

Research Article

Development of Effervescent Tablets Containing Elderberry Vine Tea Compound and Evaluation of Their Antioxidant Activity

Wanhao Sun 

College of Food Science & Nutritional Engineering, China Agriculture University, Beijing, China

Correspondence should be addressed to Wanhao Sun; 15558141912@163.com

Received 28 August 2023; Revised 20 November 2023; Accepted 22 November 2023; Published 7 December 2023

Academic Editor: Alam Zeb

Copyright © 2023 Wanhao Sun. This is an open access article distributed under the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Elderberry and vine tea, renowned for their wealth of bioactive constituents and potent antioxidant prowess, hold remarkable promise within a wide array of culinary contexts. This study presents a pioneering advancement in the realm of compound effervescent tablet formulation. Employing the response surface design optimization, this work develops a formulation comprising a total disintegrant (acid-to-base ratio 1.3:1) at 50%, elderberry extracts at 31.7%, ultra-micro vine tea powder at 8.3%, stevioside at 3.4%, PEG-6000 at 2.7%, mannitol at 2.9%, and PVP at 1%. Under these carefully optimized conditions, the resultant tablet exhibits remarkable attributes including an average disintegration time of 184.50 ± 4.65 s, foaming quantity of 30.64 ± 2.10 mg/g, and tablet hardness measuring 54.59 ± 3.41 N, with a weight uniformity well within the $\pm 5\%$. Regarding the antioxidant efficacy, the total flavonoid content attains 76.79 ± 3.76 mg RE/g, and phenolic content reaches 94.84 ± 4.95 mg GAE/g. Moreover, the EC_{50} values for scavenging 1,1-diphenyl-2-picrylhydrazyl (DPPH), hydroxyl (OH), and 2, 2'-azino-bis(3-ethylbenzothiazoline-6-sulfonic acid)(ABTS) radicals are determined at 0.4667 mg/mL, 0.9772 mg/mL, and 0.5014 mg/mL, respectively. This investigation underscores the performance of the effervescent tablet in satisfying stringent physicochemical standards, while also showcasing outstanding antioxidant potency, paving the way for the advancement of antioxidant-functional foods and sustainable exploitation of natural resources.

1. Introduction

The evolving dietary habits and sedentary lifestyles of contemporary populations have led to an upsurge in free radicals within the body, attributed to factors such as high-fat, high-sugar fast food, and limited physical activity [1]. The consequential accumulation of free radical-induced damage has been linked to a spectrum of organ-specific ailments, such as Alzheimer's disease [2], atherosclerosis, diabetes, and radiation-induced afflictions [3]. In this context, antioxidants have emerged as potent defenders against oxidative stress, curbing the impact of free radicals on cellular and tissue integrity [4]. Natural sources of antioxidants include vitamin C, carotenoids, polyphenols, and flavonoids [5].

Elderberries, distinguished for their high flavonoid content, particularly anthocyanins, stand out as a prime repository of these bioactive compounds among various berry species [6]. Studies have shown that ripe elderberries contain 47% flavanol glycosides, 34% anthocyanins, 14%

flavanols, and 3% hydroxycinnamic acids [7]. Scientific investigations have also affirmed the efficacy of elderberries in promoting mitochondrial and cellular revitalization, sustaining cellular equilibrium, and retarding the aging process [8]. This botanical asset, replete with antioxidant potency, emerges as an ideal candidate for innovative product development. Nonetheless, the utilization of elderberries remains largely confined to juices, fruit teas, and jams, which struggle with preservation challenges and susceptibility to significant anthocyanin losses during processing [9].

Vine tea, derived from the dried tender leaves of *Ampelopsis grossedentata* [10], exhibits significant antioxidant potential due to its rich content of 36 bioactive compounds, including gallic acid, butyric acid, ellagic acid, and myricetin, as demonstrated by the Folin-Ciocalteu assay [11]. However, vine tea products still remains in a preliminary processing stage due to geographical constraints and the inherent astringent taste [12]. Notably, conventional brewing methods only facilitate the infusion of a fraction of

bioactive components into the tea broth, leading to the underutilization of resources. To address these limitations, ultra-micropulverization technology emerges as a promising avenue. This technology can effectively enhance the content of water-soluble carbohydrates and the extraction of terpene synthase (TPS), thereby bolstering the scavenging activity on OH radicals [13]. Although widely adopted in black tea, green tea, Pu'er tea, and Oolong tea processing [14], its application to vine tea is largely unexplored.

Based on the aforementioned information, it is evident that elderberry and vine tea are rich in antioxidants. These properties position elderberry as a promising candidate for treating various inflammatory diseases including Huntington's disease [15], atherosclerosis [16], and UVB-induced skin photoaging [17]. Similarly, vine tea, as a traditional Chinese herbal medicine, is frequently utilized to alleviate arthritis symptoms [18], liver fibrosis [19], and atherosclerosis [20]. In a strategic endeavor to unlock the untapped potential of vine tea and fully utilize the inherent qualities of elderberry, this study introduces a pioneering approach. It deploys ultra-micropulverization technology for the treatment of vine tea, seamlessly integrated with the use of effervescent tablets. That is because elderberry extract and ultra-micro vine tea powder are highly dehydrated and hygroscopic, while the anthocyanins present in the extracts are sensitive to environmental conditions [21]. As a result, it imposes high costs of storage and transportation. To overcome this challenge, the manufacturing of tablets is considered a distinctive technique [22]. Tableting stabilizes the physical and chemical properties of the product, extending its shelf life. Furthermore, tablet production offers advantages such as improved appearance and increased consumer acceptance [23].

Response surface designs are utilized to determine the best combination of inputs that yield the most favourable outcomes. This is achieved by minimizing the average prediction variance across the design spaces. Such designs are valuable tools for optimizing products and processes as they involve iteratively adjusting one or more inputs and analysing the impact of these adjustments on the output(s) [24]. Using this method, this study is aimed at the following:

- (1) Enhancing nutrient retention in vine tea through ultra-micropulverization
- (2) Alleviating the astringency of vine tea by harnessing the natural sweet and sour fruity flavour of elderberry
- (3) Developing an experimentally validated effervescent tablet formula that possesses strong antioxidant properties and is made of natural, healthy ingredients

Through the assessment of disintegration kinetics, foaming characteristics, tablet hardness, weight consistency, and free radical scavenging activities, this investigation comprehensively unravels the potential of this novel approach. This study not only elevates the nutritional profiles of elderberry and unlocks the untapped potential of vine tea products through meticulous processing and flavour enhancement but also lays the essential theoretical foundation to meet

the evolving demands of modern society for antioxidant-rich functional foods.

2. Materials and Methods

2.1. Materials and Reagents. Vine tea: Zhangjiajie Baxingteng Tea Planting Farmers' Professional Cooperative; elderberry extract: Xi'an Michel Biotechnology Co., Ltd.; mannitol, citric acid, sodium bicarbonate: Henan Gaobao Industrial Co., Ltd.; PEG-6000: Shandong Ruisheng Excipients Co., Ltd.; polyvinyl pyrrolidone: Zhejiang Huiquan Biotechnology Co., Ltd.; anhydrous ethanol: Jinan Qiwei Chemical Co., Ltd.; total flavonoid content assay kit, total phenol content assay kit, DPPH radical scavenging activity assay kit, hydroxyl radical scavenging activity assay kit, ABTS radical scavenging activity assay kit: Beijing Solarbio Science & Technology Co., Ltd. All the materials used are food grade.

2.2. Experimental Methods

2.2.1. Ultra-Micropulverization of Vine Tea. The vine tea was introduced into the oscillating pulveriser (MF-4A, Wenling Mingda Medicine Machinery Equipment Co., Ltd., China) and crushed for 5 minutes, and then sieved through a 100-mesh sieve to yield coarse-grinding powder. This preliminary powder was further ground using an ultra-micropulveriser (MFC-35B, Wenling Mingda Medicine Machinery Equipment Co., Ltd., China) in a single round of grinding for 10 minutes and then sieved through a 200-mesh sieve to obtain the ultra-micropowder [25].

2.2.2. Tablet Preparation. The acid-base separation semidry granule pressing method was employed to prepare the effervescent tablets [26, 27], as shown in Figure 1. Stevioside, citric acid, and elderberry extract powder were finely ground and sieved through a 120-mesh sieve and then blended with ultra-micro vine tea powder. After adding a 5% polyvinyl pyrrolidone (PVP) solution, the mixture was dried at 50°C. The resulting mixture was sieved again to produce acid granules. Alkali granules were prepared using a similar pathway. Both granules were then combined, and PEG-6000 lubricant was added. The mixture was pressed under 10 Mpa for 30 seconds using a tablet press (FW-5A, Tianjin Tianguang New Optical Instrument Technology Co., Ltd., China) to produce compound effervescent tablets with consistent composition.

2.2.3. Sensory Evaluation. After undergoing rigorous professional training, ten expert assessors, aged between 19 and 50 years, were selected to conduct sensory evaluations. The evaluation employed a double-masked methodology. The assessors evaluated preferences based on four dimensions. The detailed scoring is provided in Table 1.

To address varying levels of importance attributed to individual factors in the product evaluation, a weight distribution was established based on previous research [28]. Five experienced assessors independently assigned weights to a fuzzy subset $X = \{X_1, X_2, X_3, X_4\}$ of factors related to effervescent tablets, reflecting their perceived significance. Each X_i value, ranging between 0 (not necessary) and 1 (very important), was rounded to one decimal place. Normalizing

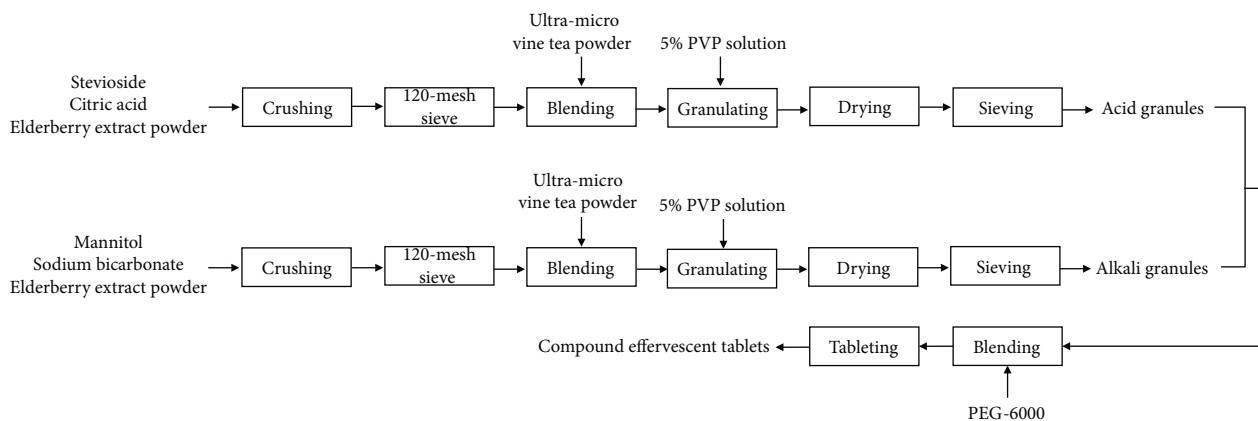


FIGURE 1: Acid-base separation semi-dry granule pressing method.

TABLE 1: Sensory evaluation criteria.

Evaluation items	Evaluation criteria	Score
Appearance (X_1)	Tablets intact and surface flat	8-10
	Tablets complete and slight flaking	4-7
	Tablets incomplete/heavy flaking	0-3
Taste (X_2)	Sweet and sour and no astringency	8-10
	Sour/sweet/slightly astringent	4-7
	Too acidic/too sweet/bitter taste	0-3
Liquor colour (X_3)	The clear solution and no precipitation	8-10
	Little cloudy solution/a bit of precipitation	4-7
	Turbid solution/much precipitation	0-3
Aroma (X_4)	Strong fruity and tea-like aroma	8-10
	Faintly fruity and tea-like aroma	4-7
	No fruit or tea aroma	0-3

and averaging this data yielded the weight set $A = \{A_1, A_2, A_3, A_4\}$. The total sensory score was then computed using Equation (1).

$$\text{Sensory Score} = X_1 \times A_1 + X_2 \times A_2 + X_3 \times A_3 + X_4 \times A_4. \quad (1)$$

2.2.4. Single-Factor Experiments for Sensory Evaluation. This study employed single-factor experiments and focused on parameters intricately contributing to the overall perception of effervescent tablets. The levels of immobilization were determined in preliminary experiments: a disintegrant content of 50% at a 1.3:1 acid-to-base ratio, complemented by a 40% combination of elderberry extracts in a 4:1 ratio with ultra-micro vine tea powder, 2.5% stevioside, 4.5% mannitol, 2% PEG-6000, and 1% PVP.

2.2.5. Response Surface Design. Building upon the outcomes of the single-factor experiments, a four-variable, three-level Box-Behnken design was employed. The utilization of Design-Expert 8.0 facilitates multiple regression analyses and the subsequent optimization [29].

2.2.6. Determination of Physicochemical Properties

(1) **Disintegration Time.** Following the guidelines of the Chinese Pharmacopoeia 2020 [30], an effervescent tablet was placed in a 250 mL beaker with 200 mL of distilled water at a temperature of $20 \pm 5^\circ\text{C}$, and the time required for complete disintegration was recorded.

(2) **Foaming Quantity.** For accurate measurements, the effervescent tablets were first ground into powder and then tested for foaming quantity [31]. The weight of effervescent tablet powder before being placed into an aqueous beaker (M_0), the total mass before foaming (M_1), and the final weight of the residue after foaming (M_2) were determined using an electronic balance (FA2104N, Shanghai Jingqi Instrument Co., Ltd., China). The foaming quantity was computed using Equation (2), which represented the amount of CO_2 gas generated in milligrams per gram of effervescent tablet.

$$\text{Foaming Quantity (mg/g)} = \frac{M_1 - M_2}{M_0} \times 1000. \quad (2)$$

(3) *Average Weight and Weight Variation.* Twenty effervescent tablets were selected to compute the average weight. The actual weight of each tablet was compared to the average value [30].

(4) *Hardness.* The hardness was evaluated using a texture analyser (TA-XT Plus, SMS Corporation, UK) with a 2 mm cylindrical probe, a force sensing element ranging up to 100 N. The probe extrusion distance was set at 2 mm, and the detection speed was 60 mm/min. The starting force was 0.4 N, and the maximum force value was measured [32].

(5) *Moisture Content.* The moisture content of the effervescent tablets was determined using the direct drying method [30]. A weighing bottle was placed in a vacuum-drying oven (DZF-6024, Shaoxing Shangli Instrument Co., Ltd., China) and heated until a consistent weight was attained. Subsequently, a precise quantity of the effervescent tablet (M_3) was weighed and added to the same bottle. The bottle was then capped, and the total weight (M_4) was recorded. The bottle and its contents were then placed back into the drying oven until a stable weight was reached (M_5). The moisture content was calculated as Equation (3).

$$\text{Moisture Content (g/100g)} = \frac{M_4 - M_5}{M_3} \times 100. \quad (3)$$

(6) *Colour.* Following the recommendations of Commission Internationale d'Eclairage (CIE), the colour was represented by L^* (brightness/darkness), a^* (redness/greenness), and b^* (yellowness/blueness) values, utilizing a colour meter (ZE6000, Nippon Denshoku, Japan). Additionally, chroma (C), hue angle (h), and yellowness index (YI) were calculated through the following equations [33].

$$\begin{aligned} C &= \sqrt{a^{*2} + b^{*2}}, \\ h &= \tan^{-1} \left(\frac{b^*}{a^*} \right), \\ \text{YI} &= 142.86 \frac{b^*}{L^*}. \end{aligned} \quad (4)$$

2.2.7. Determination of Antioxidant Content

(1) *Total Flavonoid Content (TFC).* The TFC was determined using a colorimetry method involving NaNO_2 - $\text{Al}(\text{NO}_3)_3$ - NaOH [34]. The sample was extracted in 70% alcohol for 10 minutes using an ultrasonic cleaning machine (PS-10 AD, Shenzhen Jiekang Cleaning Electric Appliance Co., Ltd., China). Subsequently, the reaction mixture was diluted to 25 mL with distilled water, and its absorbance was measured at 510 nm using a spectrophotometer (UV-5100, Shanghai Jinghong Experimental Equipment Co., Ltd., China). The TFC was quantified as rutin equivalents (RE) per gram of dry extract.

(2) *Total Polyphenol Content (TPC).* The TPC was determined using the Folin-Ciocalteu spectrophotometry method

with gallic acid as the external standard [35]. The sample was extracted in 70% alcohol for 10 minutes and neutralized with Na_2CO_3 (0.5 mL, 20%, w/v). Subsequently, the reaction mixture was diluted to 50 mL using distilled water, and its absorbance was measured at 760 nm. The TPC was quantified as gallic acid equivalents (GAE) per gram of dry extract.

2.2.8. Determination of Antioxidant Capacity

(1) *DPPH Radical Scavenging Activity.* Incorporating adaptations from the study conducted by Wu et al. [36], after being dissolved in 70% ethanol, the samples underwent 20 minutes of ultrasonication, and different sample concentrations were prepared: 0.1 mg/mL, 0.3 mg/mL, 0.5 mg/mL, 0.7 mg/mL, 1.0 mg/mL, 3.0 mg/mL, 5.0 mg/mL, 7.0 mg/mL, and 9.0 mg/mL. The experimental setup utilized a 96-well plate, with each initially receiving 100 μL of 0.1 mM DPPH chromogenic reagent, followed by the addition of 100 μL of the specific sample solution. Subsequently, the absorbance was measured at 517 nm with a microplate reader (ThermoFC, Shanghai Yilang Electromechanical Equipment Co., Ltd., China) after 30 min incubation at room temperature. DPPH radical scavenging activity was assessed using the following equation:

$$\text{DPPH (\%inhibition)} = \left(1 - \frac{A_i}{A_0} \right) \times 100\%, \quad (5)$$

where A_i and A_0 represent the absorbance of the sample and blank, respectively.

(2) *Hydroxyl Radical Scavenging Activity.* Following the methodological guidelines stipulated by the kit, the samples are dissolved in 70% ethanol and underwent 20 minutes of ultrasonication. Then, the same concentration ladder as the DPPH radical scavenging experiments was prepared. Subsequently, the reaction solution was transferred to a 96-well plate, incubated for 20 min at room temperature, and absorbance at 550 nm was measured using a microplate reader equipped with a 1 cm optical aperture. Hydroxyl radical scavenging activity was assessed using the following equation:

$$\text{Hydroxyl (\%inhibition)} = \left(\frac{A_c - A_i}{A_c - A_0} \right) \times 100\%, \quad (6)$$

where A_c , A_i , and A_0 represent the absorbance of the control, sample, and blank, respectively.

(3) *ABTS Radical Scavenging Activity.* Incorporating adaptations from the study conducted by Thaipong et al. [37], the ABTS working solution was prepared as follows: 5 mL of a 7.4 mM ABTS radical solution was mixed with 88 μL of a 2.6 mM potassium persulfate solution. The mixture was allowed to stand at room temperature in the dark for 12 to 16 hours. Prior to measurement, the working solution was diluted with 80% ethanol to achieve an absorbance range of 0.7 ± 0.05 at 734 nm. Then, 20 μL of sample solutions with the same concentration ladder as the DPPH radical scavenging experiments were combined with 200 μL of the diluted

ABTS solution. After incubating for 6 minutes, the absorbance at 734 nm was measured spectrophotometrically. The ABTS radical scavenging activity was assessed using the following equation:

$$\text{ABTS}(\% \text{inhibition}) = \left(1 - \frac{A_i}{A_0}\right) \times 100\%, \quad (7)$$

where A_i and A_0 represent the absorbance of the sample and blank, respectively.

2.2.9. Statistical Analysis. Data are presented as the mean \pm SD. The differences among the groups were compared by one-way analysis of variance (ANOVA). Correlations were calculated by the Pearson linear regression analysis. All statistics were analysed by Design-Expert (V8.0.6.1), SPSS (27.0), and GraphPad Prism (9.5.1). A P value < 0.05 indicates statistical significance. Unless otherwise noted, all data were generated from three independent experiments.

3. Results and Discussion

3.1. Weight Distribution for Sensory Evaluation. To comprehensively understand the preferences of the assessors and their collective impact on the overall sensory experience, the weight distribution of key sensory factors in the effervescent tablet assessment is presented in Table 2.

Evidently, taste (X_2) and aroma (X_4) emerge as central considerations, both bearing a weight of 0.3. This emphasis resonates with earlier findings by Liao et al. [38] and Chen [39], where the forced-decision method corroborates the significance of these attributes. Appearance (X_1) and liquor colour (X_3) carry equal weights of 0.2 each. Their acceptability is more intricately linked to the operational parameters. Therefore, the total sensory score can be calculated as equation (8), which better encapsulates the evaluative priorities.

$$\text{Sensory Score} = \text{Taste} \times 30\% + \text{Aroma} \times 30\% + \text{Appearance} \times 20\% + \text{Liquor Colour} \times 20\%. \quad (8)$$

3.2. Investigation of Single-Factor Experiments. The results of single-factor experiments are shown in Figure 2. Figure 2(a) reveals a distinct sensory score trend concerning the elderberry extract-to-ultra-micro vine tea powder ratio. The score exhibits a biphasic pattern, characterized by an initial increase followed by a decline, reaching its pinnacle at a ratio of 4:1. This optimal ratio engenders a sensorial experience dominated by a pronounced fruity aroma, complemented by the unique flavour of vine tea. The visual aspect also contributes to the sensory appeal, as the colour showcases transparency and a pleasing peach hue. At lower ratios (below 3:1), the presence of insoluble material becomes noticeable. This can be attributed to the gradual increase of insoluble substances in vine tea, such as cellulose and lignin, which leads to the formation of a natural precipitate [40]. Additionally, lower ratios accentuate the bitter and astringent taste of vine tea, as supported by the study conducted by

Wang et al., which attributes this taste to the terpenoid content in vine tea, particularly limonoids [41, 42]. At higher ratios (above 5:1), the solution exudes a robust fruity flavour, along with an appealing rosy red hue, but the tea flavour becomes less pronounced. Therefore, the response surface design selects the ratios of 3:1, 4:1, and 5:1 as the parameter range.

Compared to other nonnutritive sweeteners such as aspartame and acesulfame potassium, steviolides stand out as noncaloric sucrose substitutes and noncariogenic sweeteners, with fewer adverse effects on the neurological system [43]. The influence of total steviolide content on sensory scores, as depicted in Figure 2(b), follows a nuanced trajectory: a gradual ascent succeeded by a notable decline, reaching its pinnacle at 3%. As the quantity augments, the sweetness of the solution gradually intensifies, attaining a balance of moderate sweetness and tanginess. However, surpassing the 3.5% threshold leads to the emergence of a melange of bitterness and metallic attributes. This observation aligns with the findings of Espinoza et al. [44], and the potential cytotoxicity also sets a limit to its extensive addition [45]. Considering both the gustatory experience and safety concerns, this work focuses on a parameter range of 2.5%-3.5% for the addition of steviolide, which aligns with the range established by Skąpska et al. in the context of functional fruit-herbal beverages [46].

In comparison to sodium dodecyl sulfate and magnesium stearate, PEG-6000 exhibits the advantage of being free from metal ions, thus effectively circumventing the catalysis of anthocyanin oxidation [47]. The impact of PEG-6000 addition on the sensory score of effervescent tablets, illustrated in Figure 2(c), unfolds in a two-phased manner, with an initial ascent followed by a subsequent descent, culminating at its zenith at 3%. Within this context, a PEG-6000 addition of less than 2% yields a rough tablet surface prone to detachment. Conversely, a concentration surpassing 5% poses the risk of punch sticking [48]. Consequently, a judicious parameter range of 2%-4% has been delineated.

The quantity of disintegrant holds sway over both taste and disintegration time at a predefined ratio. Figure 2(d) underscores this ascending-then-descending pattern, with the zenith achieved at 50%. Given the fixed acid-to-base ratio of 1.3:1 in disintegrants, the solution's acidity proportionally escalates with an augmented disintegrant content. Notably, a disintegrant content less than 45% yields a bland disintegrated solution, while an excess of 55% elicits an irksome acidity. This typically necessitates the introduction of sugar to attain a balanced sour-sweet profile [49]. The amount of disintegrant also profoundly influences the disintegration time: a higher quantity of disintegrant correlates with progressively shorter disintegration times. Ultimately, this trend culminates in a disintegration time of less than 4 minutes with a disintegrant content exceeding 45%, which remains well within the acceptable range for commercially available edible effervescent tablets [50]. A comprehensive synthesis of sensory evaluation and disintegration time considerations yields a parameter range of 45%-55% for optimal response surface design.

TABLE 2: Weight distribution for sensory evaluation.

Professional assessors	Degree of affiliation				Normalization			
	X_1	X_2	X_3	X_4	X_1	X_2	X_3	X_4
1	0.3	0.6	0.2	0.2	0.2	0.5	0.2	0.2
2	0.2	0.5	0.2	0.5	0.1	0.4	0.2	0.4
3	0.2	0.3	0.4	0.5	0.1	0.2	0.3	0.4
4	0.3	0.6	0.2	0.7	0.2	0.3	0.1	0.4
5	0.3	0.4	0.2	0.4	0.2	0.3	0.2	0.3
Average					0.2	0.3	0.2	0.3

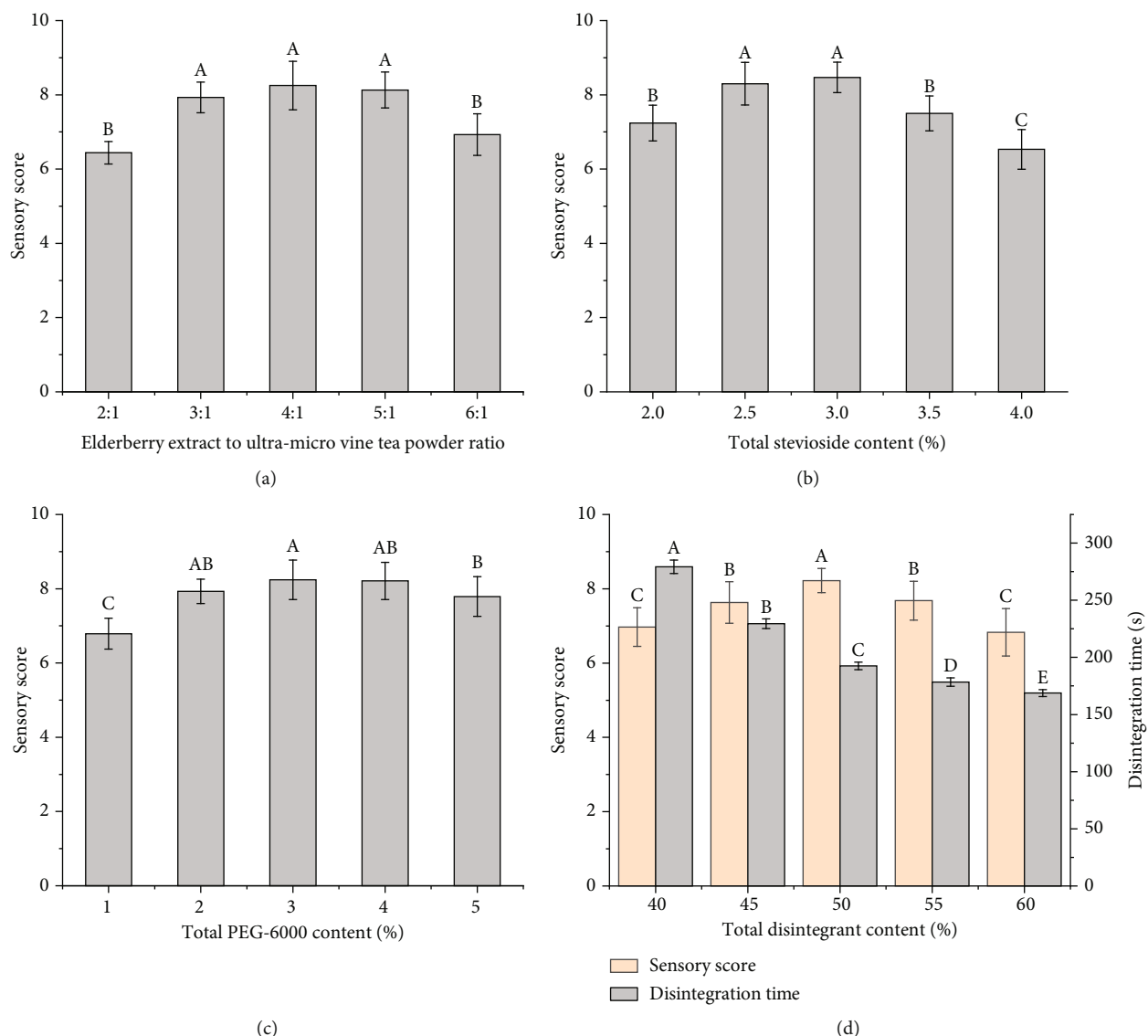


FIGURE 2: Optimal parameter ranges via single-factor experiments. Data are represented mean \pm SD ($n = 10$). Bars labelled with distinct letters (A, B, C, D, and E) indicate statistical significance at $P < 0.05$. (a) Elderberry extract-to-ultra-micro vine tea powder ratio: influence on sensory scores. (b) Total stevioside content: impact on sensory scores. (c) Total PEG-6000 content: effect on sensory scores. (d) Total disintegrant content: relationship with sensory scores.

3.3. Response Surface Optimization and Analysis

3.3.1. Response Surface Design and Analysis of Variance. Guided by the results gained from the single-factor experiments, this study conducted the response surface design according to 2.2.5. The sensory scores (Y) serve as the response values, and the factors and levels are shown in Table 3. Detailed results of these multifaceted experiments are presented in Tables 4 and 5.

Through regression fitting and variance analysis, a quadratic multinomial regression equation that describes the relationship between sensory scores (Y) and the four independent variables is derived in the following equation:

$$Y = 8.39 - 0.23A + 0.46B - 0.13C - 0.037D + 0.21AB - 0.02AC - 0.15AD + 0.022BC + 0.005BD + 0.06CD - 0.013A^2 - 0.25B^2 - 0.21C^2 - 0.65D^2. \quad (9)$$

As shown in Table 5, the regression equation model boasts a F value of 144.98, accompanied by a low significance level ($P < 0.0001$). This outcome substantiates the model's profound significance and its congruence with the established equation. In contrast, the lack of fit test reveals an F value of 0.55, coupled with a P value of 0.7861, surpassing the conventional significance threshold of 0.05. Thus, this particular term stands as inconsequential, signifying a marginal model error. The metrics of $R^2 = 0.9941$ and $\text{Adj } R^2 = 0.9873$ illustrate that this model elucidates 98.73% of the variance observed in sensory scores. The coefficient of variation (CV) is a mere 0.7%, undercutting the accepted 5% threshold, a testament to the model's robust adaptability.

The orchestration of influence among the quartet of factors shaping the overarching scores of the effervescent tablets emerges with utmost clarity from the discerning lens of the F and P values in the ANOVA. This hierarchical arrangement unfolds as follows: reigning supreme is the total stevioside content (B), positioned above the ratio of elderberry extract-to-ultra-micro vine tea powder (A), succeeded by the total PEG-6000 content (C), and culminating with the total disintegrant content (D). Systematically, the facets that wield a resounding impact, adorned with a statistical significance of ($P < 0.01$), reveal themselves as the primary actors, namely, terms A , B , and C , alongside the secondary personas B^2 , C^2 , D^2 , and the intriguing interactions of AB and AD . Marching to a slightly muted tune but nevertheless embracing significance ($P < 0.05$), the primary role is attributed to term D . However, adopting a cloak of insignificance ($P > 0.05$), the secondary term A^2 , and the cross terms AC , BC , BD , and CD fade into the backdrop. A refined optimization model (Equation (10)) emerges.

$$Y = 8.39 - 0.23A + 0.46B - 0.13C - 0.037D + 0.21AB - 0.15AD - 0.25B^2 - 0.21C^2 - 0.65D^2. \quad (10)$$

3.3.2. Interactive Analysis. In order to gain insights into the combined effects of different factors on the sensory score, this study performed an interactive analysis. The response surface and contour maps are presented in Figure 3, offering a visual representation of their effects on the sensory score.

As illustrated in Figure 3(a), the 3D surface plot representing the interaction between the elderberry extract-to-ultra-micro vine tea powder ratio and total stevioside content exhibits a relatively steep slope, accompanied by a large ovality in the contour map. These characteristics indicate a significant impact of this interaction on the sensory score ($P < 0.05$). According to the results of the single-factor experiment, as the elderberry extract-to-ultra-micro vine tea powder ratio decreased, the main constraint on sensory scores was the bitterness of the vine tea, but the refreshing sweetness of the stevioside can mask this drawback well. As mentioned in previous studies, dry stevia leaves have been used to sweeten traditional bitter drinks such as mate tea [51]. On the other hand, the berry sweetness of the elderberry itself and the herbal sweetness of the stevioside are somewhat incompatible when the above ratios are too high, even leading to slightly astringent aftertaste. Thus, within certain limits, the interaction of these two factors (AB) should not be omitted.

Similarly, in Figure 3(b), as the elderberry extract-to-ultra-micro vine tea powder ratio and total disintegrant content increase, the sensory score initially rises and then declines. The evident curve radian of the response surface and the large ovality of the contour map further emphasize the considerable influence of this interaction (AD) on the sensory score ($P < 0.05$). It is supposed that at the lower the elderberry extract-to-ultra-micro vine tea powder ratio, citric acid can act as a bitter taste modifier similar to stevioside. This hypothesis is confirmed by Sotoyama et al., who found that citric acid can suppress the bitter taste of olopatadine hydrochloride orally disintegrating tablets [52]. Similarly, in higher ratio, a sour-sweet solution presents a balance of both sour and sweet flavours. Many drinks on the market such as fruit vinegars are made using this principle of neutralization [53].

Conversely, the interactions between the other four groups of factors (AC , BC , BD , and CD) show relatively weaker effects on the sensory score, as evident from Figures 3(c)–3(f). These findings are consistent with the ANOVA results, reinforcing the notion that these interactions may have less pronounced impacts on the overall sensory evaluation. Of these, the amount of total PEG-6000 does not have a significant interaction with any of the other variables, which attributes to the fact that at this range of additions, it has little effect on mouthfeel and mainly affects the morphology of the tablets [54].

3.3.3. Validation of Optimum Conditions. Based on the interactive analysis, this study proceeds to validate the optimal conditions, which encompass an elderberry extract-to-ultra-micro vine tea powder ratio of 3.8 : 1, coupled with precise content of stevioside, PEG-6000, and disintegrant at 3.41%, 2.73%, and 49.87%, respectively. This configuration

TABLE 3: Factors and levels of response surface design.

Levels	Factors			
	A Elderberry extract-to-ultra-micro vine tea powder ratio	B Total stevioside content (%)	C Total PEG-6000 content (%)	D Total disintegrant content (%)
-1	3:1	2.5	2	45
0	4:1	3	3	50
1	5:1	3.5	4	55

TABLE 4: Design of response surface tests and results.

Run	A	B	C	D	Response
1	0	0	-1	1	7.53
2	0	0	0	0	8.42
3	-1	0	1	0	8.3
4	1	-1	0	0	7.18
5	0	0	1	-1	7.37
6	0	0	0	0	8.31
7	1	0	0	1	7.38
8	-1	0	0	1	8.11
9	0	-1	0	1	6.98
10	0	-1	-1	0	7.77
11	0	1	0	1	7.86
12	1	1	0	0	8.54
13	1	0	0	-1	7.68
14	0	-1	1	0	7.28
15	0	0	-1	-1	7.74
16	0	1	-1	0	8.55
17	1	0	1	0	7.81
18	0	1	0	-1	7.99
19	0	0	0	0	8.44
20	0	1	1	0	8.28
21	-1	-1	0	0	8.1
22	0	-1	0	-1	7.13
23	-1	1	0	0	8.62
24	1	0	-1	0	8.07
25	-1	0	-1	0	8.48
26	0	0	1	1	7.4
27	-1	0	0	-1	7.79

is projected to yield a sensory score of 8.62. However, considering practicality, a refined adjustment was made; the elderberry extract-to-ultra-micro vine tea powder ratio remains at 3.8:1, while the stevioside, PEG-6000, and disintegrant contents are adjusted to 3.4%, 2.7%, and 50%, respectively. In line with these parameters, a trio of parallel tests is executed, culminating in a sensory score of 8.57 ± 0.11 . Impressively, the relative error value stands at a mere -0.58% in comparison to the prognosticated value. This notable convergence underscores the potential of these optimized conditions. In the forthcoming phase, compre-

hensive assessments encompassing both physical and chemical attributes will illuminate the potency and promise of this refined formulation.

3.4. Quality Inspection of Effervescent Tablets. Following the procedure outlined in Sections 2.2.6 and 2.2.7, a comprehensive evaluation of the physicochemical properties of the effervescent tablets is conducted. Notably, the mean disintegration time is determined to be 184.50 ± 4.65 s, accompanied by a foaming quantity of 30.64 ± 2.10 mg/g, and a weight variation of less than $\pm 5\%$. These three essential

TABLE 5: ANOVA for the fitted quadratic model.

Source	Sum of squares	Df	Mean square	F value	P value (Prob > F)	Significance
Model	6.25	14	0.45	144.98	<0.0001	**
A	0.63	1	0.63	203.32	<0.0001	**
B	2.55	1	2.55	828.19	<0.0001	**
C	0.21	1	0.21	66.75	<0.0001	**
D	0.016	1	0.016	5.24	0.0409	*
AB	0.18	1	0.18	57.33	<0.0001	**
AC	1.60×10^{-3}	1	1.60×10^{-3}	0.52	0.4847	Not significant
AD	0.096	1	0.096	31.23	0.0001	**
BC	2.03×10^{-3}	1	2.03×10^{-3}	0.66	0.433	Not significant
BD	1.00×10^{-4}	1	1.00×10^{-4}	0.032	0.8599	Not significant
CD	0.014	1	0.014	4.68	0.0514	Not significant
A ²	8.90×10^{-4}	1	8.90×10^{-4}	0.29	0.6006	Not significant
B ²	0.34	1	0.34	109.78	<0.0001	**
C ²	0.24	1	0.24	79.5	<0.0001	**
D ²	2.26	1	2.26	733.23	<0.0001	**
Residual	0.037	12	3.08×10^{-3}			
Lack of Fit	0.027	10	2.71×10^{-3}	0.55	0.7861	Not significant
Pure Error	9.80×10^{-3}	2	4.90×10^{-3}			
Cor Total	6.28	26				

Note: * indicates statistically significant, $P < 0.05$; ** indicates highly significant, $P < 0.01$.

parameters demonstrate compliance with the specifications delineated in the 2020 edition of the Chinese Pharmacopoeia [30], indicating that the product has an appropriate disintegration rate, effervescent effect, and consistent tablet quality. The texture analyser data reveal that the tablets exhibit a hardness of $54.59 \pm 3.41 N$. This property is crucial as it ensures the tablets can withstand the various stages of production, packaging, and transportation without being damaged [55].

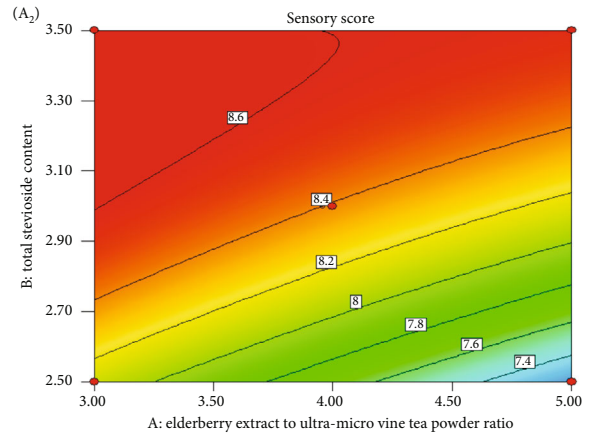
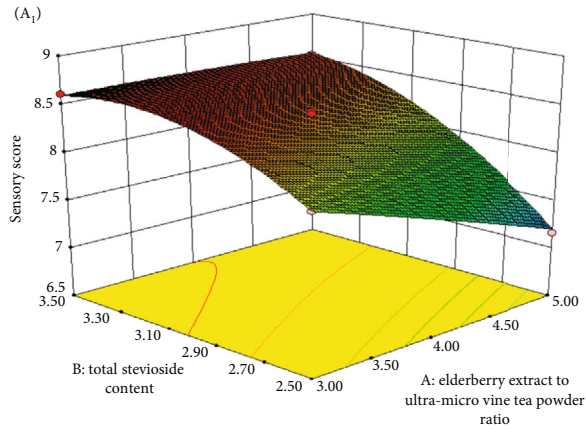
The colour properties provide an objective assessment of the product's appearance. The L^* , a^* , and b^* values indicate a relatively brighter appearance with a red shift on the red-green axis and a reduced yellow shift on the blue-yellow axis. C , h , and YI values suggest a softly saturated colour resembling natural berries, akin to jambolan effervescent tablets developed by Minh [56]. Notably, no signs of artificial colour additions are observed.

Regarding ingredient content, the moisture content is determined to be 3.52 ± 0.13 g/100 g. This value falls below the national standard limit of 5% [30]. The lower moisture content can prolong the shelf life and help mitigate the risk of microbial contamination. Moreover, these effervescent tablets exhibit a remarkable TFC of 76.79 ± 3.76 mg RE/g and TPC of 94.84 ± 4.95 mg GAE/g. Due to the abundance of flavonoids and polyphenols, along with their C6-C3-C6 basic structural units and numerous free phenol hydroxyl groups, the effervescent tablets are expected to exhibit a remarkable activity to scavenge free radicals by providing

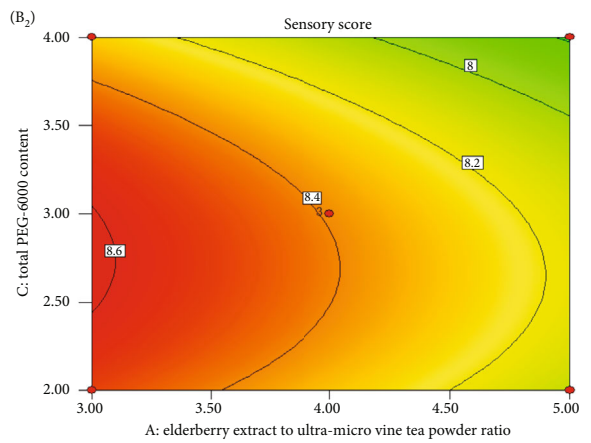
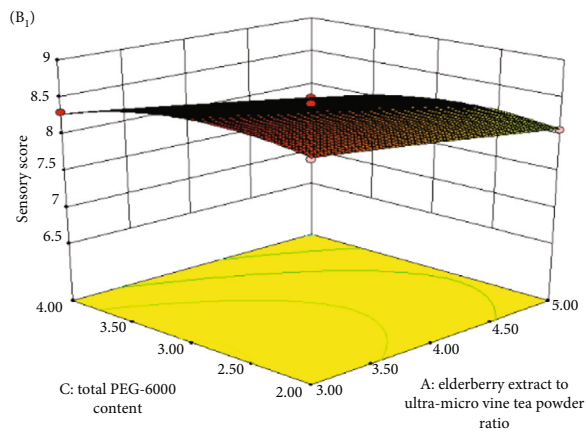
hydrogen protons [57]. The quantification of this activity will be discussed in the following section.

3.5. Detection of Antioxidant Activity. To assess the antioxidant capacity comprehensively, this study performs three established techniques: DPPH, hydroxyl, and ABTS radical scavenging activity experiments [58]. By comparing the antioxidant potential of solutions across diverse concentrations and ascertaining median effect concentration (EC_{50}) values, an insight into the product's efficacy in combating oxidative stress can be gained. The radical scavenging rate curve is shown in Figure 4.

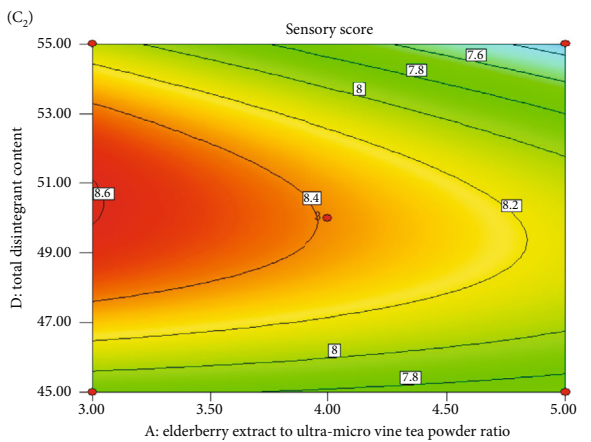
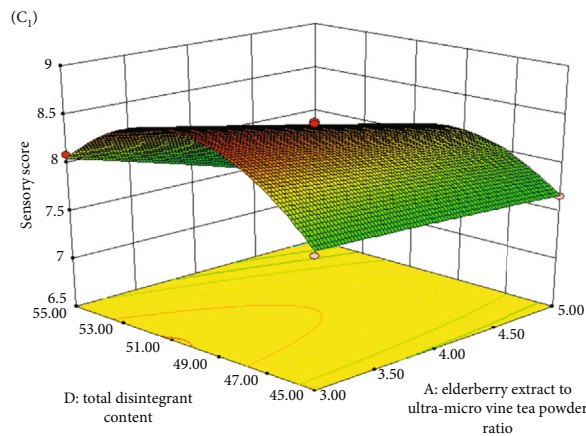
According to the EC_{50} values, the inhibition rates of DPPH, hydroxyl, and ABTS radical are calculated as 0.4667 mg/mL ($R^2 = 0.9819$), 0.9772 mg/mL ($R^2 = 0.9932$), and 0.5014 mg/mL ($R^2 = 0.9856$), respectively. In comparison with other antioxidant solid beverages available in the market, this effervescent tablet demonstrates significantly lower EC_{50} values than the *Aronia melanocarpa* effervescent tablet [59] and fermented *Ginkgo biloba* seeds solid beverage [60], while slightly exceeding the EC_{50} of the *Moringa oleifera* leaf extract effervescent tablet [61]. It is noteworthy that the effervescent tablet's raw material is not a direct anthocyanin extract from elderberries, but rather a concentrated powder designed to preserve taste. Despite this difference, the effervescent tablet still exhibits a well-balanced and satisfactory antioxidant activity, as indicated by its competitive EC_{50} values.



(a)



(b)



(c)

FIGURE 3: Continued.

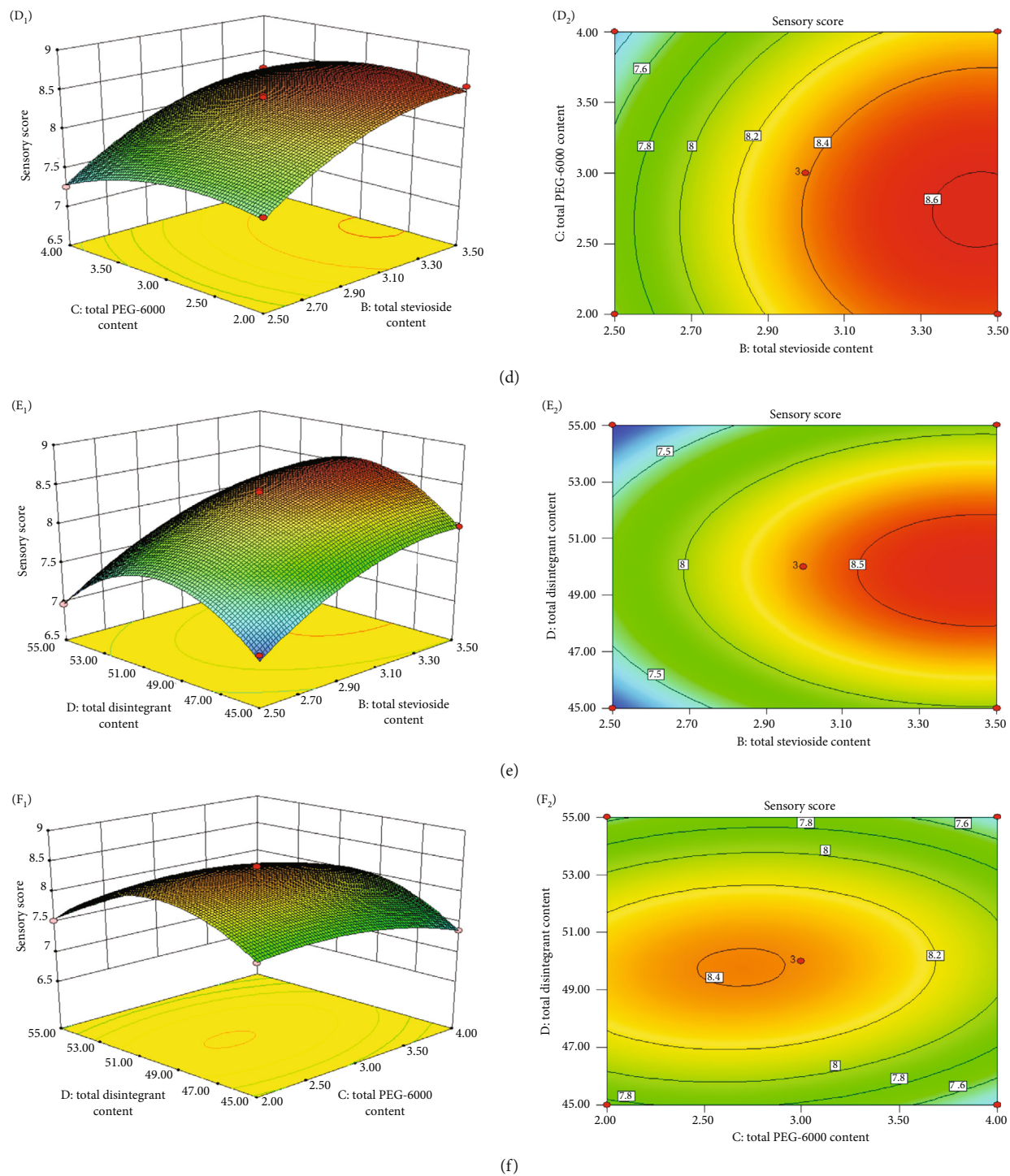


FIGURE 3: Response surface and contour maps for different interactive analyses. The curvature of the response surface plot and the elliptical shape of the contour plot reflect the magnitude of the impact of factor interactions on the sensory score. (a) Interaction between elderberry extract-to-ultra-micro vine tea powder ratio and total stevioside content. (b) Interaction between elderberry extract-to-ultra-micro vine tea powder ratio and total PEG-6000 content. (c) Interaction between elderberry extract-to-ultra-micro vine tea powder ratio and total disintegrant content. (d) Interaction between total stevioside content and total PEG-6000 content. (e) Interaction between total stevioside content and total disintegrant content. (f) Interaction between total PEG-6000 content and total disintegrant content.

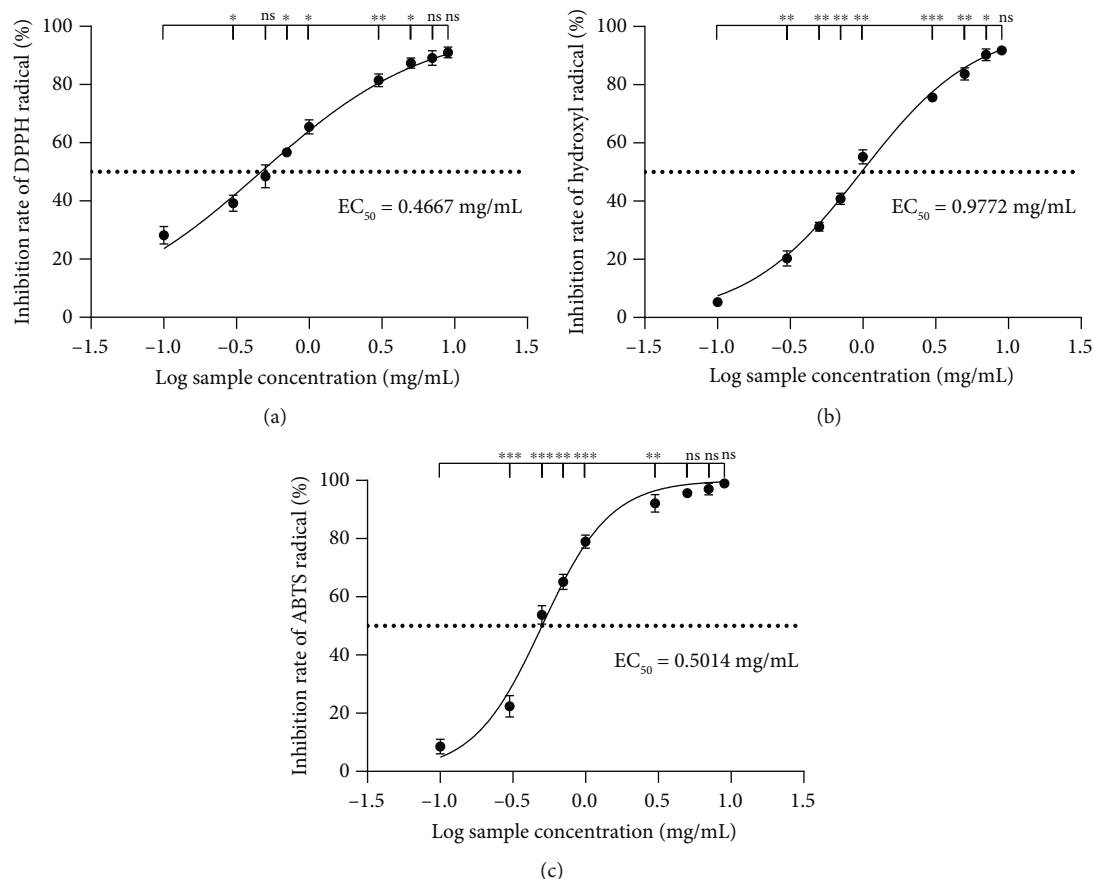


FIGURE 4: Radical scavenging activity of compound effervescent tablets. Data are represented as mean \pm SD ($n = 3$). Asterisks mark statistically significant difference (* $P < 0.05$, ** $P < 0.01$, and *** $P < 0.001$). ns: no statistical significance. (a) DPPH radical inhibition activity at different concentrations. (b) Hydroxyl radical inhibition activity at different concentrations. (c) ABTS radical inhibition activity at different concentrations.

4. Conclusion

In conclusion, this study has successfully developed an effervescent tablet with high levels of antioxidants. The optimized formulation comprises 50% total disintegrant with a balanced acid-to-base ratio of 1.3:1, a blend of elderberry extracts and ultra-micro vine tea powder constituting 40% at a ratio of 3.8:1, 3.4% stevioside, 2.7% PEG-6000, 2.9% mannitol, and 1% PVP. The resulting tablets exhibit an attractive external appearance, clear solution, delightful sweetness, captivating aroma, and commendable physicochemical attributes. Notably, the DPPH, hydroxyl, and ABTS radical scavenging activities are quantified with EC₅₀ values of 0.4667 mg/mL, 0.9772 mg/mL, and 0.5014 mg/mL, respectively, highlighting their robust antioxidant potential. This effervescent tablet not only offers a novel approach to harness the benefits of elderberry and vine tea but also presents an innovative concept for antioxidant-enriched functional foods. Future investigations could explore the aromatic characteristics of vine tea to further enhance the nutritional value of the product.

Data Availability

The data used to support the findings of this study are available from the corresponding author upon request.

Conflicts of Interest

The author declares that there is no conflict of interest regarding the publication of this paper.

Acknowledgments

The author gratefully acknowledges the technical and financial assistance from the College of Food Science & Nutritional Engineering, China Agricultural University.

References

- [1] Y. A. M. Elkhateeb and M. R. Alshammary, "Effects of fast foods in relation to free radicals and antioxidants," *American Journal of Laboratory Medicine*, vol. 2, no. 6, pp. 156–162, 2017.
- [2] A. Atlante, D. Valenti, V. Latina, and G. Amadoro, "Role of oxygen radicals in Alzheimer's disease: focus on tau protein," *Oxygen*, vol. 1, no. 2, pp. 96–120, 2021.
- [3] V. Z. Lankin, A. K. Tikhaze, and A. M. Melkumyants, "Malondialdehyde as an important key factor of molecular mechanisms of vascular wall damage under heart diseases development," *International Journal of Molecular Sciences*, vol. 24, no. 1, 2023.

- [4] J. Viña, C. Borrás, and J. Miquel, "Theories of ageing," *IUBMB Life*, vol. 59, no. 4–5, pp. 249–254, 2007.
- [5] T. Miyazawa, C. Abe, G. C. Burdeos, A. Matsumoto, and M. Toda, "Food antioxidants and aging: theory, current evidence and perspectives," *Nutraceuticals*, vol. 2, no. 3, pp. 181–204, 2022.
- [6] R. Domínguez, L. Zhang, G. Rocchetti et al., "Elderberry (*Sambucus nigra* L.) as potential source of antioxidants. Characterization, optimization of extraction parameters and bioactive properties," *Food Chemistry*, vol. 330, article 127266, 2020.
- [7] M. Mikulic-Petkovsek, A. Ivancic, S. Gacnik et al., "Biochemical characterization of black and green mutant elderberry during fruit ripening," *Plants*, vol. 12, no. 3, p. 504, 2023.
- [8] X. Hu, Y. Yang, S. Tang et al., "Anti-aging effects of anthocyanin extracts of *Sambucus canadensis* caused by targeting mitochondrial-induced oxidative stress," *International Journal of Molecular Sciences*, vol. 24, no. 2, p. 1528, 2023.
- [9] O. E. Pascariu and F. Israel-Roming, "Bioactive compounds from elderberry: extraction, health benefits, and food applications," *Processes*, vol. 10, no. 11, p. 2288, 2022.
- [10] T. Murakami, M. Miyakoshi, D. Araho et al., "Hepatoprotective activity of tocha, the stems and leaves of *Ampelopsis grosedentata*, and ampelopsin," *BioFactors*, vol. 21, no. 1–4, pp. 175–178, 2004.
- [11] W. Bi, C. He, Y. Ma et al., "Investigation of free amino acid, total phenolics, antioxidant activity and purine alkaloids to assess the health properties of non-*Camellia* tea," *Acta Pharmaceutica Sinica B*, vol. 6, no. 2, pp. 170–181, 2016.
- [12] R. Cai, X. Li, C. Li et al., "Standards-based UPLC-Q-exactive Orbitrap MS systematically identifies 36 bioactive compounds in *Ampelopsis grosedentata* (vine tea)," *Separations*, vol. 9, no. 11, p. 329, 2022.
- [13] J. Hu, Y. Chen, and D. Ni, "Effect of superfine grinding on quality and antioxidant property of fine green tea powders," *LWT-Food Science and Technology*, vol. 45, no. 1, pp. 8–12, 2012.
- [14] W. Gao, F. Chen, X. Wang, and Q. Meng, "Recent advances in processing food powders by using superfine grinding techniques: a review," *Comprehensive Reviews in Food Science and Food Safety*, vol. 19, no. 4, pp. 2222–2255, 2020.
- [15] M. H. Moghaddam, A. H. Bayat, N. Eskandari et al., "Elderberry diet ameliorates motor function and prevents oxidative stress-induced cell death in rat models of Huntington disease," *Brain Research*, vol. 1762, article 147444, 2021.
- [16] C. L. Millar, G. H. Norris, C. Jiang et al., "Long-term supplementation of black elderberries promotes hyperlipidemia, but reduces liver inflammation and improves HDL function and atherosclerotic plaque stability in apolipoprotein E-knockout mice," *Molecular Nutrition & Food Research*, vol. 62, no. 23, article 1800404, 2018.
- [17] P. Lin, E. Hwang, H. T. T. Ngo, S. A. Seo, and T. H. Yi, "*Sambucus nigra* L. ameliorates UVB-induced photoaging and inflammatory response in human skin keratinocytes," *Cytototechnology*, vol. 71, no. 5, pp. 1003–1017, 2019.
- [18] Y. Sun, S. Liu, S. Yang et al., "Mechanism of dihydromyricetin on inflammatory diseases," *Frontiers in Pharmacology*, vol. 12, article 794563, 2022.
- [19] J. Q. Ma, Y. Z. Sun, Q. L. Ming, Z. K. Tian, H. X. Yang, and C. M. Liu, "Ampelopsin attenuates carbon tetrachloride-induced mouse liver fibrosis and hepatic stellate cell activation associated with the SIRT1/TGF- β 1/Smad3 and autophagy pathway," *International Immunopharmacology*, vol. 77, article 105984, 2019.
- [20] Y. Luo, S. Lu, X. Dong, L. Xu, G. Sun, and X. Sun, "Dihydromyricetin protects human umbilical vein endothelial cells from injury through ERK and Akt mediated Nrf2/HO-1 signaling pathway," *Apoptosis*, vol. 22, no. 8, pp. 1013–1024, 2017.
- [21] P. Moser, T. C. B. Gallo, L. A. C. Zuanon, G. E. Pereira, and V. R. Nicoletti, "Water sorption and stickiness of spray-dried grape juice and anthocyanins stability," *Journal of Food Processing and Preservation*, vol. 42, no. 12, Article ID e13830, 2018.
- [22] L. P. Zea, Y. A. Yusof, M. G. Aziz, C. N. Ling, and N. A. M. Amin, "Compressibility and dissolution characteristics of mixed fruit tablets made from guava and pitaya fruit powders," *Powder Technology*, vol. 247, pp. 112–119, 2013.
- [23] S. Yousefi, Z. Emam-Djomeh, and S. M. Mousavi, "Effect of carrier type and spray drying on the physicochemical properties of powdered and reconstituted pomegranate juice (*Punica granatum* L.)," *Journal of Food Science and Technology*, vol. 48, no. 6, pp. 677–684, 2011.
- [24] L. I. Mulargia, E. Lemmens, K. Korompokis et al., "Tailoring the formulation of sugar-snap cookies to lower *in vitro* starch digestibility: A response surface modelling approach," *Food Chemistry*, vol. 435, article 137601, 2024.
- [25] J. Xu, Y. Liu, N. Zhang, S. Xiong, L. Zhang, and J. Wang, "Effect of superfine grinding on the physicochemical properties of straw mushroom (*Volvariella volvacea*) powders rich in VITAMIND₂," *Journal of Food Processing and Preservation*, vol. 46, no. 12, Article ID e17192, 2022.
- [26] Y. Ji, Q. Wu, W. Shan, B. Zhang, and Z. Zhang, "Preparation and quality evaluation of red ginseng effervescent tablet solid beverage," *Ginseng Research*, vol. 33, no. 3, pp. 9–12, 2021.
- [27] W. J. Xu, C. L. Chen, J. J. Meng, Y. J. Liu, S. Deng, and Y. P. Lv, "Preparation of a blueberry effervescent tablet and effect of its formulation on quality," *Food Research and Development*, vol. 44, no. 1, pp. 102–109, 2023.
- [28] M. Yu, J. Ma, X. Wang et al., "Peanut sprout yogurt: increased antioxidant activity and nutritional content and sensory evaluation by fuzzy mathematics," *Journal of Food Processing and Preservation*, vol. 46, no. 7, Article ID e16663, 2022.
- [29] I. Sopyan, D. Gozali, and R. K. Guntina, "Design-expert software (DOE): an application tool for optimization in pharmaceutical preparations formulation," *International Journal of Applied Pharmaceutics*, vol. 14, pp. 55–63, 2022.
- [30] Chinese Pharmacopoeia Commission, *Pharmacopoeia of the People's Republic of China (2020)*, China Medical Science Press, Beijing, China, 2022.
- [31] D. Zheng, J. Wang, L. Zhang, F. Wang, and H. Wang, "Optimized preparation and quality analysis of bamboo leaf extract effervescent tablets," *Food Science*, vol. 37, no. 8, pp. 39–44, 2016.
- [32] S. H. Lee, H. W. Kim, and H. J. Park, "Plaque removal effectiveness of 3D printed dental hygiene chews with various infill structures through artificial dog teeth," *Heliyon*, vol. 8, no. 3, article e09096, 2022.
- [33] L. E. Ordóñez-Santos, J. Martínez-Girón, and M. E. Arias-Jaramillo, "Effect of ultrasound treatment on visual color, vitamin C, total phenols, and carotenoids content in cape gooseberry juice," *Food Chemistry*, vol. 233, pp. 96–100, 2017.

- [34] G. Jia, H. Zhao, D. Hou, T. Sun, and W. Lin, "Quantitative determination of total flavonoids from *Polygonatum sibiricum* by spectrophotometry," *IOP Conference Series: Materials Science and Engineering*, vol. 677, no. 2, article 022126, 2019.
- [35] D. Kondhare and H. Lade, "Phytochemical profile, aldose reductase inhibitory, and antioxidant activities of Indian traditional medicinal *Coccinia grandis* (L.) fruit extract," *3 Biotech*, vol. 7, no. 6, p. 378, 2017.
- [36] Y. Wu, L. Xu, X. Liu et al., "Effect of thermosonication treatment on blueberry juice quality: total phenolics, flavonoids, anthocyanin, and antioxidant activity," *LWT*, vol. 150, article 112021, 2021.
- [37] K. Thaipong, U. Boonprakob, K. Crosby, L. Cisneros-Zevallos, and D. H. Byrne, "Comparison of ABTS, DPPH, FRAP, and ORAC assays for estimating antioxidant activity from guava fruit extracts," *Journal of Food Composition and Analysis*, vol. 19, no. 6-7, pp. 669-675, 2006.
- [38] F. Liao, R. Wen, L. Yang, R. Yang, Q. He, and E. Zhang, "Oil tea soup formula optimization based on orthogonal test and fuzzy mathematics sensory evaluation," *Journal of Food Science and Engineering*, vol. 7, no. 9, pp. 435-442, 2017.
- [39] B. Chen, *Preparation of the Effervescent Tablets of the Whole Fruit Freeze-Dried Powder of Indigo Fruit*, Shenyang Agricultural University, 2021.
- [40] R. C. V. Carneiro, L. Ye, N. Baek, G. H. A. Teixeira, and S. F. O'Keefe, "Vine tea (*Ampelopsis grossedentata*): A review of chemical composition, functional properties, and potential food applications," *Journal of Functional Foods*, vol. 76, article 104317, 2021.
- [41] D. Wang, J. Liu, J. Lu, and S. Zheng, "ChemInform abstract: two new Limonoids of *Ampelopsis grossedentata* Hand.-Mazz," *Indian Journal of Chemistry*, vol. 30, no. 27, pp. 240-242, 1999.
- [42] A. Roy and S. Saraf, "Limonoids: overview of significant bioactive triterpenes distributed in plants kingdom," *Biological & Pharmaceutical Bulletin*, vol. 29, no. 2, pp. 191-201, 2006.
- [43] A. M. Orellana-Paucar, "Steviol glycosides from *Stevia rebaudiana*: an updated overview of their sweetening activity, pharmacological properties, and safety aspects," *Molecules*, vol. 28, no. 3, p. 1258, 2023.
- [44] M. I. Espinoza, J. P. Vincken, M. Sanders, C. Castro, M. Stieger, and E. Agosin, "Identification, quantification, and sensory characterization of Steviol glycosides from differently processed *Stevia rebaudiana* commercial extracts," *Journal of Agricultural and Food Chemistry*, vol. 62, no. 49, pp. 11797-11804, 2014.
- [45] T. Pasqualli, P. E. E. Chaves, C. P. L. Da Veiga, É. A. Serpa, L. F. S. D. Oliveira, and M. M. Machado, "Steviol, the active principle of the stevia sweetener, causes a reduction of the cells of the immunological system even consumed in low concentrations," *Immunopharmacology and Immunotoxicology*, vol. 42, no. 5, pp. 504-508, 2020.
- [46] S. Skąpska, K. Marszałek, Ł. Woźniak, J. Szczepańska, J. Danielczuk, and K. Zawada, "The development and consumer acceptance of functional fruit-herbal beverages," *Foods*, vol. 9, no. 12, p. 1819, 2020.
- [47] Y. Li, "Effects of different metal ions on the stability of anthocyanins as indicators," *IOP Conference Series: Earth and Environmental Science*, vol. 300, no. 5, article 52015, 2019.
- [48] S. Paul, L. J. Taylor, B. Murphy et al., "Powder properties and compaction parameters that influence punch sticking propensity of pharmaceuticals," *International Journal of Pharmaceutics*, vol. 521, no. 1-2, pp. 374-383, 2017.
- [49] F. Oboh and J. Imafidon, "Antioxidant and sensory properties of new beverage formulations composed of palm sugar, *Aframomum melegueta*, and citric acid," *Beverages*, vol. 4, no. 3, p. 59, 2018.
- [50] M. Meisner, P. Duda, B. Szulc-Musioł, and B. Sarecka-Hujar, "Characteristics of commercial effervescent tablets using selected pharmacopeial and novel analytical methods," *Applied Sciences*, vol. 13, no. 5, p. 3171, 2023.
- [51] A. D. Kinghorn and D. D. Soejarto, "Discovery of terpenoid and phenolic sweeteners from plants," *Pure and Applied Chemistry*, vol. 74, no. 7, pp. 1169-1179, 2002.
- [52] M. Sotoyama, S. Uchida, S. Tanaka et al., "Citric acid suppresses the bitter taste of olopatadine hydrochloride orally disintegrating tablets," *Biological & Pharmaceutical Bulletin*, vol. 40, no. 4, pp. 451-457, 2017.
- [53] A. Vilela, "Microbial dynamics in sour-sweet wine vinegar: impacts on chemical and sensory composition," *Applied Sciences*, vol. 13, no. 13, p. 7366, 2023.
- [54] P. A. Apsari, D. N. E. Sari, A. P. Kusuma, and O. Indrati, "Effervescent tablet formulation melinjo seed extract (*Gnetum gnemon* L.) using PEG 6000 as lubricant and citric Acid - Tartaric acids as acid sources," *Jurnal Eksakta*, vol. 18, no. 1, pp. 30-41, 2018.
- [55] J. Suksaeree, C. Monton, L. Charoenchai, N. Chankana, and T. Wunnakup, "Optimization of process and formulation variables for Semha-Pinas extract effervescent tablets using the Box-Behnken design," *AAPS PharmSciTech*, vol. 24, no. 1, p. 52, 2023.
- [56] N. P. Minh, "Production of effervescent tablet from jambolan (*Syzygium cumini* L.) fruit," *Research on Crops*, vol. 23, pp. 888-895, 2022.
- [57] R. Harun, A. Mahmuda, U. Jalal et al., "Antioxidant, cytotoxic, antibacterial and thrombolytic activities of *Centella asiatica* L.: possible role of phenolics and flavonoids," *Clinical Phytoscience*, vol. 9, no. 1, 2023.
- [58] S. Chaurasia, "Phytochemical analysis, antioxidant potential and radical scavenging activity of *Lomatium dissectum*: an ancient plant of North America," *Biology and Life Sciences Forum*, vol. 12, no. 1, 2022.
- [59] S. Wu, Y. Yuan, J. Yin et al., "Characteristics of effervescent tablets of *Aronia melanocarpa*: response surface design and antioxidant activity evaluation," *Journal of Food Measurement and Characterization*, vol. 16, no. 4, pp. 2969-2977, 2022.
- [60] J. Zhu, Y. Liang, C. Wu et al., "Process optimization for development of a novel solid beverage with high antioxidant activity and acceptability from fermented Ginkgo biloba seeds," *Journal of Food Measurement and Characterization*, vol. 16, no. 6, pp. 4630-4640, 2022.
- [61] A. C. Iwansyah, H. Fauzi, W. Cahyadi et al., "Development, physiochemical and sensory evaluation of a new effervescent tablet formulation based on *Moringa oleifera* leaves extract," *International Journal of Food Engineering*, vol. 19, no. 3-4, pp. 133-141, 2023.