analysis time reduction and commercial viability83. Improvements in microfabrication systems have similarly aided in advancing biosensor technology and utility<sup>84</sup>. Emerging nanomaterials, such as nanoparticles and nanofibers have featured in these, paving the way for this miniaturization trend<sup>85</sup>.

Such functional nanomaterials enhance electrochemical biosensors in two ways: refining the response features of the electrode by increasing its surface area for instance and assisting in robust attachment of the bioreceptor/recognition entity. With greater surface to area volume ratios, nanomaterials lend greater catalytic prowess, ensure biocompatibility and achieve lower mass transfer resistance. This translates to improved selectivity, sensitivity, time efficiency and cost effectiveness for the biosensor<sup>86-88</sup>. Similarly, the increase in transducer surface area delivers greater conductivity and sensitivity, promotes greater interaction capacity<sup>89</sup> and lowers detection limits<sup>90</sup>. These are all ideal features of a biosensing interface. An excellent example of a nano-biosensor, capable of pesticide residue detection in concentrations as low as 0.4 pM, has been reported in by Verma et al.,86. Furthermore, the inclusion of other nanomaterials at the transducer level, for example carbon nanotubes<sup>91</sup>, can increase electron transfer and increase the transducer activity92. Evidence of these improvements is in the slow but gradual replacement of traditional enzyme-substrate biosensors by nano-biosensor technology93. Nano-biosensors have been developed for the agriculture and food processing industries to identify and quantify pesticides, herbicides, pathogenic microorganisms and other microbial contamination such as viruses and bacteria, hormones, glucose, as well as the presence of insects or fungus<sup>94-96</sup>.

Another medium of interest is microfluidics which provides throughput processing, reduces sample and reagents volume (down to the nanolitre)<sup>56</sup>, increases sensitivity, and employs a single platform for both sample preparation and detection<sup>97</sup>. Microfluidics are portable, disposable, offer real-time detection, and simultaneous analysis of different analytes in a single device with exceptional accuracy<sup>98,99</sup>. For example, microfluidic nano-biosensor for the detection of pathogenic species like *Salmonella* have already been proposed recently<sup>100</sup>.

It is envisaged that the ever improving analytical properties of electrochemical transducers will even allow for the detection of multiple analytes simultaneously<sup>101</sup>. However, despite these promising advances and the potential of nanomaterial-based biosensors, realistically their application within food matrices is still in the very early stages of development<sup>102</sup>. Compared with other biosensing forays, for example in medicinal biosensor technology through favored point of care and home diagnostics for pregnancy, glucose content, biosensing in food production and processing or screening has not been embraced as readily<sup>103-105.</sup>

## **Concluding Remarks and Future Directions**

Although biosensors display clear advantages over traditional methods, the perfect biosensor does not as yet exist<sup>106</sup> and there are many obstacles in its development to be overcome<sup>107</sup>. Presently, many biosensors are not easily implementable, if only because so few are currently available commercially<sup>107</sup>.

Nonetheless, it is almost inevitable that the future of biosensors will involve partnership with information communications technology to assist food producers, retailers, authorities and even consumers, in their decision making108 by equipping them with the necessary tools and data to improve their decision-making process. This will ultimately, enable greater management of the natural resources<sup>109-112</sup>. Moreover, inspired by mammalian sensory networks, new sensor systems

being developed have the potential to revolutionize food analysis<sup>113,114</sup>. Biomimetic sensors, such as electronic tongues and electronic noses are based on biosensor technologies<sup>115</sup> and we expect that their exploitation of arrays of low specificity sensors capable of detecting multiple signals will allows a more complete analysis of food quality. Inspiration for these developments and applications comes from the electronic tongues that form the basis for food authenticity and safety sensor systems<sup>116,117</sup>, or similarly, electronic noses that can detect unique volatile compounds within the tea, wine, coffee, and spice industries<sup>118-120</sup>.

The combination of different types of biosensors has great promise: the fusion of electronic tongues with electronic noses and may further increase the identification capabilities of such a biomimetic system, as precisely as it does within the biological system<sup>121</sup>. The advantage of real time monitoring in food manufacture, especially of dairy products<sup>122</sup> and brewed products<sup>123</sup>, further enhances the usefulness of biosensors and drives the push for their commercially availability to general public<sup>124</sup>. The inherent specificity, sensitivity, and adaptability of biosensors make them the ideal candidate for use as a safety net throughout the food industry improving product quality with minimal investment<sup>124</sup>, both now and for the foreseeable future. The opportunity afforded through biosensing, particularly in situ and safety analysis at all levels of the supply chain, as well as authenticity and guality analysis by the consumers themselves, make biosensors food productions tool of the future.

#### References

- Wu, M.Y.-C.; Hsu, M.-Y.; Chen, S.-J.; Hwang, D.-K.; Yen, T.-H.; Cheng, C.-M. Point-of-care detection devices for food safety monitoring: Proactive disease prevention. *Trends in Biotechnology*; **35**(4): 288-300: (2017)
- Baiano, A. Applications of hyperspectral imaging for quality assessment of liquid based and semi-liquid food products: A review. Journal of Food Engineering; 214 (Supplement C): 10-15: (2017)
- Cacciola, F.; Dugo, P.; Mondello, L. Multidimensional liquid chromatography in food analysis. *TRAC Trends in Analytical Chemistry*: (2017)
- Sun, H.; Ge, X.; Lv, Y.; Wang, A. Application of accelerated solvent extraction in the analysis of organic contaminants, bioactive and nutritional compounds in food and feed. *Journal of Chromatography A*; 12371-23: (2012)
- Verdu, C.F.; Gatto, J.; Freuze, I.; Richomme, P.; Laurens, F.; Guilet, D. Comparison of two methods, uhplc-uv and uhplc-ms/ms, for the quantification of polyphenols in cider apple juices. *Molecules (Basel, Switzerland)*; 18(9): 10213-10227: (2013)
- Frenich, A.G.; Vidal, J.L.M.; López, T.L.; Aguado, S.C.; Salvador, I.M. Monitoring

multi-class pesticide residues in fresh fruits and vegetables by liquid chromatography with tandem mass spectrometry. *Journal of Chromatography A*; **1048**(2): 199-206: (2004)

- Alshannaq, A.; Yu, J.H. Occurrence, toxicity, and analysis of major mycotoxins in food. Int *J Environ Res Public Health*; **14**(6): (2017)
- Yeni, F.; Acar, S.; Polat, Ö.G.; Soyer, Y.; Alpas, H. Rapid and standardized methods for detection of foodborne pathogens and mycotoxins on fresh produce. *Food Control*; 40359-367: (2014)
- Välimaa, A.-L.; Tilsala-Timisjärvi, A.; Virtanen, E. Rapid detection and identification methods for listeria monocytogenes in the food chain – a review. *Food Control*; 55103-114: (2015)
- 10. Vasilescu, A.; Marty, J.-L. Electrochemical aptasensors for the assessment of food quality and safety. *TRAC Trends in Analytical Chemistry*; 7960-70: (2016)
- 11. Bahadır, E.B.; Sezgintürk, M.K. Applications of commercial biosensors in clinical, food, environmental, and biothreat/biowarfare analyses. *Analytical Biochemistry*; 478107-120: (2015)
- 12. Perumal, V.; Hashim, U. Advances in biosensors: Principle, architecture and

applications. *Journal of Applied Biomedicine*; 12(1): 1-15: (2014)

- Thévenot, D.R.; Toth, K.; Durst, R.A.; Wilson, G.S. Electrochemical biosensors: Recommended definitions and classification1international union of pure and applied chemistry: Physical chemistry division, commission i.7 (biophysical chemistry); analytical chemistry division, commission v.5 (electroanalytical chemistry).1. *Biosensors and Bioelectronics*; **16**(1): 121-131: (2001)
- Sharma, A.; Goud, K.Y.; Hayat, A.; Bhand, S.; Marty, J.L. Recent advances in electrochemical-based sensing platforms for aflatoxins detection. *Chemosensors*; 5(1): (2017)
- Idili, A.; Amodio, A.; Vidonis, M.; Feinberg-Somerson, J.; Castronovo, M.; Ricci, F. Foldingupon-binding and signal-on electrochemical DNA sensor with high affinity and specificity. *Analytical Chemistry*; *86*(18): 9013-9019: (2014)
- Yoo, E.-H.; Lee, S.-Y. Glucose biosensors: An overview of use in clinical practice. *Sensors* (*Basel, Switzerland*); **10**(5): 4558-4576: (2010)
- 17. Arora, P.; Sindhu, A.; Kaur, H.; Dilbaghi, N.; Chaudhury, A. An overview of transducers as platform for the rapid detection of foodborne pathogens. *Applied Microbiology and Biotechnology*; **97**(5): 1829-1840: (2013)
- Arduini, F.; Cinti, S.; Scognamiglio, V.; Moscone, D. Nanomaterials in electrochemical biosensors for pesticide detection: Advances and challenges in food analysis. *Microchimica Acta*; **183**(7): 2063-2083: (2016)
- Bazin, I.; Tria, S.A.; Hayat, A.; Marty, J.-L. New biorecognition molecules in biosensors for the detection of toxins. *Biosensors and Bioelectronics*; 87285-298: (2017)
- Gaudin, V. Advances in biosensor development for the screening of antibiotic residues in food products of animal origin – a comprehensive review. *Biosensors and Bioelectronics*; 90363-377: (2017)
- Velusamy, V.; Arshak, K.; Korostynska, O.; Oliwa, K.; Adley, C. An overview of foodborne pathogen detection: In the perspective of biosensors. *Biotechnology Advances*; *28*(2): 232-254: (2010)

- Bettazzi, F.; Marrazza, G.; Minunni, M.; Palchetti, I.; Scarano, S. Biosensors and related bioanalytical tools. *Comprehensive Analytical Chemistry*, 771-33: (2017)
- Chandra, S.; Miller, A.D.; Wong, D.K.Y. Evaluation of physically small p-phenylacetatemodified carbon electrodes against fouling during dopamine detection in vivo. *Electrochimica Acta*; 101225-231: (2013)
- Chandra, S.; Miller, A.D.; Bendavid, A.; Martin, P.J.; Wong, D.K.Y. Minimizing fouling at hydrogenated conical-tip carbon electrodes during dopamine detection in vivo. *Analytical Chemistry*; 86(5): 2443-2450: (2014)
- 25. Chandra, S.; Siraj, S.; Wong, D.K.Y. Recent advances in biosensing for neurotransmitters and disease biomarkers using microelectrodes. *ChemElectroChem*; 4(4): 822-833: (2017)
- Ali, J.; Najeeb, J.; Ali, M.A.; Aslam, M.F.; Raza, A. Biosensors: Their fundamentals, designs, types and most recent impactful applications: A review. *Journal of Biosensors* & *Bioelectronics*; 8(1): (2017)
- 27. Putzbach, W.; Ronkainen, J.N. Immobilization techniques in the fabrication of nanomaterialbased electrochemical biosensors: A review. *Sensors*; **13**(4): (2013)
- Srivastava, S.; Kumar, V.; Ali, M.A.; Solanki, P.R.; Srivastava, A.; Sumana, G.; Saxena, P.S.; Joshi, A.G.; Malhotra, B.D. Electrophoretically deposited reduced graphene oxide platform for food toxin detection. *Nanoscale*; *5*(7): 3043-3051: (2013)
- Devi, R.; Yadav, S.; Nehra, R.; Yadav, S.; Pundir, C.S. Electrochemical biosensor based on gold coated iron nanoparticles/chitosan composite bound xanthine oxidase for detection of xanthine in fish meat. *Journal of Food Engineering*; **115**(2): 207-214: (2013)
- Turner, A.P.F. Biosensors: Sense and sensibility. *Chemical Society Reviews*; 42(8): (2013)
- Harper, A.; Anderson, M.R. Electrochemical glucose sensors—developments using electrostatic assembly and carbon nanotubes for biosensor construction. *Sensors*; 10(9): (2010)
- Bhalla, N.; Jolly, P.; Formisano, N.; Estrela,
  P. Introduction to biosensors. *Essays in Biochemistry*; *60*(1): 1-8: (2016)

- Kim, K.-P.; Singh, A.; Bai, X.; Leprun, L.; Bhunia, A. Erratum: Kim, k.-p.; singh, a.K.; bai, x.; leprun, l.; bhunia, a.K. Novel pcr assays complement laser biosensor-based method and facilitate listeria species detection from food. Sensors 2015, 15, 22672–22691. Sensors; 17(5): 945-945: (2017)
- Sharma, H.; Mutharasan, R. Review of biosensors for foodborne pathogens and toxins. *Sensors and Actuators B: Chemical*; 183535-549: (2013)
- Biji, K.B.; Ravishankar, C.N.; Mohan, C.O.; Srinivasa Gopal, T.K. Smart packaging systems for food applications: A review. *Journal of Food Science and Technology*; 52(10): 6125-6135: (2015)
- Law, J.W.-F.; Ab Mutalib, N.-S.; Chan, K.-G.; Lee, L.-H. Rapid methods for the detection of foodborne bacterial pathogens: Principles, applications, advantages and limitations. 2014; Vol. 5.
- Nychas, G.-J.E.; Panagou, E.Z.; Mohareb, F. Novel approaches for food safety management and communication. *Current Opinion in Food Science*; 1213-20: (2016)
- Vidal, J.C.; Bonel, L.; Ezquerra, A.; Hernández, S.; Bertolín, J.R.; Cubel, C.; Castillo, J.R. Electrochemical affinity biosensors for detection of mycotoxins: A review. *Biosensors* and *Bioelectronics*; 49146-158: (2013)
- McGrath, T.F.; Elliott, C.T.; Fodey, T.L. Biosensors for the analysis of microbiological and chemical contaminants in food. *Analytical* and Bioanalytical Chemistry; 403(1): 75-92: (2012)
- Korotkaya, E. Biosensors: Design, classification, and applications in the food industry. *Foods and Raw Materials*; 2(2): (2014)
- Justino, C.I.L.; Freitas, A.C.; Pereira, R.; Duarte, A.C.; Rocha Santos, T.A.P. Recent developments in recognition elements for chemical sensors and biosensors. *TRAC Trends in Analytical Chemistry*; 682-17: (2015)
- 42. Rotariu, L.; Lagarde, F.; Jaffrezic-Renault, N.; Bala, C. Electrochemical biosensors for fast detection of food contaminants – trends and perspective. *TRAC Trends in Analytical Chemistry*; 7980-87: (2016)

- Ye, W.; Guo, J.; Bao, X.; Chen, T.; Weng, W.; Chen, S.; Yang, M. Rapid and sensitive detection of bacteria response to antibiotics using nanoporous membrane and graphene quantum dot (gqds)-based electrochemical biosensors. *Materials*; **10**(6): 603: (2017)
- Ribeiro, F.W.P.; Barroso, M.F.; Morais, S.; Viswanathan, S.; de Lima-Neto, P.; Correia, A.N.; Oliveira, M.B.P.P.; Delerue-Matos, C. Simple laccase-based biosensor for formetanate hydrochloride quantification in fruits. *Bioelectrochemistry*; 957-14: (2014)
- 45. Andjelkovic, U.; Gavrovic-Jankulovic, M.; Martinovic, T.; Josic, D. Omics methods as a tool for investigation of food allergies. *TRAC Trends in Analytical Chemistry*: (2017)
- Malhotra, B.D.; Srivastava, S.; Ali, M.A.; Singh, C. Nanomaterial-based biosensors for food toxin detection. *Applied Biochemistry and Biotechnology*; **174**(3): 880-896: (2014)
- Singh, R.; Mukherjee, M.D.; Sumana, G.; Gupta, R.K.; Sood, S.; Malhotra, B.D. Biosensors for pathogen detection: A smart approach towards clinical diagnosis. *Sensors* and Actuators B: Chemical; 197385-404: (2014)
- Cao, M.; Li, Z.; Wang, J.; Ge, W.; Yue, T.; Li, R.; Colvin, V.L.; Yu, W.W. Food related applications of magnetic iron oxide nanoparticles: Enzyme immobilization, protein purification, and food analysis. *Trends in Food Science & Technology*; 27(1): 47-56: (2012)
- 49. Inbaraj, B.S.; Chen, B.H. Nanomaterialbased sensors for detection of foodborne bacterial pathogens and toxins as well as pork adulteration in meat products. *Journal of Food and Drug Analysis*; **24**(1): 15-28: (2016)
- 50. Mehrotra, P. Biosensors and their applications a review. 2016; Vol. 6.
- Lavecchia, T.; Tibuzzi, A.; Giardi, M.T. Biosensors for functional food safety and analysis. Giardi, M.T.; Rea, G.; Berra, B., Eds. Springer US: Boston, MA, 2010; pp 267-281.
- Adley, C.C. Past, present and future of sensors in food production. *Foods*; *3*(3): 491-510: (2014)
- Castillo, G.; Spinella, K.; Poturnayová, A.; Šnejdárková, M.; Mosiello, L.; Hianik, T. Detection of aflatoxin b1 by aptamer-based

biosensor using pamam dendrimers as immobilization platform. *Food Control*; 529-18: (2015)

- 54. Cesarino, I.; Moraes, F.C.; Lanza, M.R.V.; Machado, S.A.S. Electrochemical detection of carbamate pesticides in fruit and vegetables with a biosensor based on acetylcholinesterase immobilised on a composite of polyaniline–carbon nanotubes. *Food Chemistry*, **135**(3): 873-879: (2012)
- 55. Jayas, D.S. The role of sensors and bioimaging in monitoring food quality. *Resource Magazine* 2017, p 12.
- Weng, X.; Neethirajan, S. Ensuring food safety: Quality monitoring using microfluidics. *Trends in Food Science & Technology*; 6510-22: (2017)
- 57. Moreb, N.A.; Priyadarshini, A.; Jaiswal, A.K. Knowledge of food safety and food handling practices amongst food handlers in the republic of ireland. *Food Control;* 80341-349: (2017)
- Ruiz-Valdepeñas Montiel, V.; Gutiérrez, M.L.; Torrente-Rodríguez, R.M.; Povedano, E.; Vargas, E.; Reviejo, Á.J.; Linacero, R.; Gallego, F.J.; Campuzano, S.; Pingarrón, J.M. Disposable amperometric polymerase chain reaction-free biosensor for direct detection of adulteration with horsemeat in raw lysates targeting mitochondrial DNA. *Analytical Chemistry*; 89(17): 9474-9482: (2017)
- Ahmed, A.; Rushworth, J.V.; Hirst, N.A.; Millner, P.A. Biosensors for whole-cell bacterial detection. 2014; Vol. 27, pp 631-646.
- Thakur, M.S.; Ragavan, K.V. Biosensors in food processing. *Journal of Food Science* and Technology; 50(4): 625-641: (2013)
- 61. Bülbül, G.; Hayat, A.; Andreescu, S. Portable nanoparticle-based sensors for food safety assessment. *Sensors*; **15**(12): (2015)
- Garg, M.; Mehrotra, S. Biosensors. In Principles and applications of environmental biotechnology for a sustainable future, Singh, R.L., Ed. Springer Singapore: Singapore, 2017; pp 341-363.
- Alves, R.C.; Barroso, M.F.; González-García, M.B.; Oliveira, M.B.P.P.; Delerue-Matos, C. New trends in food allergens detection: Toward biosensing strategies. *Critical Reviews*

*in Food Science and Nutrition*; **56**(14): 2304-2319: (2016)

- Nordin, N.; Yusof, N.A.; Abdullah, J.; Radu, S.; Hushiarian, R. A simple, portable, electrochemical biosensor to screen shellfish for vibrio parahaemolyticus. *AMB Express*; 7(1): 41: (2017)
- Jun, J.W.; Kim, H.J.; Yun, S.K.; Chai, J.Y.; Park, S.C. Eating oysters without risk of vibriosis: Application of a bacteriophage against vibrio parahaemolyticus in oysters. *International Journal of Food Microbiology*; 18831-35: (2014)
- Castro-Ibáñez, I.; López-Gálvez, F.; Gil, M.I.; Allende, A. Identification of sampling points suitable for the detection of microbial contamination in fresh-cut processing lines. *Food Control*; 59841-848: (2016)
- 67. Singh, A.; Poshtiban, S.; Evoy, S. Recent advances in bacteriophage based biosensors for food-borne pathogen detection. *Sensors* (*Basel*); **13**(2): 1763-1786: (2013)
- Wang, Y.; Salazar, J.K. Culture-independent rapid detection methods for bacterial pathogens and toxins in food matrices. *Comprehensive Reviews in Food Science* and Food Safety; **15**(1): 183-205: (2016)
- Zeng, Y.; Zhu, Z.; Du, D.; Lin, Y. Nanomaterialbased electrochemical biosensors for food safety. *Journal of Electroanalytical Chemistry*; 781147-154: (2016)
- Farahi, R.H.; Passian, A.; Tetard, L.; Thundat, T. Critical issues in sensor science to aid food and water safety. *ACS Nano*; 6(6): 4548-4556: (2012)
- Ligaj, M.; Tichoniuk, M.; Gwiazdowska, D.; Filipiak, M. Electrochemical DNA biosensor for the detection of pathogenic bacteria aeromonas hydrophila. *Electrochimica Acta*; 12867-74: (2014)
- Poltronieri, P.; Mezzolla, V.; Primiceri, E.; Maruccio, G. Biosensors for the detection of food pathogens. *Foods*; *3*(3): 511-526: (2014)
- Luo, C.; Lei, Y.; Yan, L.; Yu, T.; Li, Q.; Zhang, D.; Ding, S.; Ju, H. A rapid and sensitive aptamer-based electrochemical biosensor for direct detection of escherichia coli o111. *Electroanalysis*; *24*(5): 1186-1191: (2012)
- 74. Zwietering, M.H.; den Besten, H.M.W.

Microbial testing in food safety: Effect of specificity and sensitivity on sampling plans—how does the oc curve move. *Current Opinion in Food Science*; 1242-51: (2016)

- Ma, X.; Jiang, Y.; Jia, F.; Yu, Y.; Chen, J.; Wang, Z. An aptamer-based electrochemical biosensor for the detection of salmonella. *Journal of Microbiological Methods*; 9894-98: (2014)
- Idil, N.; Mattiasson, B. Imprinting of microorganisms for biosensor applications. Sensors (Basel, Switzerland); 17(4): (2017)
- Raghu, P.; Madhusudana Reddy, T.; Reddaiah, K.; Kumara Swamy, B.E.; Sreedhar, M. Acetylcholinesterase based biosensor for monitoring of malathion and acephate in food samples: A voltammetric study. *Food Chemistry*, 142188-196: (2014)
- Oliveira, T.M.B.F.; Fátima Barroso, M.; Morais, S.; de Lima-Neto, P.; Correia, A.N.; Oliveira, M.B.P.P.; Delerue-Matos, C. Biosensor based on multi-walled carbon nanotubes paste electrode modified with laccase for pirimicarb pesticide quantification. *Talanta*; 106137-143: (2013)
- Chen, T.; Cheng, G.; Ahmed, S.; Wang, Y.; Wang, X.; Hao, H.; Yuan, Z. New methodologies in screening of antibiotic residues in animalderived foods: Biosensors. *Talanta*; 175435-442: (2017)
- Ayrapetyan, M.; Oliver, J.D. The viable but non-culturable state and its relevance in food safety. *Current Opinion in Food Science*; 8127-133: (2016)
- Yang, T.; Huang, H.; Zhu, F.; Lin, Q.; Zhang, L.; Liu, J. Recent progresses in nanobiosensing for food safety analysis. *Sensors*; *16*(7): (2016)
- Cho, I.-H.; Radadia, A.D.; Farrokhzad, K.; Ximenes, E.; Bae, E.; Singh, A.K.; Oliver, H.; Ladisch, M.; Bhunia, A.; Applegate, B., et al. Nano/micro and spectroscopic approaches to food pathogen detection. *Annual Review* of *Analytical Chemistry*; 7(1): 65-88: (2014)
- Warriner, K.; Reddy, S.M.; Namvar, A.; Neethirajan, S. Developments in nanoparticles for use in biosensors to assess food safety and quality. *Trends in Food Science & Technology*; 40(2): 183-199: (2014)
- 84. Derkus, B. Applying the miniaturization

technologies for biosensor design. *Biosensors* and *Bioelectronics*; 79901-913: (2016)

- Shruthi, G.S.; Amitha, C.V.; Blessy Baby, M. Biosensors: A modern day achievement. *Journal of Instrumentation Technology*; 2(1): 26-39: (2014)
- Verma, M.L. Nanobiotechnology advances in enzymatic biosensors for the agri-food industry. *Environmental Chemistry Letters*; 1-6: (2017)
- Zhu, C.; Yang, G.; Li, H.; Du, D.; Lin, Y. Electrochemical sensors and biosensors based on nanomaterials and nanostructures. *Analytical Chemistry*; **87**(1): 230-249: (2015)
- Anu Bhushani, J.; Anandharamakrishnan, C. Electrospinning and electrospraying techniques: Potential food based applications. *Trends in Food Science & Technology*; *38*(1): 21-33: (2014)
- Eivazzadeh-Keihan, R.; Pashazadeh, P.; Hejazi, M.; de la Guardia, M.; Mokhtarzadeh, A. Recent advances in nanomaterial-mediated bio and immune sensors for detection of aflatoxin in food products. *TRAC Trends in Analytical Chemistry*, 87112-128: (2017)
- Rai, M.; Jogee, P.S.; Ingle, A.P. Emerging nanotechnology for detection of mycotoxins in food and feed. *International Journal of Food Sciences and Nutrition*, 2015, Vol. *66*(4), p.363-370; 66(4): 363-370: (2015)
- Barsan, M.M.; Ghica, M.E.; Brett, C.M.A. Electrochemical sensors and biosensors based on redox polymer/carbon nanotube modified electrodes: A review. *Analytica Chimica Acta;* 8811-23: (2015)
- Yang, N.; Chen, X.; Ren, T.; Zhang, P.; Yang, D. Carbon nanotube based biosensors. *Sensors* and Actuators B: Chemical; 207690-715: (2015)
- Sharma, T.K.; Ramanathan, R.; Rakwal, R.; Agrawal, G.K.; Bansal, V. Moving forward in plant food safety and security through nanobiosensors: Adopt or adapt biomedical technologies? *PROTEOMICS*; **15**(10): 1680-1692: (2015)
- Sekhon, B.S. Nanotechnology in agri-food production: An overview. *Nanotechnology, Science and Applications*; 7: (2014)
- 95. Pashazadeh, P.; Mokhtarzadeh, A.;

Hasanzadeh, M.; Hejazi, M.; Hashemi, M.; de La Guardia, M. Nano-materials for use in sensing of salmonella infections: Recent advances. *Biosensors and Bioelectronics;* 871050-1064: (2017)

- Sharma, R.; Ragavan, K.V.; Thakur, M.S.; Raghavarao, K.S.M.S. Recent advances in nanoparticle based aptasensors for food contaminants. *Biosensors and Bioelectronics*; 74612-627: (2015)
- Luka, G.; Ahmadi, A.; Najjaran, H.; Alocilja, E.; DeRosa, M.; Wolthers, K.; Malki, A.; Aziz, H.; Althani, A.; Hoorfar, M. Microfluidics integrated biosensors: A leading technology towards labon-a-chip and sensing applications. *Sensors*; *15*(12): (2015)
- Rackus, D.G.; Shamsi, M.H.; Wheeler, A.R. Electrochemistry, biosensors and microfluidics: A convergence of fields. *Chemical Society Reviews*; 44(15): 5320-5340: (2015)
- Huang, Y.; Shi, Y.; Yang, H.Y.; Ai, Y. A novel single-layered mos2 nanosheet based microfluidic biosensor for ultrasensitive detection of DNA. *Nanoscale*; 7(6): 2245-2249: (2015)
- 100. Kim, G.; Moon, J.-H.; Moh, C.-Y.; Lim, J.-g. A microfluidic nano-biosensor for the detection of pathogenic salmonella. *Biosensors and Bioelectronics*; 67(Supplement C): 243-247: (2015)
- Vigneshvar, S.; Sudhakumari, C.C.; Senthilkumaran, B.; Prakash, H. Recent advances in biosensor technology for potential applications – an overview. *Frontiers in Bioengineering and Biotechnology*; 411: (2016)
- 102. Pilolli, R.; Monaci, L.; Visconti, A. Advances in biosensor development based on integrating nanotechnology and applied to food-allergen management. *TRAC Trends in Analytical Chemistry*; 4712-26: (2013)
- Qin, C.; Tao, L.; Phang, Y.H.; Zhang, C.; Chen, S.Y.; Zhang, P.; Tan, Y.; Jiang, Y.Y.; Chen, Y.Z. The assessment of the readiness of molecular biomarker-based mobile health technologies for healthcare applications. *Scientific Reports*; 5(October): 17854-17854: (2015)
- 104. Narsaiah, K.; Jha, S.N.; Bhardwaj, R.; Sharma, R.; Kumar, R. Optical biosensors for

food quality and safety assurance—a review. *Journal of Food Science and Technology*; *49*(4): 383-406: (2012)

- 105. Quesada-González, D.; Merkoçi, A. Mobile phone-based biosensing: An emerging "diagnostic and communication" technology. *Biosensors and Bioelectronics*; 92(June 2016): 549-562: (2017)
- Templier, V.; Roux, A.; Roupioz, Y.; Livache, T. Ligands for label-free detection of whole bacteria on biosensors: A review. *TRAC Trends in Analytical Chemistry*; 7971-79: (2016)
- 107. Valderrama, W.B.; Dudley, E.G.; Doores, S.; Cutter, C.N. Commercially available rapid methods for detection of selected food-borne pathogens. *Critical Reviews in Food Science* and Nutrition; 56(9): 1519-1531: (2016)
- 108. Zhang, D.; Liu, Q. Biosensors and bioelectronics on smartphone for portable biochemical detection. *Biosensors and Bioelectronics*; 75273-284: (2016)
- 109. Seo, S.-M.; Kim, S.-W.; Jeon, J.-W.; Kim, J.-H.; Kim, H.-S.; Cho, J.-H.; Lee, W.-H.; Paek, S.-H. Food contamination monitoring via internet of things, exemplified by using pocket-sized immunosensor as terminal unit. *Sensors and Actuators B: Chemical*; 233148-156: (2016)
- 110. Liu, Y.; Han, W.; Zhang, Y.; Li, L.; Wang, J.; Zheng, L. An internet-of-things solution for food safety and quality control: A pilot project in china. *Journal of Industrial Information Integration*; 31-7: (2016)
- 111. Parastar, H.; Shaye, H. Mvc app: A smartphone application for performing chemometric methods. *Chemometrics and Intelligent Laboratory Systems*; 147105-110: (2015)
- 112. Qian, J.-P.; Yang, X.-T.; Wu, X.-M.; Zhao, L.; Fan, B.-L.; Xing, B. A traceability system incorporating 2d barcode and rfid technology for wheat flour mills. *Computers and Electronics in Agriculture*; 8976-85: (2012)
- 113. Lu, L.; Hu, X.; Zhu, Z. Biomimetic sensors and biosensors for qualitative and quantitative analyses of five basic tastes. *TRAC Trends in Analytical Chemistry*; 8758-70: (2017)
- 114. Sliwinska, M.; Wisniewska, P.; Dymerski, T.; Namiesnik, J.; Wardencki, W. Food analysis using artificial senses. *Journal of Agricultural* and Food Chemistry; **62**(7): 1423-1448:

(2014)

- Cetó, X.; Voelcker, N.H.; Prieto-Simón, B. Bioelectronic tongues: New trends and applications in water and food analysis. *Biosensors and Bioelectronics*; 79608-626: (2016)
- 116. Peris, M.; Escuder-Gilabert, L. Electronic noses and tongues to assess food authenticity and adulteration. *Trends in Food Science & Technology*; 5840-54: (2016)
- 117. Facure, M.H.M.; Mercante, L.A.; Mattoso, L.H.C.; Correa, D.S. Detection of trace levels of organophosphate pesticides using an electronic tongue based on graphene hybrid nanocomposites. *Talanta*; 16759-66: (2017)
- Dong, Q.; Du, L.; Zhuang, L.; Li, R.; Liu, Q.; Wang, P. A novel bioelectronic nose based on brain-machine interface using implanted electrode recording in vivo in olfactory bulb. *Biosensors and Bioelectronics*; 49263-269: (2013)
- 119. Glass, J.B.; Chen, S.; Dawson, K.S.; Horton, D.R.; Vogt, S.; Ingall, E.D.; Twining, B.S.; Orphan, V.J. Trace metal imaging of sulfatereducing bacteria and methanogenic archaea at single-cell resolution by synchrotron x-ray fluorescence imaging. *Geomicrobiology Journal*; 1-9: (2017)

- Zhou, H.; Luo, D.; GholamHosseini, H.; Li, Z.; He, J. Identification of chinese herbal medicines with electronic nose technology: Applications and challenges. *Sensors*; *17*(5): 1073-1073: (2017)
- 121. Di Rosa, A.R.; Leone, F.; Cheli, F.; Chiofalo, V. Fusion of electronic nose, electronic tongue and computer vision for animal source food authentication and quality assessment – a review. *Journal of Food Engineering*; 21062-75: (2017)
- 122. Nguyen-Boisse, T.T.; Saulnier, J.; Jaffrezic-Renault, N.; Lagarde, F. Highly sensitive conductometric biosensors for total lactate, dand I-lactate determination in dairy products. *Sensors and Actuators B: Chemical*; 179232-239: (2013)
- 123. Vargas, E.; Conzuelo, F.; Ruiz, A.M.; Campuzano, S.; Ruiz-Valdepeñas Montiel, V.; González de Rivera, G.; López-Colino, F.; Reviejo, J.Á.; Pingarrón, M.J. Automated bioanalyzer based on amperometric enzymatic biosensors for the determination of ethanol in low-alcohol beers. *Beverages*; 3(2): (2017)
- Scognamiglio, V.; Arduini, F.; Palleschi, G.; Rea, G. Biosensing technology for sustainable food safety. *TRAC Trends in Analytical Chemistry*; 621-10: (2014)