



Article Using a Smartphone-Based Colorimetric Device with Molecularly Imprinted Polymer for the Quantification of Tartrazine in Soda Drinks

Christian Jacinto ^{1,2}, Ily Maza Mejía ^{1,2}, Sabir Khan ^{1,2,3,4}, Rosario López ^{1,2}, Maria D. P. T. Sotomayor ^{3,*} and Gino Picasso ²

- ¹ Laboratory of Instrumental Analysis Environment, Faculty of Sciences, National University of Engineering, Av. Tupac Amaru 210, Rimac 15333, Lima, Peru; christian@uni.edu.pe (C.J.); imaza@uni.edu.pe (I.M.M.); sabir@ufersa.edu.br (S.K.); rclopez@uni.edu.pe (R.L.)
- ² Technology of Materials for Environmental Remediation Group (TecMARA), Faculty of Sciences, National University of Engineering, Av. Tupac Amaru 210, Rimac 15333, Lima, Peru; gpicasso@uni.edu.pe

³ Chemistry Institute-Araraquara-SP, São Paulo State University (UNESP), Araraquara 14801-900, Brazil

- ⁴ Department of Natural Sciences, Mathematics, and Statistics, Federal Rural University of the Semi-Arid, Mossoro 59625-900, Brazil
- * Correspondence: m.sotomayor@unesp.br

Abstract: The present study reports the development and application of a rapid, low-cost in-situ method for the quantification of tartrazine in carbonated beverages using a smartphone-based colorimetric device with molecularly imprinted polymer (MIP). The MIP was synthesized using the free radical precipitation method with acrylamide (AC) as the functional monomer, N,N'methylenebisacrylamide (NMBA) as the cross linker, and potassium persulfate (KPS) as radical initiator. The smartphone (RadesPhone)-operated rapid analysis device proposed in this study has dimensions of $10 \times 10 \times 15$ cm and is illuminated internally by light emitting diode (LED) lights with intensity of 170 lux. The analytical methodology involved the use of a smartphone camera to capture images of MIP at various tartrazine concentrations, and the subsequent application of the Image-J software to calculate the red, green, blue (RGB) color values and hue, saturation, value (HSV) values from these images. A multivariate calibration analysis of tartrazine in the range of 0 to 30 mg/L was performed, and the optimum working range was determined to be 0 to 20 mg/L using five principal components and a limit of detection (LOD) of 1.2 mg/L was obtained. Repeatability analysis of tartrazine solutions with concentrations of 4, 8, and 15 mg/L (n = 10) showed a coefficient of variation (% RSD) of less than 6%. The proposed technique was applied to the analysis of five Peruvian soda drinks and the results were compared with the UHPLC reference method. The proposed technique showed a relative error between 6% and 16% and % RSD lower than 6.3%. The results of this study demonstrate that the smartphone-based device is a suitable analytical tool that offers an on-site, cost-effective, and rapid alternative for the quantification of tartrazine in soda drinks. This color analysis device can be used in other molecularly imprinted polymer systems and offers a wide range of possibilities for the detection and quantification of compounds in various industrial and environmental matrices that generate a color change in the MIP matrix.

Keywords: smartphone; molecularly imprinted polymers (MIP); tartrazine; colorimetric detection; UHPLC

1. Introduction

Tartrazine (TAR) is a highly popular yellow dye that is widely applied in a variety of food and beverage products, including sweets, juices, and general food products. Several studies reported in the literature have raised serious concerns regarding the potential health risks associated with the use of TAR in food products; these studies have linked Tartrazine to a wide range of negative health problems, including headaches, neurotoxicity,



Citation: Jacinto, C.; Maza Mejía, I.; Khan, S.; López, R.; Sotomayor, M.D.P.T.; Picasso, G. Using a Smartphone-Based Colorimetric Device with Molecularly Imprinted Polymer for the Quantification of Tartrazine in Soda Drinks. *Biosensors* 2023, *13*, 639. https://doi.org/ 10.3390/bios13060639

Received: 8 May 2023 Revised: 5 June 2023 Accepted: 6 June 2023 Published: 9 June 2023



Copyright: © 2023 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (https:// creativecommons.org/licenses/by/ 4.0/). genotoxicity, and even cancer. Bearing that in mind, consumers are advised to take note of the potential risks associated with tartrazine and consider limiting their intake of products that contain this dye [1]. The recommended daily intake of tartrazine in humans ranges between 0 and 7 mg/L5 mg/kg of body weight [2]. Azo dyes, such as tartrazine, are known to undergo reductive biotransformation during metabolism, and this can lead to the release of toxic and potentially carcinogenic compounds in the body. The presence of these compounds in the human body affects the DNA, as it causes mutagenic effects, and increases the risk of cancer [3]. Therefore, regulating the levels of tartrazine in food products is essentially crucial as it helps to ensure food safety and protect public health [4].

Several studies have proposed a plethora of techniques for the determination of tartrazine. High-performance liquid chromatography (HPLC) is the technique most widely applied for TAR determination due to its high sensitivity and selectivity [5]; however, this technique is found to be costly and its application often requires the use of significant amounts of toxic solvents [6,7]. Spectrophotometric methods are also widely used for TAR determination, though their specificity may be limited due to spectral overlap with other species in the matrix [8,9]. Among the more recent analytical techniques that have been applied for TAR determination are paper chromatography [10] and voltammetry [11,12]. A comprehensive outline of other methods used for TAR determination in food products can be found in Rovina's work [13].

Molecularly imprinted polymers (MIPs) are synthetic analogs of biological antibodyantigen systems [14]; these materials function via a "lock and key" mechanism with the selective binding of the molecule to which they are designed during synthesis. MIPs offer the potential specificity and selectivity of biological receptors with the explicit advantages of durability under environmental conditions and at low costs [15]. MIPs can be employed for a wide range of purposes, including the separation and purification of chemical compounds [16–18], catalysis [19], detection and quantification of substances in complex samples [20–22], and in the fabrication of sensory and biosensor devices [23–25]. MIPs can also be used as optical sensors for dyes, where they modulate the changes in the optical properties of the polymer upon binding to the dye. For this reason, they have been used for the determination of textile [26,27] and food colorants [28–30].

The use of smartphones in digital color analysis is an innovative technique that allows for accurate and easy measurement and effective comparison of the color of different objects or surfaces [31,32]. Under this technique, the smartphone camera is used to capture the images of colored objects which are intended to be analyzed. In addition, image processing algorithms are applied aiming at extracting the numerical color values in different color spaces, including the following: Red, green, blue (RGB); cyan, magenta, yellow, and key (black) (CMYK); hue, saturation, value (HSV); and CIELAB [33]. Specific image processing software, such as Adobe Photoshop, Image-J, MATLAB, etc. are used for color quantification under the appropriate color space. A relationship between the quantified image data and the analyte concentration can then be established using multivariate calibration models, such as partial least-squares (PLS) [34] and machine learning [35]. Currently, the widespread use of mobile applications (apps) and the high computing power of smartphones have led to an increasingly growing application of these devices in digital color analyses for point-of-care testing (POCT) [32]. POCT involves the conduct of rapid and simple sample detection tests using a portable instrument at the sampling site which helps in significantly reducing the time and cost of the test [36]. This testing technique has been used in various areas, including clinical diagnosis [37,38], food safety [39,40], environmental testing [41,42], among others [43,44].

Previous works reported in the literature have employed MIP for tartrazine determination analyses. Arabzadeh [45] synthesized and used MIP for tartrazine adsorption. Córdova [29] reported the use of modified MIP in core-shell type nanoparticles for tartrazine determination in carbonated beverages through reflectance. This work is the first to use MIP for colorimetric quantification of tartrazine using a smartphone. Other similar works have been reported to determine antibiotics, such as sulfamethoxazole [46], fungal toxins, such as aflatoxin B1 [47], and dyes, such as malachite green [48], basic red 9 [49], and Erythrosine B [50]. In our present work, MIP will be used for tartrazine determination in soda drinks through the application of color analysis. To conduct this analysis, an image capture device (RadesPhone) which uses a smartphone camera will be constructed, and the image of the MIP exposed to aqueous samples of tartrazine will be obtained. After obtaining the digital image of the MIP, the appropriate software will be used to extract the values of RGB, HSV, L, and I color channels, and PLS multivariate regression analysis will be conducted to determine tartrazine in the samples. The application of color analysis with the constructed device opens up a wide range of possibilities for the use of different types and forms of MIP, as well as for the rapid detection and quantification of different compounds in industrial and environmental matrices at low-cost and without the need of specialized personnel.

2. Materials and Methods

2.1. Chemic als and Reagents

All reagents used in this work were of analytical grade. Solutions were prepared in ultrapure water (18 M Ω at 25 °C). Tartrazine (TAR), N,N'-methylenebisacrylamide (NMBA), acrylamide (AA), and potassium persulfate (KPS) were purchased from Sigma-Aldrich. Sodium hydroxide, methanol (HPLC grade), concentrated ammonia solution (23–28%), and hydrochloric acid (10%) were purchased from Merck.

The infrared spectrum was obtained using a Shimadzu IRPrestige 21 in ATR mode to show the functionalization and polymerization of the MIP through the presence of different functional groups. The light intensity measurement was performed with luxmeter UT383 Mini Light Meters. UHPLC was performed using a Thermo ScientificTM VanquishTM Duo UHPLC system with a C18 column of 150×4.6 mm, UV detector, and particle size of 5 µm.

2.2. Synthesis of Molecularly Imprinted Polymer (MIP)

The MIP was prepared based on the precipitation method using the general synthesis procedure proposed by Arabzadeh [45], but it was modified to a molar ratio of template, functional monomer, and cross-linking agent of 1:2:100, respectively. For this, 0.1 mmol of tartrazine (template) and 0.2 mmol of acrylamide (functional monomer) were mixed in 40 mL of ultrapure water (porogenic solvent) and remained under agitation for 2 h. The solution was bubbled with N₂ for 10 min to remove oxygen, and then 10 mmol of N,N'-methylenebis (acrylamide) (cross linker) was added. Next, 0.185 mmol of potassium persulfate (initiator) was added, and the solution was continuously treated with nitrogen for another 10 min. Finally, the reaction flask was placed in a water bath at 60 °C for 22 h under constant agitation. After obtaining the MIP, tartrazine was extracted using a Soxhlet extraction system with an 8:2 (v/v) water/ammonia solution until the washing solutions showed no UV signal of tartrazine. The obtained material was washed with a mixture of ethanol and water, dried at room temperature in a vacuum desiccator for 1 day, ground in an agate mortar, and passed through a 250 µm sieve (No. 60).

2.3. Adsorption Process

To evaluate the effect of pH and adsorption time, the MIP prepared under the new conditions was used. To adjust the procedure and obtain an amount of MIP that could be used in image analysis, 15 mg of MIP and 15 mL of tartrazine sample at pH 3.0 were placed in a 20 mL flask, agitated in a homogenizer for 10 min and filtered through Whatman No. 40 filter paper. The solid was removed from the paper and dried in an oven at 60 °C to make it available for image analysis.

2.4. Construction of the RadesPhone Device

Both the sample holder and the imaging device were constructed using a 3D printer. The sample holder was a 3×3 cm platform with a thickness of 3 mm, and it had a space for the solid sample in the center (Figure 1A). The RadesPhone imaging device was

constructed with dimensions of $15 \times 15 \times 10$ cm and was completely painted black on the inside (Figure 1B). A previous study determined that the dimensions of the imaging device and the height of the smartphone did not influence image capture [49]. Two arrays of LED lamps were placed at the top corners of the device providing a light intensity of 170 lux, which operated by connecting them to the power outlet.



Figure 1. RadesPhone device for image acquisition. (A) Sample holder with dimensions of 3×3 cm and height of 3 mm; (B) camera for image acquisition.

2.5. Image Acquisition with RadesPhone

A Samsung Galaxy S20 FE smartphone, model SM-G780F, with a 12 MP rear camera was used for image capture. The image capture settings were adjusted with the camera sensor sensitivity set to automatic ISO, a camera shutter speed of 1/60 s, an exposure EV of 0, and an automatic white balance. The sample holder containing the MIP exposed to tartrazine solutions was placed in the corresponding RadesPhone imaging device for image capture (Figure 2).



Figure 2. RadesPhone device with the smartphone for imaging the MIP on the sample holder.

2.6. Analysis of Soda Drinks by UHPLC

The reference quantities of tartrazine in the carbonated beverages were quantified using UHPLC measurements as described above. The mobile phase consisted of a mixture of methanol and water (20:80 v/v), with a flow rate of 1 mL/min, a wavelength of 430 nm, and an injected sample volume of 20 μ L.

3. Results

3.1. Synthesis of MIP Obtained by Precipitation

The MIP prepared by Arabzadeh [45] presented the characteristic yellow color of tartrazine, even after being washed for a week. This background color reduces the sensitivity of the MIP for its use in subsequent color analysis when exposed to tartrazine solutions. To improve the sensitivity of the MIP, it was proposed to modify the template:MF:ME molar ratio to 1:2:100. This proportion increases the amount of ME in the MIP synthesis, which produces a significant decrease in the intensity of the yellow color of tartrazine in the resulting MIP. This new MIP was employed in tartrazine adsorption studies for subsequent color analysis using a smartphone.

Some characterization tests were performed on the newly synthesized MIP to verify that the MIP was obtained, as well as some tartrazine adsorption tests in aqueous solutions. Figure 3A shows the infrared spectra of the MIP and tartrazine, displaying the characteristic peaks of the polymer at 1675 cm⁻¹ and 3342 cm⁻¹, corresponding to the C=O and -NH-vibration peaks, respectively. Additionally, tartrazine exhibits peaks at 3453, 1472, and 1176 cm⁻¹ corresponding to the hydroxyl (-OH), sulfonate (-SO₃Na), and carboxylate (-COONa) linkages, respectively, and shows that the MIP does not exhibit appreciable signals of tartrazine. Figure 3B compares the spectra of the MIP with N-N'-methylene-bisacrylamide (cross linker). The C-H doublet peak of the out-of-plane deformation of the vinyl group (992 and 910 cm⁻¹) disappears in the MIP, indicating that polymerization was carried out through the C=C double bond of this monomer. The C=O and -NH peaks of the monomer (at 1661 cm⁻¹ and 3314 cm⁻¹) remain in the MIP since the cross linker is present in a higher proportion in the MIP.

The effect of pH is studied to achieve the maximum adsorption capacity of the polymer (Figure 4). It has been observed that the maximum adsorption occurs at a pH of 3.0. This preference for adsorption at pH 3.0 is due to the fact that the sulfonate group of the polymer has a pKa of 3.2, indicating that it is in its ionic form at this pH. Under these conditions, the interaction between tartrazine and the MIP primarily occurs through hydrogen bonding and electrostatic interactions. The electrostatic interactions are established between the non-dissociated amide group present on the polymer surface and the sulfonate group of tartrazine [51].

$$P - NH_2 + H^+ \leftrightarrow P - NH_3^+$$

$$T - SO_3Na \rightarrow T - SO_3^- + Na^+$$

$$T - SO_3^- + P - NH_3^+ \rightarrow T - SO_3^- + NH_3 - P$$

Molecular imprinting also has an effect on the adsorption capacity of the MIP for tartrazine. This effect has been evaluated by studying the selectivity of the MIP compared to another similar dye, such as Sunset Yellow CFC. In a study conducted by Arabzadeh [45], the selectivity factor of the MIP for tartrazine relative to Sunset Yellow CFC was determined to be 22.5, while the NIP showed a value of 0.73. These results highlight the impact of molecular imprinting on the selective adsorption of tartrazine.



Figure 3. ATR-mode FTIR spectra of (**A**) MIP and tartrazine, (**B**) MIP and N,N'-methylbisacrylamide cross-linking monomer.

The effect of contact time of MIP on tartrazine adsorption was evaluated at different times up to 3 h and is shown in Figure 5. The adsorption rate of tartrazine was rapid initially, with maximum adsorption achieved at 10 min. It was also observed that at contact times greater than 30 min, desorption of tartrazine occurred, resulting in a decrease in % removal, although after 60 min it increased again.



Figure 4. Effect of pH on tartrazine removal. Measurements were made using 6 mg of MIP with 2 mL of tartrazine solution and stirred for 1 h.



Figure 5. Effect of MIP contact time on tartrazine adsorption. Measurements were made using 6 mg of MIP with 2 mL of tartrazine solution at pH 3.0.

The general procedure for color analysis using the RadesPhone device is as follows. The MIP exposed to the tartrazine samples is filtered and dried in an oven at 60 °C. Subsequently, it is placed in the sample holder along with the RadesPhone device (Figure 6). The digital image of the sample was obtained using the smartphone camera. With the assistance of the free software Image-J, we obtained the RGB values from a 16×16 pixel region in the image. By converting these values to the other channels (using Equations (1)–(5)), we obtained the corresponding HSV, L, and I values. With the values of these channels, we perform a multivariate regression PLS and select the number of components that provide the lowest prediction error. Moreover, we select the working range and evaluate the repeatability and detection limit, as well as the application to soft drink samples.



Figure 6. General scheme of color analysis for the determination of tartrazine using MIP. (**A**) Capturing 16×16 pixel image of MIP with tartrazine; (**B**) decomposition of the image into RGB; (**C**) transformation to HSV, L, and I color channels; (**D**) obtaining the table of values of the color channels; (**E**) multivariate regression with the values of the channels obtained from each sample.

Since RGB values are not the only ways to represent color, conversion to other channels is performed. The H, S, and V channels represent the hue, saturation, and value components of an image. The hue channel describes the color tone, while saturation refers to the amount of color present in the image, and the value describes the intensity or brightness of the color. The L channel is a component of the CIELAB color space used to measure color differences perceived by the human eye and describes the lightness of a color. The I channel refers to the intensity component in the HSI (hue, saturation, intensity) color space, which is often used to describe the image in terms of color and brightness. RGB values are obtained from the image using Image-J. The other HSV, L, and I values are calculated from Equations (1)–(5):

V

$$H = \begin{cases} undefined, & if \max = min \\ 60^{\circ} \times \frac{G-B}{\max - min} + 0^{\circ}, & if \max = R \\ 60^{\circ} \times \frac{G-B}{\max - min} + 360^{\circ}, & if \max = R \\ 60^{\circ} \times \frac{B-R}{\max - min} + 120^{\circ}, & if \max = G \\ 60^{\circ} \times \frac{R-G}{\max - min} + 240^{\circ}, & if \max = B \end{cases}$$
(1)

$$S = \begin{cases} 0, & if \max = 0\\ 1 - \frac{\min}{\max}, & in another case \end{cases}$$
(2)

$$= max$$
 (3)

$$L = \frac{1}{2}(\max + \min) \tag{4}$$

$$I = \frac{R+G+B}{3} \tag{5}$$

These values can be obtained from websites, such as https://www.peko-step.com/ es/tool/hslrgb.html. A detailed explanation of what each color channel represents in its color space is described in the literature [34,52,53].

The RGB, HSV, L, and I values obtained for the standards and samples are presented in Table 1:

R В v L Sample G I н S 1 ppm 194.5 ± 2.4 174.7 ± 2.1 28.9 ± 23.4 139.1 ± 5.4 52.3 ± 0.6 84.3 ± 12.3 76.0 ± 1.0 43.3 ± 4.0 5 ppm 208.1 ± 1.2 172.5 ± 2.8 3.1 ± 0.1 128.0 ± 1.2 49.3 ± 0.6 98.0 ± 0.0 81.0 ± 0.0 41.0 ± 0.0 10 ppm 206.5 ± 1.3 151.8 ± 0.7 1.9 ± 0.1 120.0 ± 0.1 43.7 ± 0.6 99.0 ± 0.0 80.3 ± 0.6 40.3 ± 0.6 15 ppm 198.9 ± 0.8 140.8 ± 1.3 2.0 ± 0.0 114.0 ± 0.7 42.3 ± 0.6 98.0 ± 0.0 77.3 ± 0.6 39.0 ± 0.0 106.9 ± 0.2 42.0 ± 0.0 36.3 ± 0.6 132.7 ± 0.6 0.9 ± 0.1 72.7 ± 0.6 20 ppm 187.0 ± 1.0 99.0 ± 0.0 105.8 ± 0.2 41.3 ± 0.6 186.0 ± 1.0 130.0 ± 1.0 1.3 ± 0.6 98.7 ± 0.6 72.3 ± 0.6 36.3 ± 0.6 25 ppm 1.7 ± 0.6 102.6 ± 0.7 41.0 ± 0.0 70.3 ± 0.6 36.7 ± 0.6 181.0 ± 1.0 125.0 ± 1.0 98.3 ± 0.6 30 ppm 203.8 ± 1.7 42.5 ± 1.4 78.4 ± 2.2 148.0 ± 1.8 1.9 ± 0.1 116.4 ± 1.6 97.8 ± 1.6 40.5 ± 1.3 M1 192.3 ± 2.0 136.2 ± 1.1 1.5 ± 0.1 111.2 ± 1.6 41.5 ± 1.3 97.3 ± 2.0 74.5 ± 1.3 37.2 ± 1.0 M2 М3 192.4 ± 1.2 137.4 ± 1.5 1.6 ± 0.5 111.1 ± 0.8 40.8 ± 0.8 94.4 ± 1.5 75.4 ± 1.4 37.1 ± 1.0 M4 193.9 ± 1.7 136.9 ± 1.7 1.4 ± 0.3 110.8 ± 0.9 42.9 ± 1.0 97.6 ± 1.4 75.2 ± 1.1 36.3 ± 1.5 150.4 ± 1.6 1.6 ± 0.5 117.6 ± 1.4 97.3 ± 1.4 M5 205.0 ± 1.1 43.4 ± 0.5 80.6 ± 1.3 40.8 ± 1.1

Table 1. RGB, HSV, L, and I results obtained for tartrazine standards and soft drink samples.

3.2. Multivariate PLS Regression

PLS is a multivariate calibration method that seeks factors that explain the highest possible covariance between a set of predictor variables, X, and response variables, Y. A training matrix was developed consisting of 21 samples with concentrations of 0, 5, 10, 15, 20, 25, and 30 mg/L of tartrazine at pH 3.0, prepared in triplicate. PLS first performs a principal component analysis (PCA) to reduce the dimensionality of the original variables. A regression model is then developed using these principal components. Cross-validation is used to evaluate the performance of the model, which assesses the ability of the PLS model to predict the responses of a data sample that was not used in calibration. To evaluate the number of principal components that provide the best prediction, a graph was plotted showing the use of these components with the coefficient of determination R^2 between the measured value and the predicted value. The number of components that provide the highest R² value will define the number of components to use. Figure 7A shows the graph for selecting the number of principal components, which indicates that four components provide the best regression model. After selecting the four components to use, the PLS model is evaluated by plotting the measured values against the predicted values (Figure 7B). This graph shows that there is no good linear fit beyond 20 mg/L, and thus the working range will be reduced to 20 mg/L.

The PLS model was re-evaluated using the range of 0 to 20 mg/L and the results are shown in Figure 8. It can be seen that the model improved and it will be used for future sample predictions. A summary of the performance results of both evaluated models is shown in Table 2.

Table 2. Summary of R² values according to the studied concentration range.

Range (mg/L)	Number of Components	R² (Calibration)	R² (Validation)	
0–30	4	0.980973	0.967121	
0–20	5	0.995304	0.967887	



Figure 7. Performance graphs of the PLS regression model using the range of 0 to 30 mg/L of tartrazine. (**A**) Graph of the number of principal components used with the R² calibration and validation; (**B**) graph between the actual (real) and predicted (calculated) values by the PLS model using four principal components.

Table 3 shows the performance metrics of the obtained PLS model. Repeatability was tested at different concentration levels, showing that variability increases as concentration decreases. The detection limit was calculated based on the standard deviation of the regression of the PLS model between measured and predicted values [54]. This value is related to the initial yellow color of the MIP and, as previously mentioned, the cross linker ratio was modified during synthesis to obtain an MIP suitable for color analysis. The results obtained are satisfactory, demonstrating that color analysis using the RadesPhone device and the MIP for tartrazine determination is appropriate.



Figure 8. Performance graphs of the PLS regression model using the range of 0 to 20 mg/L of tartrazine. (**A**) Graph of the number of principal components used with the R² calibration and validation; (**B**) graph between the actual (real) and predicted (calculated) values by the PLS model using five principal components.

Range (mg/L)	Number of Components	Repeatability (<i>n</i> = 10)		Limit of Detection (LOD) (mg/L)	
		mg/L	Found (mg/L)	% RSD	
0–20	4	4 8 15	4.6 7.2 14.5	5.3 3.3 4.6	1.2



The results of the proposed RadesPhone method were compared to those obtained by UHPLC for real samples of soda drinks. A typical chromatogram and the analytical curve is shown in Figure 9.

Figure 9. UHPLC chromatogram of 1 mg/L tartrazine, mobile phase methanol and water (20:80 v/v), flow rate of 1 mL/min, wavelength of 430 nm, and injected sample volume of 20 µL. Thermo ScientificTM VanquishTM Duo UHPLC system liquid chromatograph with 150 × 4.6 mm C18 column, UV detector, and particle size of 5 µm.

The comparison results are shown in Table 4. Despite the low concentrations being worked with, the results are satisfactory. In general, the proposed method's results are slightly higher (except for M2) than those of the UHPLC reference method, possibly due to the adsorption of other yellow dyes that the samples may contain, such as sunset yellow, which is common in these types of foods.

Am	ount of Tartrazine Fo (mg/L, <i>n</i> = 3)	und			
Sample	RadesPhone	UHPLC	Relative Error (%)	RSD (%)	
M1	13.6 ± 0.8	12.8 ± 0.2	6.3	5.9	
M2	17.4 ± 0.6	19.4 ± 0.3	-10.3	3.5	
M3	11.2 ± 0.7	9.6 ± 0.2	16.7	6.3	
M4	18.1 ± 0.7	16.8 ± 0.1	7.7	4.6	
M5	14.0 ± 0.6	12.6 ± 0.3	15.9	4.3	

Table 4. Amount of tartrazine found in soda drinks by the proposed RadesPhone and UHPLC method.

This method is comparable to the results obtained by Córdova [29] in his study on soda drinks samples. In that study, results were reported with a relative error between 14% and 23% (alternative method between 7.7% and 16%), and a relative standard deviation between 12% and 20% (alternative method between 3.5% and 6.3%).

4. Conclusions

In the present work, a color analysis device (RadesPhone) using MIP has been developed for the quantitative analysis of tartrazine in soda drinks. The MIP used in this study was synthesized using the precipitation method and had previously been employed in studies involving the adsorption of tartrazine in food samples. In this work, the same MIP was utilized for both the adsorption of tartrazine and the subsequent color analysis using a smartphone. Color analysis has been demonstrated to be a powerful tool for quantifying tartrazine in carbonated beverages using the RadesPhone device and MIP. The method, which involves transforming the digital image into RGB color values and employing multivariate regression PLS, has demonstrated excellent repeatability and low prediction errors. Comparing the results obtained with a similar study that utilized a modified MIP for tartrazine determination in soda drinks, the proposed method with the smartphone exhibited lower prediction errors (23% relative error in the modified MIP method compared to 16% in the proposed smartphone method) and reduced variability (20% RSD in the similar method compared to 6.3% in the proposed method). The utilization of smartphones in color analysis with the RadesPhone device enables the development of a simple and cost-effective method for determining soda drinks. This approach eliminates the need for solvents or specialized equipment and allows for untrained personnel to perform the analysis, following the principles of point-of-care testing (POCT). The results obtained by color analysis were slightly higher than those obtained by UHPLC, possibly due to interferences present in the color analysis and the higher selectivity of the chromatographic method. However, the RadesPhone device and the color analysis methodology with multivariate analysis offer several opportunities for the use of MIP in different systems and analytes, allowing for the development of fast, cost-effective, and easy-to-use methods of analysis. Furthermore, the developed MIP can be prepared to selectively adsorb and improve its sensitivity by obtaining intensely colored compounds. MIP can be supported on various surfaces, such as paper, glass, PET, and others. After immersing the MIP-supported surface in an aqueous solution of the analyte, selective adsorption of a colored compound takes place. Alternatively, the surface can be treated to develop color, followed by direct analysis using the image analysis device. This approach enhances the practicality of the analysis process.

Author Contributions: Conceptualization, S.K., R.L., M.D.P.T.S. and G.P.; formal analysis, S.K. and G.P.; funding acquisition, R.L. and M.D.P.T.S.; investigation, C.J.; methodology, C.J. and I.M.M.; project administration, R.L. and G.P.; resources, M.D.P.T.S.; software, C.J.; validation, C.J., I.M.M. and S.K.; writing—original draft, C.J., I.M.M., S.K., R.L. and G.P.; writing—review and editing, M.D.P.T.S. All authors have read and agreed to the published version of the manuscript.

Funding: This project was supported by PROYECTO #067-FONDECYT-2021 and CNPq-Brazil (301728/2019-4).

Institutional Review Board Statement: Not applicable.

Informed Consent Statement: Not applicable.

Data Availability Statement: Not applicable.

Acknowledgments: The authors wish to express their sincerest gratitude to São Paulo State University (UNESP), Chemistry Institute-Araraquara-SP, Brazil, Department of Natural Sciences, Mathematics, and Statistics, Federal Rural University of the Semi-Arid, Mossoró, Rio Grande do Norte, Brazil, and Faculty of Sciences, National University of Engineering of Lima, Peru, and project PE501080434-2022-PROCIENCIA.

Conflicts of Interest: The authors declare no conflict of interest.

References

- Yamjala, K.; Nainar, M.S.; Ramisetti, N.R. Methods for the analysis of azo dyes employed in food industry—A review. *Food Chem.* 2016, 192, 813–824. [CrossRef] [PubMed]
- EFSA Panel on Food Additives and Nutrient Sources Added to Food. Scientific Opinion on the re-evaluation Tartrazine (E 102). EFSA J. 2009, 7, 1331. [CrossRef]
- Atlı Şekeroğlu, Z.; Güneş, B.; Kontaş Yedier, S.; Şekeroğlu, V.; Aydın, B. Effects of tartrazine on proliferation and genetic damage in human lymphocytes. *Toxicol. Methods* 2017, 27, 370–375. [CrossRef] [PubMed]

- 4. Liu, L.; Sun, H.; Xiao, L.; Yang, Z.Q.; Han, J.; Gong, X.; Hu, Q. Development of a highly sensitive fluorescence method for tartrazine determination in food matrices based on carbon dots. *Anal. Bioanal. Chem.* **2021**, *413*, 1485–1492. [CrossRef]
- 5. Dinç, E.; Aktas, A.H.; Baleanu, D.; Ustundag, O. Simultaneous determination of tartrazine and allura red in commercial preparation by chemometric HPLC method. *J. Food Drug Anal.* **2006**, *14*, 284. [CrossRef]
- 6. Alp, H.; Başkan, D.; Yaşar, A.; Yaylı, N. Simultaneous determination of sunset yellow FCF, allura red AC, quinoline yellow WS, and tartrazine in food samples by RP-HPLC. *J. Chem.* **2018**, 2018, 6486250. [CrossRef]
- Agbokponto, J.E.; Kpaibe, A.P.S.; Yemoa, L.A.; Assanhou, A.G.; Ganfon, H.; Gbassi, G.K.; Aké, M.; Amin, N.C.; Agbokponto, J.E.; Kpaibe, A.P.S.; et al. Simultaneous Determination by HPLC-UV Vis of Tartrazine and Sunset Yellow in Soft Drinks Sold in Benin. *Am. J. Anal. Chem.* 2022, 13, 277–288. [CrossRef]
- 8. Lawal, A.; Gwaram, N.S.; Abdulkarim, S. The Spectrophotometric Determination of Tartrazine in Some Selected Beverages: A Case Study of Katsina Town, Nigeria. *FUDMA J. Sci.* **2020**, *4*, 685–689.
- 9. Sahraei, R.; Farmany, A.; Mortazavi, S.S. A nanosilver-based spectrophotometry method for sensitive determination of tartrazine in food samples. *Food Chem.* **2013**, *138*, 1239–1242. [CrossRef]
- 10. Gharaghani, F.M.; Akhond, M.; Hemmateenejad, B. A three-dimensional origami microfluidic device for paper chromatography: Application to quantification of Tartrazine and Indigo carmine in food samples. *J. Chromatogr. A* **2020**, *1621*, 461049. [CrossRef]
- 11. Lipskikh, O.I.; Korotkova, E.I.; Barek, J.; Vyskocil, V.; Saqib, M.; Khristunova, E.P. Simultaneous voltammetric determination of Brilliant Blue FCF and Tartrazine for food quality control. *Talanta* **2020**, *218*, 121136. [CrossRef]
- 12. Ziyatdinova, G.K.; Budnikov, H.C. Voltammetric Determination of Tartrazine on an Electrode Modified with Cerium Dioxide Nanoparticles and Cetyltriphenylphosphonium Bromide. *J. Anal. Chem.* **2022**, 77, 664–670. [CrossRef]
- 13. Rovina, K.; Siddiquee, S.; Shaarani, S.M. A review of extraction and analytical methods for the determination of tartrazine (E 102) in foodstuffs. *Crit. Rev. Anal. Chem.* **2017**, *47*, 309–324. [CrossRef]
- 14. Parisi, O.I.; Francomano, F.; Dattilo, M.; Patitucci, F.; Prete, S.; Amone, F.; Puoci, F. The Evolution of Molecular Recognition: From Antibodies to Molecularly Imprinted Polymers (MIPs) as Artificial Counterpart. *J. Funct. Biomater.* **2022**, *13*, 12. [CrossRef]
- 15. Belbruno, J.J. Molecularly Imprinted Polymers. Chem. Rev. 2019, 119, 94–119. [CrossRef]
- López, R.; Khan, S.; Torres, S.E.; Wong, A.; Sotomayor, M.D.P.T.; Picasso, G. Synthesis and Characterization of Magnetic Molecularly Imprinted Polymer for the Monitoring of Amoxicillin in Real Samples Using the Chromatographic Method. *Magnetochemistry* 2023, *9*, 92. [CrossRef]
- 17. Barhate, C.L.; Donnell, A.F.; Davies, M.; Li, L.; Zhang, Y.; Yang, F.; Black, R.; Zipp, G.; Zhang, Y.; Cavallaro, C.L.; et al. Microscale purification in support of high-throughput medicinal chemistry. *Chem. Commun.* **2021**, *57*, 11037–11040. [CrossRef]
- Srivastava, N.; Singh, A.; Kumari, P.; Nishad, J.H.; Gautam, V.S.; Yadav, M.; Bharti, R.; Kumar, D.; Kharwar, R.N. Advances in extraction technologies: Isolation and purification of bioactive compounds from biological materials. *Nat. Bioact. Compd. Technol. Adv.* 2021, 409–433. [CrossRef]
- 19. Li, X.; Wan, J.; Wang, Y.; Yan, Z.; Chi, H.; Ding, S. Mechanism of accurate recognition and catalysis of diethyl phthalate (DEP) in wastewater by novel MIL100 molecularly imprinted materials. *Appl. Catal. B Environ.* **2020**, *266*, 118591. [CrossRef]
- Cui, B.; Liu, P.; Liu, X.; Liu, S.; Zhang, Z. Molecularly imprinted polymers for electrochemical detection and analysis: Progress and perspectives. J. Mater. Res. Technol. 2020, 9, 12568–12584. [CrossRef]
- Cengiz, N.; Guclu, G.; Kelebek, H.; Capanoglu, E.; Selli, S. Application of Molecularly Imprinted Polymers for the Detection of Volatile and Off-Odor Compounds in Food Matrices. ACS Omega 2021, 7, 15258–15266. [CrossRef] [PubMed]
- Arshad, U.; Mujahid, A.; Lieberzeit, P.; Afzal, A.; Bajwa, S.Z.; Iqbal, N.; Roshan, S. Molecularly imprinted polymeric coatings for sensitive and selective gravimetric detection of artemether. *RSC Adv.* 2020, *10*, 34355–34363. [CrossRef] [PubMed]
- 23. López, R.; Khan, S.; Wong, A.; del Pilar Taboada Sotomayor, M.; Picasso, G. Development of a New Electrochemical Sensor Based on Mag-MIP Selective Toward Amoxicillin in Different Samples. *Front. Chem.* **2021**, *9*, 146. [CrossRef] [PubMed]
- Mortari, B.; Khan, S.; Wong, A.; Sotomayor, M.D.P.T. A spot test for direct quantification of acid green 16 adsorbed on a molecularly imprinted polymer through diffuse reflectance measurements. *Anal. Methods* 2021, 13, 453–461. [CrossRef] [PubMed]
- Foguel, M.V.; Pedro, N.T.B.; Wong, A.; Khan, S.; Zanoni, M.V.B.; Sotomayor, M.D.P.T. Synthesis and evaluation of a molecularly imprinted polymer for selective adsorption and quantification of Acid Green 16 textile dye in water samples. *Talanta* 2017, 170, 244–251. [CrossRef]
- Foguel, M.V.; Pedro, N.T.B.; Zanoni, M.V.B.; Sotomayor, M.D.P.T. Molecularly Imprinted Polymer (MIP): A Promising Recognition System for Development of Optical Sensor for Textile Dyes. *Procedia Technol.* 2017, 27, 299–300. [CrossRef]
- 27. Mortari, B.; Khan, S.; Wong, A.; Fireman Dutra, R.A.; Taboada Sotomayor, M.D.P.T. Next generation of optodes coupling plastic antibody with optical fibers for selective quantification of Acid Green 16. *Sens. Actuators B Chem.* **2020**, *305*, 127553. [CrossRef]
- George, A.; Rose Cherian, A.; Jacob, B.; Varghese, A.; Maiyalagan, T. Design optimisation and fabrication of amino acid based molecularly imprinted sensor for the selective determination of food additive tartrazine. *Food Chem.* 2023, 404, 134673. [CrossRef]
- Ruiz-Córdova, G.A.; Villa, J.E.L.; Khan, S.; Picasso, G.; Del Pilar Taboada Sotomayor, M. Surface molecularly imprinted core-shell nanoparticles and reflectance spectroscopy for direct determination of tartrazine in soft drinks. *Anal. Chim. Acta* 2021, 1159, 338443. [CrossRef]
- 30. Mohammadinejad, A.; Abouzari-Lotf, E.; Aleyaghoob, G.; Rezayi, M.; Kazemi Oskuee, R. Application of a transition metal oxide/carbon-based nanocomposite for designing a molecularly imprinted poly (l-cysteine) electrochemical sensor for curcumin. *Food Chem.* **2022**, *386*, 132845. [CrossRef]

- 31. Saadati, M. Smartphone-Based Digital Image Analysis for Determination of Some Food Dyes in Commercial Products. *Food Anal. Methods* **2021**, *14*, 2367–2374. [CrossRef]
- 32. Liu, J.; Geng, Z.; Fan, Z.; Liu, J.; Chen, H. Point-of-care testing based on smartphone: The current state-of-the-art (2017–2018). *Biosens. Bioelectron.* **2019**, 132, 17–37. [CrossRef]
- Fan, Y.; Li, J.; Guo, Y.; Xie, L.; Zhang, G. Digital image colorimetry on smartphone for chemical analysis: A review. *Measurement* 2021, 171, 108829. [CrossRef]
- 34. Böck, F.C.; Helfer, G.A.; da Costa, A.B.; Dessuy, M.B.; Ferrão, M.F. PhotoMetrix and colorimetric image analysis using smartphones. *J. Chemom.* **2020**, *34*, e3251. [CrossRef]
- 35. Hassanijalilian, O.; Igathinathane, C.; Doetkott, C.; Bajwa, S.; Nowatzki, J.; Haji Esmaeili, S.A. Chlorophyll estimation in soybean leaves infield with smartphone digital imaging and machine learning. *Comput. Electron. Agric.* **2020**, *174*, 105433. [CrossRef]
- Xing, G.; Ai, J.; Wang, N.; Pu, Q. Recent progress of smartphone-assisted microfluidic sensors for point of care testing. *TrAC Trends Anal. Chem.* 2022, 157, 116792. [CrossRef]
- Zhang, S.; Li, Z.; Wei, Q. Smartphone-based cytometric biosensors for point-of-care cellular diagnostics. *Nanotechnol. Precis. Eng.* 2020, 3, 32. [CrossRef]
- Saddique, Z.; Faheem, M.; Habib, A.; UlHasan, I.; Mujahid, A.; Afzal, A. Electrochemical Creatinine (Bio)Sensors for Point-of-Care Diagnosis of Renal Malfunction and Chronic Kidney Disorders. *Diagnostics* 2023, 13, 1737. [CrossRef]
- Lin, B.; Yu, Y.; Cao, Y.; Guo, M.; Zhu, D.; Dai, J.; Zheng, M. Point-of-care testing for streptomycin based on aptamer recognizing and digital image colorimetry by smartphone. *Biosens. Bioelectron.* 2018, 100, 482–489. [CrossRef]
- 40. Guo, L.; Zhao, D.M.; Chen, S.; Yu, Y.L.; Wang, J.H. Smartphone-Integrated Photoacoustic Analytical Device for Point-of-Care Testing of Food Contaminant Azodicarbonamide. *Anal. Chem.* **2022**, *94*, 14004–14011. [CrossRef]
- Yadav, P.; Yadav, L.; Laddha, H.; Agarwal, M.; Gupta, R. Upsurgence of smartphone as an economical, portable, and consumerfriendly analytical device/interface platform for digital sensing of hazardous environmental ions. *Trends Environ. Anal. Chem.* 2022, *36*, e00177. [CrossRef]
- 42. Lyu, Z.; Ding, S.; Tieu, P.; Fang, L.; Li, X.; Li, T.; Pan, X.; Engelhard, M.H.; Ruan, X.; Du, D.; et al. Single-Atomic Site Catalyst Enhanced Lateral Flow Immunoassay for Point-of-Care Detection of Herbicide. *Research* 2022, 2022, 9823290. [CrossRef] [PubMed]
- 43. Tao, D.; Xiao, X.; Lan, X.; Xu, B.; Wang, Y.; Khazalwa, E.M.; Pan, W.; Ruan, J.; Jiang, Y.; Liu, X.; et al. An Inexpensive CRISPR-Based Point-of-Care Test for the Identification of Meat Species and Meat Products. *Genes* **2022**, *13*, 912. [CrossRef] [PubMed]
- Suan Ng, S.; Ling Lee, H.; Bothi Raja, P.; Doong, R. an Recent Advances in Nanomaterial-based Optical Biosensors as Potential Point-of-Care Testing (PoCT) Probes in Carcinoembryonic Antigen Detection. *Chem. An Asian J.* 2022, 17, e202200287. [CrossRef] [PubMed]
- Arabzadeh, N.; Khosravi, A.; Mohammadi, A.; Mahmoodi, N.M.; Khorasani, M. Synthesis, characterization, and application of nano-molecularly imprinted polymer for fast solid-phase extraction of tartrazine from water environment. *Desalin. Water Treat.* 2015, 54, 2452–2460. [CrossRef]
- 46. Lamaoui, A.; Karrat, A.; Amine, A. Molecularly imprinted polymer integrated into paper-based analytical device for smartphonebased detection: Application for sulfamethoxazole. *Sens. Actuators B Chem.* **2022**, *368*, 132122. [CrossRef]
- Sergeyeva, T.; Yarynka, D.; Piletska, E.; Linnik, R.; Zaporozhets, O.; Brovko, O.; Piletsky, S.; El, A. Development of a smartphonebased biomimetic sensor for aflatoxin B1 detection using molecularly imprinted polymer membranes. *Talanta* 2019, 201, 204–210. [CrossRef]
- Lowdon, J.W.; Diliën, H.; van Grinsven, B.; Eersels, K.; Cleij, T.J. Colorimetric Sensing of Amoxicillin Facilitated by Molecularly Imprinted Polymers. *Polymer* 2021, 13, 2221. [CrossRef]
- 49. Kaymaz, C.; Güner, H.; Akbulut, M.; Güven, O. A smartphone-based colorimetric PET sensor platform with molecular recognition via thermally initiated RAFT-mediated graft copolymerization. *Sens. Actuators B Chem.* **2019**, 296, 126653. [CrossRef]
- Elfadil, D.; Della Pelle, F.; Compagnone, D.; Amine, A. Green Synthesis of Molecularly Imprinted Polymers for Dispersive Magnetic Solid-Phase Extraction of Erythrosine B Associated with Smartphone Detection in Food Samples. *Materials* 2022, 15, 7653. [CrossRef]
- Al-Degs, Y.S.; Abu-Surrah, A.S.; Ibrahim, K.A. Preparation of highly selective solid-phase extractants for Cibacron reactive dyes using molecularly imprinted polymers. *Anal. Bioanal. Chem.* 2009, 393, 1055–1062. [CrossRef]
- 52. García, A.; Erenas, M.M.; Marinetto, E.D.; Abad, C.A.; De Orbe-Paya, I.; Palma, A.J.; Capitán-Vallvey, L.F. Mobile phone platform as portable chemical analyzer. *Sens. Actuators B Chem.* **2011**, *156*, 350–359. [CrossRef]
- 53. John, C.R.; Russ, J.C. Introduction to Image Processing and Analysis; CRC Press: Boca Raton, FL, USA, 2017.
- 54. Ostra, M.; Ubide, C.; Vidal, M.; Zuriarrain, J. Detection limit estimator for multivariate calibration by an extension of the IUPAC recommendations for univariate methods. *Analyst* **2008**, *133*, 532–539. [CrossRef]

Disclaimer/Publisher's Note: The statements, opinions and data contained in all publications are solely those of the individual author(s) and contributor(s) and not of MDPI and/or the editor(s). MDPI and/or the editor(s) disclaim responsibility for any injury to people or property resulting from any ideas, methods, instructions or products referred to in the content.