A NUTRACEUTICAL FORMULATED DRINK FROM EDIBLE BIRD'S NEST AND COLLAGEN WITH HIGH BIOAVAILABILITY PROPERTIES

SHAMALA DEVI A/P NADARAJU

A project submitted in fulfilment of the requirements for the award of the degree of Master of Science

School of Chemical and Energy Engineering Faculty of Engineering Universiti Teknologi Malaysia

AUGUST 2022

ACKNOWLEDGEMENT

In preparing this thesis, I was in contact with many people, researchers, academicians, and practitioners. They have contributed towards my understanding and thoughts. In particular, I wish to express my sincere appreciation to my supervisor, Dr Lee Ting Hun, for encouragement, guidance, critics and friendship. I am also very thankful to Dr Fadilah Hamdan and Mr Lee Chia Hau for their guidance, advices and motivation. Without their continued support and interest, this thesis would not have been the same as presented here.

I am also indebted to my classmates for the stimulating discussion and encouragement. My sincere appreciation also extends to all my lecturers who have provided assistance at various occasions. Their views and tips are useful indeed.

Last but not least, I would like to thank my parents, sisters, cousins, friends and family for supporting me spiritually throughout the successful of this study journey. With their love, patient, and relentless encouragement that make this work more complete.

ABSTRACT

Nutraceuticals drinks formulated from phytochemicals or functional foods that contain natural bioactive and chemical compounds contributes to healthpromoting, disease-preventing or therapeutic properties. In current trends, there is a high demand for nutraceutical products and functional food, particularly drinks or beverages. Formulation of nutraceutical drink containing Edible Bird Nest's (EBN) extract and collagen type II can result in preventing bone health disease as it gives great benefits and improve the bone strength. The aim of this study is to develop a nutraceutical drink from EBN extract and collagen type II for bone health properties with high bioavailability of collagen content. In this study, six different formulation was carried out with different concentrations of EBN extract (0%, 0.03% 0.025%, 0.050%, 0.075%, 1%) and two samples of collagen drink were purchased for benchmarking from market such as Kinohibitsu and Ecolite. All sample were analysed with *in vitro* bioavailability analysis and sensory analysis using Hedonic Test. Findings indicated that F4 (formulation 4) with 1% of EBN extract and 0.03% collagen type II showed the best results and contribute to high bioavailability of collagen (85.61%) when comparing with formulation of EBN extract with C1 (control 1) 0.03% of collagen type II, without EBN and F1 0.03% of collagen type II, EBN 0.025%. Findings indicated that with the increase in the addition of EBN extract caused significant difference (p<0.05) in absorption of total collagen into the body when compare with C1 and F1. Sensory analysis findings revealed EBN collagen drink of F4 with 1% of EBN extract and 0.03% collagen type II is the most preferred among consumers. It caused significant difference (p<0.05) in appearance, color, aroma, taste and overall acceptability for sensory analysis. In summary, nutraceutical drink of formulation 4 of 1% EBN extract and 0.03% collagen type II has high bioavailability of collagen and consumer acceptability.

ABSTRAK

Minuman Nutraseutikal yang dirumus daripada fitokimia atau makanan berfungsi yang mengandungi sebatian bioaktif dan kimia semulajadi menyumbang kepada sifat menggalakkan kesihatan, mencegah penyakit atau terapeutik. Dalam trend semasa, terdapat permintaan yang tinggi untuk produk nutraseutikal dan makanan berfungsi, terutamanya minuman. Formulasi minuman nutraseutikal yang mengandungi ekstrak Edible Bird Nest (EBN) dan kolagen jenis II boleh menyebabkan mencegah penyakit kesihatan tulang kerana ia memberi manfaat yang besar dan meningkatkan kekuatan tulang. Matlamat kajian ini adalah untuk membuat minuman nutraseutikal daripada ekstrak EBN dan kolagen jenis II untuk sifat kesihatan tulang dengan bioavailabiliti tinggi kandungan kolagen. Dalam kajian ini, enam rumusan berbeza telah dijalankan dengan kepekatan berbeza ekstrak EBN (0%, 0.03% 0.025%, 0.050%, 0.075%, 1%) dan dua sampel minuman kolagen telah dibeli untuk penanda aras daripada pasaran seperti Kinohibitsu dan Ecolite. Semua sampel dianalisis untuk analisis bioavailabiliti in vitro dan analisis deria menggunakan Ujian Hedonik. Dapatan menunjukkan bahawa F4 (formulasi 4) dengan 1% ekstrak EBN dan 0.03% kolagen jenis II menunjukkan hasil terbaik dan menyumbang kepada bioavailabiliti tinggi kolagen (85.61%) apabila dibandingkan dengan formulasi ekstrak EBN dengan C1 (kawalan 1) 0.03% kolagen jenis II, tanpa EBN dan F1 (formulasi 1) 0.03% daripada kolagen jenis II, EBN 0.025%. Dapatan kajian menunjukkan peningkatan dalam penambahan ekstrak EBN menyebabkan perbezaan ketara (p<0.05) dalam penyerapan jumlah kolagen ke dalam badan jika dibandingkan dengan C1 dan F1. Penemuan analisis deria mendedahkan minuman kolagen EBN F4 dengan 1% ekstrak EBN dan 0.03% kolagen jenis II adalah yang paling disukai di kalangan pengguna. Ia menyebabkan perbezaan ketara (p<0.05) dalam rupa, warna, aroma, rasa dan penerimaan keseluruhan untuk analisis deria. Secara ringkasnya, minuman nutraseutikal F4 dengan 1% ekstrak EBN dan 0.03% kolagen jenis II mempunyai bioavailabiliti tinggi kolagen dan kebolehterimaan pengguna.

TABLE OF CONTENTS

TITLE

	DECLARATION			iii	
	DEDICATION				
	ACKNOWLEDGEMENT				
	ABST	RACT		vi	
	ABSTRAK				
	LIST OF TABLES			xi	
	LIST OF FIGURES			xii	
	LIST OF ABBREVIATIONS			xiii	
	LIST	OF SYMI	BOLS	xiv	
CHAPTER	81	INTROI	DUCTION	1	
	1.1	Backgrou	and of study	1	
	1.2	Problem Statement		4	
	1.3	Objectives		5	
	1.4	Scope of	Objectives	6	
	1.5	Significa	nt of study	7	
CHAPTER	2	LITERA	TURE REVIEW	9	
	2.1	Nutraceu	tical and Functional Food	9	
	2.2	Edible Bi	ird Nest (EBN)	10	
		2.2.1 C	urrent Economy Trend of EBN	11	
		2.2.2 C	omposition of EBN	11	
	2.3	EBN App	plication in Food Products	16	
	2.4	EBN App	plication in Therapeutic Products	16	
	2.5	Pharmaco	ological Properties of EBN	17	
			BN Anti-inflammatory and Enhance Immune coperties	17	
		2.5.2 E	BN Anti-hypertension Properties	18	

		2.5.3	EBN An	tioxidant and Anti-aging Properties.	19
		2.5.4	Mechani Propertie	ism of EBN in Improving Bone Health	20
	2.6	Collag	gen		22
		2.6.1	Collager	n Type II	23
		2.6.2	Structure	e of Collagen	24
		2.6.3	Sources	of Collagen	25
		2.6.4	Biologic	al Properties of Collagen	25
		2.6.5	Collager	Application	26
2.7		Rosehip Extract			28
		2.7.1	Biologic Rosehip	al Properties and Application of Extract	29
	2.8	Amin	o Acid (L	ysine)	30
	2.9	Ascor	bic Acid		31
	2.10	Peach	Extract		33
		2.10.1	In Vitro	Bioavailability Analysis	34
	2.11	Senso	ry Study		35
CHAPTER	3	RESE	CARCH N	IETHODOLOGY	37
	3.1	Mater	ials		39
	3.2	EBN (Collagen I	Drink Preparation	39
	3.3	Formulation of EBN Collagen Drink			39
	3.4	Sample Analy		S	40
		3.4.1	In vitro I	Bioavailability Collagen Analysis	41
			3.4.1.1	Preparation of Pancretin-bile Acid Solution	41
			3.4.1.2	Preparation of Dialysis Tube	42
		3.4.2	Sensory	Analysis	42
		3.4.3	Statistic	al Analysis	43
CHAPTER	4	RESU	JLT & DI	SCUSSION	45
	4.1	Formu	lation of	EBN Collagen drink	45
	4.2	In Viti	ro Bioava	ilability Collagen Analysis	47
	4.3	Sensory Study			51

CHAPTER 5 CONCLUSION

REFERENCES

57

55

LIST OF TABLES

TABLE NO.	TITLE	PAGE
Table 2.1	EBN Composition Report by Different Researchers.	14
Table 2.2	Pharmacological Properties of (EBN) Beneficial to Human Health and its Contributing Nutritional Component.	21
Table 2.3	Different Types of Collagen and its Benefit to Human Health (Kwatra, 2020).	23
Table 3.1	EBN collagen drink formulation	40
Table 4.1	The appearance of EBN Collagen Drink with different EBN extract concentration	46
Table 4.2	The <i>in vitro</i> bioavailability of total collagen present in the EBN Collagen Drink and other sample obtained from Market	50
Table 4.3	Sensory acceptance of EBN Collagen Drink and other sample obtained from Market	53

LIST OF FIGURES

FIGURE NO.	TITLE	PAGE
Figure 3.1	Flow chart of research activities for stage 1 and stage 2	38
Figure 4.1	<i>In vitro</i> bioavailability of total collagen present in the EBN collagen drink and other sample obtained from market.	51
Figure 4.2	Sensory acceptance of EBN collagen drink and other sample obtained from market.	54

LIST OF SYMBOLS

°C	-	Degree Celcius
%	-	Percentage
-	-	Negative
+	-	Positive
<	-	Less than
>	-	More than
±	-	Plus-Minus
Р	-	Probability
hr	-	Hour

Collagen added beverages are the new current trend on the global market now (Bilek & Bayram, 2015). Manufactures and product development scientists have formulated and developed varieties of collagen-based drink products such as cappuccino collagen, soy collagen, cocoa collagen and juice-based collagen (León-López et al., 2020) Adding collagen to develop functional beverages contribute to a good source of benefits. In the composition of human diets, they contribute valuable nutritive fibres and protein sources (Neklyudov, 2003). Collagen synthesis reduces when human age, causing tissues to become thinner, weaker, and less elastic. Adding collagen to develop beverages help consumers maintain healthy body tissue and body structure (King'ori, 2011).

Collagen provides healthy creatine and amino acid in new muscle growth after exercises (Czajka et al., 2018a). Collagen added beverages are beneficial to health and essential in maintaining good health. They help to sustain and improve skin hydration, elasticity, and reduce osteoarthritis pain. Studies have shown that daily consumption of collagen helps in slowing the aging process by making bones denser and less brittle (Czajka et al., 2018a). Collagen added beverages helps reduce wrinkles on the face or skin and increase hair's quantity, scalp coverage, and thickness. Nails grow faster and reduce to break or chip by consuming collagen beverages.

Collagen can be found and obtain from many natural sources. One of the promising collagen source can be found in Edible Bird's Nest (EBN) extract. EBN consumption provides multiple advantages and benefits to human health (Chye et al., 2017a). EBN has been used widely as functional food and therapeutic remedies due to its essential composition and biological activity. It has been applied as an intermediate ingredient for many food products to improve product's the variety, quality, taste, and bioavailability according to consumers need. Based on the previous studies' conducted, many value-added products were created using EBN such as beverages, noodles, chocolates, and ice-creams (Hanisah et al., 2021). EBN consists high percentage of protein and carbohydrates, followed by ash, fat, and minerals. The protein and carbohydrates content in EBN make up 60% and 30% of the overall EBN, respectively. EBN claims to be healthy and has a high nutritional

level of alcohol consumption, smoking, less exercise, as well as aging. This collagen contains all nine essential amino acids including glycine, and its proline concentration is close to twenty times higher than that found in other protein-rich foods that help to enhance this formulation (Sionkowska et al., 2020). Additionally, collagen intake also helps to decrease joint pain, reduce wrinkles, and improve skin with anti-aging properties (Czajka et al., 2018a).

Adding EBN extract to the diet helps to prevent bone diseases such as osteoarthritis (Chua et al., 2013). EBN are rich in protein-containing glycosaminoglycans, including non-sulfated chondroitin glycosaminoglycan that has similar properties to the matrix of cartilage and it helps to promote the collagen content with high bioavailability and absorbed into the body easily. EBN extracts and its beneficial bioactive compounds will be considered an asset for this formulation of a nutraceutical drink in preventing bone diseases. EBN extract enhances collagen content and contribute effectively to the formulation of drinks that are absorbed into the body, and indicate high bioavailability. Therefore, EBN helps to prevent osteoarthritis and regenerate cartilage at the same time (Chua et al., 2013).

Additional functional ingredients are therefore needed in order to create effective functional beverages. Having ordinary diet drinks is good for health and it would be better to consume drinks that have added ingredients with greater value such as EBN extract and collagen type II. Therefore, the consumption of nutraceutical drinks that developed with EBN and collagen type II extract is more nutritious and effective (Bilek & Bayram, 2015).

1.3 Objectives

The objectives of the research are:

To formulate a nutraceutical drink containing EBN extract and collagen type
 II for bone health properties with high bioavailability of collagen content.

- 2. To investigate the bioavailability of total collagen of formulated EBN collagen drink through *in vitro* study and benchmarking against the collected similar market samples.
- 3. To evaluate the sensory attributes of the formulated EBN collagen drink through sensory study

1.4 Scope of Objectives

The scopes of the research are:

- Formulating nutraceutical drink with six different formulation containing EBN extract and collagen type II as active ingredients and rosehip extract, amino acid, ascorbic acid and peach extract as supplemented ingredients. The EBN extract concentration is varies for all the six formulation and EBN extract limits to 1% due to the price from EBN collagen drink in the market.
- 2. Investigating the bioavailability of total collagen in six different formulation of nutraceutical EBN collagen drink and two samples drink that purchased from market through *in vitro* study. *In vitro* digestion study help to examine the absorption and release of food components under simulated gastrointestinal conditions.
- 3. Evaluating the sensory attributes of the nutraceutical EBN collagen drink and two sample drink that purchased from market in terms of appearance, colour, aroma, taste and overall acceptability through sensory study. The sensory test is to perform with 20 panellist and to select the most preferred formulations.

1.5 Significant of study

This study is expected to produce a formulation of a nutraceutical drink using EBN extract and collagen type II as its active ingredient for bone health properties with high bioavailability of collagen content. Additional ingredient like rosehip extract, ascorbic acid, amino acid and peach extract create effectiveness to the drink. This formulation is beneficial for many range of consumers especially active people and the elder generation since it also maintain healthy gut and promotes immune function. Bone health problem is a chronic condition that effects millions people in the world. Examples are people with bone infections, osteoporosis, rickets and osteoarthritis. Current drugs are more to reducing pain and symptoms, yet there are side effects. This formulation will increase consumer demand for all natural and non-modified functional ingredient.

Apart from that, this formulation is easy to consume with improve taste and can be market with affordable price for all range of consumers. It also can emergence with variety of EBN based product that will create wider market and consumer will have more choices to choose. Therefore, this study help to produce a formulation that have good and excellent consumer acceptance and palatable. In addition, all the findings from this research will contribute to a good source of healthy beverage to the consumers with bone health problem in the market.

REFERENCES

- Abeln, M., Albers, I., Peters-Bernard, U., Flächsig-Schulz, K., Kats, E., Kispert, A., Tomlinson, S., Gerardy-Schahn, R., Münster-Kühnel, A., & Weinhold, B. (2019). Sialic Acid Is A Critical Fetal Defense Against Maternal Complement Attack. *The Journal of Clinical Investigation*. 422–436.
- Akhtar, N., & Haqqi, T. M. (2012). Current Nutraceuticals In The Management of Osteoarthritis: A Review. *Therapeutic Advances in Musculoskeletal Disease*. 4(3), 181.
- Albishtue, A. A., Yimer, N., Zakaria, M. Z. A., Haron, A. W., Yusoff, R., Assi, M. A., & Almhanawi, B. H. (2018). Edible Bird's Nest Impact On Rats' Uterine Histomorphology, Expressions of Genes of Growth Factors And Proliferating Cell Nuclear Antigen, And Oxidative Stress Level. *Veterinary World*. 71–79.
- Arseni, L., Lombardi, A., & Orioli, D. (2018). From Structure to Phenotype: Impact of Collagen Alterations on Human Health. *International Journal of Molecular Sciences.* 1407.
- Avila R, I., Rodríguez Barroso, L. G., & Sánchez, M. L. (2018). Collagen: A Review On Its Sources And Potential Cosmetic Applications. *Journal of Cosmetic Dermatology*. 20–26.
- Azmi, N. A., Lee, T. H., Lee, C. H., Hamdan, N., & Cheng, K. K. (2021). Differentiation Unclean And Cleaned Edible Bird's Nest Using Multivariate Analysis of Amino Acid Composition Data. *Pertanika Journal of Science and Technology*. 677–691.
- Babji, A. S., Nurfatin, M. H., Etty Syarmila, I. K., & Masitah, M. (2015). Secrets of Edible Bird Nest.
- Bach, F., Helm, C. V., Bellettini, M. B., Maciel, G. M., & Haminiuk, C. W. I. (2017). Edible Mushrooms: A Potential Source of Essential Amino Acids, Glucans And Minerals. *International Journal of Food Science & Technology*. 2382–2392.
- Baker, D. H. (2007). Lysine, Arginine, and Related Amino Acids: An Introduction to the 6th Amino Acid Assessment Workshop. *The Journal of Nutrition*. 1599-1601.

- Banga, S., Kumar, V., Suri, S., Kaushal, M., Prasad, R., & Kaur, S. (2019). Nutraceutical Potential of Diet Drinks: A Critical Review on Components, Health Effects, and Consumer Safety. 39(3), 272–286.
- Bashir, M. J. K., Xian, T. M., Shehzad, A., Sethupahi, S., Choon Aun, N., & Abu Amr, S. (2016). Sequential Treatment For Landfill Leachate By Applying Coagulation-Adsorption Process. *Geosystem engineering*. 9-20.
- Bello, A. B., Kim, D., Kim, D., Park, H., & Lee, S. H. (2020). Engineering and Functionalization of Gelatin Biomaterials: From Cell Culture to Medical Applications. *Tissue Engineering Part B: Reviews*. 164-180.
- Benjamin, R. M. (2010). Bone Health: Preventing Osteoporosis. Public Health Reports. 125(3), 368.
- Bento, C., Gonçalves, A. C., Silva, B., & Silva, L. R. (2020). Peach (Prunus Persica):Phytochemicals and Health Benefits. *Food Reviews International*. 1-32.
- Bilek, S. E., & Bayram, S. K. (2015). Fruit Juice Drink Production Containing Hydrolyzed Collagen. *Journal of Functional Foods*. 14, 562–569.
- Blidi, O. El, Omari, N. El, Balahbib, A., Ghchime, R., Menyiy, N. El, Ibrahimi, A., Kaddour, K. Ben, Bouyahya, A., Chokairi, O., & Barkiyou, M. (2021).
 Extraction Methods, Characterization And Biomedical Applications Of Collagen: A Review. *Biointerface Research in Applied Chemistry*. 13587–13613.
- Carolina Font, I, F., Gradziel, T. M., Gogorcena, Y., & Moreno, M. Á. (2014). Phenotypic Diversity Among Local Spanish And Foreign Peach And Nectarine [Prunus Persica (L.) Batsch] Accessions. *Euphytica*. 261-277.
- Castaneda-Arriaga, R., Pérez-González, A., Reina, M., Alvarez-Idaboy, J. R., & Galano, A. (2018). Comprehensive Investigation of the Antioxidant and Prooxidant Effects of Phenolic Compounds: A Double-Edged Sword in the Context of Oxidative Stress. *Journal of Physical Chemistry B*. 6198–6214.
- Chambial, S., Dwivedi, S., Shukla, K. K., John, P. J., & Sharma, P. (2013). Vitamin C in Disease Prevention and Cure: An Overview. *Indian Journal of Clinical Biochemistry*. 28(4), 314.
- Chan, F. K. L., Cryer, B., Goldstein, J. L., Lanas, A., Peura, D. A., Scheiman, J. M., Simon, L. S., Singh, G., Stillman, M. J., Wilcox, C. M., Berger, M. F., Breazna, A., & Dodge, W. (2010). A Novel Composite Endpoint To Evaluate The Gastrointestinal (GI) Effects Of Nonsteroidal Antiinflammatory Drugs

Through The Entire GI Tract. The Journal of Rheumatology. 37(1), 167–174.

- Chapko, M. J., & Seo, H. S. (2019). Characterizing Product Temperature-Dependent Sensory Perception Of Brewed Coffee Beverages: Descriptive Sensory Analysis. *Food Research International*. 612–621.
- Chua, K. H., Lee, T. H., Nagandran, K., Md Yahaya, N. H., Lee, C. T., Tjih, E. T. T.,
 & Abdul Aziz, R. (2013). Edible Bird's Nest Extract As A Chondro-Protective Agent For Human Chondrocytes Isolated From Osteoarthritic Knee: In Vitro Study. *BMC Complementary and Alternative Medicine*. 13.
- Chua, L. S., & Zukefli, S. N. (2016). A Comprehensive Review of Edible Bird Nests And Swiftlet Farming. *Journal of Integrative Medicine*. 14(6), 415–428.
- Chye, S. M., Kin Tai, S., Yian Koh, R., & Yen Ng, K. (2017a). A Mini Review on Medicinal Effects of Edible Bird's Nest. Letters in Health and Biological Sciences. 1–3.
- Chye, S. M., Kin Tai, S., Yian Koh, R., Yen Ng, K. (2017b). A Mini Review On Medicinal Effects Of Edible Bird's Nest. Letters in Health and Biological Sciences. 65-7.
- Cong, L., Bremer, P., & Mirosa, M. (2020). Functional Beverages in Selected Countries of Asia Pacific Region. *A Review Beverages 2020.* 6(2), 21.
- Czajka, A., Kania, E. M., Genovese, L., Corbo, A., Merone, G., Luci, C., & Sibilla, S. (2018a). Daily Oral Supplementation With Collagen Peptides Combined With Vitamins And Other Bioactive Compounds Improves Skin Elasticity And Has A Beneficial Effect On Joint And General Wellbeing. *Nutrition Research*. 57, 97–108.
- Czajka, A., Kania, E. M., Genovese, L., Corbo, A., Merone, G., Luci, C., & Sibilla, S. (2018b). Daily Oral Supplementation With Collagen Peptides Combined With Vitamins And Other Bioactive Compounds Improves Skin Elasticity And Has A Beneficial Effect On Joint And General Wellbeing. *Nutrition Research.* 57, 97–108.
- Dai, Y., Cao, J., Wang, Y., Chen, Y., & Jiang, L. (2021). A Comprehensive Review Of Edible Bird's Nest. *Food Research International*. 140.
- Daneault, A., Prawitt, J., Fabien Soulé, V., Coxam, V., & Wittrant, Y. (2017). Biological Effect Of Hydrolyzed Collagen On Bone Metabolism. *Critical Reviews in Food Science and Nutrition*. 1922-1937
- Daud, N. 'Aliah, Sarbini, S. R., Babji, A. S., Mohamad Yusop, S., & Lim, S. J.

(2019). Characterization of Edible Swiftlet's Nest As A Prebiotic Ingredient Using A Simulated Colon Model. *Annals of Microbiology*. 69(12), 1235–1246.

- Daud, N., Mohamad Yusop, S., Babji, A. S., Lim, S. J., Sarbini, S. R., & Hui Yan, T. (2021). Edible Bird's Nest: Physicochemical Properties, Production, and Application of Bioactive Extracts and Glycopeptides. *Food Reviews International*. 177–196.
- Davidović, S. M., Veljović, M. S., Pantelić, M. M., Baošić, R. M., Natić, M. M., Dabić, D. Č., Pecić, S. P., & Vukosavljević, P. V. (2013). Physicochemical, Antioxidant And Sensory Properties Of Peach Wine Made From Redhaven Cultivar. *Journal of Agricultural and Food Chemistry*. 1357–1363.
- Dickinson, E. (2017). Biopolymer-Based Particles As Stabilizing Agents For Emulsions And Foams. *Food Hydrocolloids*. 219–231.
- Du, J., Cullen, J. J., & Buettner, G. R. (2012). Ascorbic Acid: Chemistry, Biology And The Treatment Of Cancer. Biochimica et Biophysica Acta (BBA) -Reviews on Cancer. 1826(2), 443–457.
- Egerton, S., Culloty, S., Whooley, J., Stanton, C., & Ross, R. P. (2018). Characterization Of Protein Hydrolysates From Blue Whiting (Micromesistius Poutassou) And Their Application In Beverage Fortification. *Food Chemistry*. 698–706.
- Farage, M. A., Miller, K. W., & Maibach, H. I. (2017). Degenerative Changes in Aging Skin. *Textbook of Aging Skin.* 15–30.
- Feng, M., & Betti, M. (2017). Transepithelial Transport Efficiency Of Bovine Collagen Hydrolysates In A Human Caco-2 Cell Line Model. *Food Chemistry*. 224, 242–250.
- Figueiredo, J. S. B., Santos, G. L. M., Lopes, J. P. A., Fernandes, L. B., Silva, F. N.,
 Faria, R. B., Rocha, A. C. S., Farias, P. K. S., Lima, W. J. N., Durães, C. A.
 F., Xavier, A. R. E. de O., de CARVALHO, B. M. A., Careli, R. T., de
 ALMEIDA, A. C., & Brandi, I. V. (2019). Sensory Evaluation Of Fermented
 Dairy Beverages Supplemented With Iron And Added By Cerrado Fruit
 Pulps. *Food Science and Technology*. 39, 410–414.
- Fini, M., Torricelli, P., Giavaresi, G., Carpi, A., Nicolini, A., & Giardino, R. (2001). Effect of L-Lysine And L-Arginine On Primary Osteoblast Cultures From Normal And Osteopenic Rats. *Biomedicine & Pharmacotherapy*. 213–220.

- Fizl, S. M., Lim, C. K., & Mustafa, A. R. (2013). Roosting And Nest-Building Behaviour of The White-Nest Swiftlet Aerodramus Fuciphagus (Thunberg) (Aves: Apodidae) in Farmed Colonies. *Raffles Bulletin of Zoology*. 225-235.
- García-Coronado, J. M., Martínez-Olvera, L., Elizondo-Omaña, R. E., Acosta-Olivo,
 C. A., Vilchez-Cavazos, F., Simental-Mendía, L. E., & Simental-Mendía, M.
 (2019). Effect Of Collagen Supplementation On Osteoarthritis Symptoms: A
 Meta-Analysis Of Randomized Placebo-Controlled Trials. *International Orthopaedics*. 43, 531–538.
- García-Lafuente, A., Guillamón, E., Villares, A., Rostagno, M. A., & Martínez, J. A. (2009). Flavonoids As Anti-Inflammatory Agents: Implications In Cancer And Cardiovascular Disease. *Inflammation Research : Official Journal of the European Histamine Research Society*. 537–552.
- Gaspar-Pintiliescu, A., Stanciuc, A. M., & Craciunescu, O. (2019). Natural Composite Dressings Based On Collagen, Gelatin And Plant Bioactive Compounds For Wound Healing: A Review. *International Journal of Biological Macromolecules*. 138, 854–865.
- Ghassem, M., Arihara, K., Mohammadi, S., Sani, N. A., & Babji, A. S. (2017). Identification Of Two Novel Antioxidant Peptides From Edible Bird's Nest (Aerodramus Fuciphagus) Protein Hydrolysates. *Food & Function*. 2046– 2052.
- Gottardi, R., Hansen, U., Raiteri, R., Loparic, M., Düggelin, M., Mathys, D., Friederich, N. F., Bruckner, P., & Stolz, M. (2016). Supramolecular Organization of Collagen Fibrils in Healthy and Osteoarthritic Human Knee and Hip Joint Cartilage. Plos one. 10-11.
- Gul, K., Singh, A. K., & Jabeen, R. (2015). Nutraceuticals and Functional Foods: The Foods for the Future World. Critical reviews in food science and nutrition, 56(16), 2617-2627.
- Güler, E., Bak, T., Karadeniz, T., & Muradoğlu, F. (2021). Relationships of Fruit Characteristics of Rosehips (Rosa canina L.) Grown in Bolu City Center. *Journal of the Institute of Science and Technology*. 831–838.
- Haghani, A., Mehrbod, P., Safi, N., Aminuddin, N. A., Bahadoran, A., Omar, A. R., & Ideris, A. (2016). In Vitro And In Vivo Mechanism Of Immunomodulatory And Antiviral Activity Of Edible Bird's Nest (EBN) Against Influenza A Virus (IAV) Infection. *Journal of Ethnopharmacology*. 185, 327–340.

- Hamzah, Z., Hulwani Ibrahim, N., Hussin, K., Hashim, O., & Lee, B.-B. (2013). Nutritional Properties of Edible Bird Nest. *Journal of Asian Scientific Research*, 3, 600–607.
- Hanisah, E., Farahniza, Z., Maaruf, A., & Abdul Salam, B. (2021). Antioxidative
 Properties of Edible Bird's Nest Mincroparticulates Incorporated Into Red
 Dates Drink. In Edible Bird Nest Industry Conference, Putrajaya, Malaysia.
 58
- Hao, Q. L., & Rahman, A. O. (2016). Swiftlets and Edible Bird's Nest Industry in Asia. Pertanika Journal of Scholarly Research Reviews. 2(1).
- Harris, R. B., Fonseca, F. L. A., Sharp, M. H., & Ottinger, C. R. (2021). Functional Characterization of Undenatured Type II Collagen Supplements: Are They Interchangeable. *Journal of Dietary Supplements*, 1-16.
- He, Y., Li, Z., Alexander, P. G., Ocasio-Nieves, B. D., Yocum, L., Lin, H., & Tuan,
 R. S. (2020). Pathogenesis of Osteoarthritis: Risk Factors, Regulatory
 Pathways in Chondrocytes, and Experimental Models. *Biology*. 1–32.
- Hou, Z., Imam, M. U., Ismail, M., Azmi, N. H., Ismail, N., Ideris, A., & Mahmud, R. (2015). Lactoferrin And Ovotransferrin Contribute Toward Antioxidative Effects Of Edible Bird's Nest Against Hydrogen Peroxide-Induced Oxidative Stress In Human SH-SY5Y Cells. *Bioscience, Biotechnology, and Biochemistry*. 1570–1578.
- Huang, T., Tu, Z., Zou, Z., Shangguan, X., Wang, H., & Bansal, N. (2020). Glycosylated Fish Gelatin Emulsion: Rheological, Tribological Properties And Its Application As Model Coffee Creamers. *Food Hydrocolloids*. 102.
- Hur, S. J., Lim, B. O., Decker, E. A., & McClements, D. J. (2011). In Vitro Human Digestion Models For Food Applications. *Food Chemistry*. 125(1), 1–12.
- Hwang, E., Park, S. W., & Yang, J.-E. (2020). Anti-Aging, Anti-Inflammatory, And Wound-Healing Activities of Edible Bird's Nest In Human Skin Keratinocytes And Fibroblasts. *Pharmacognosy Magazine*. 16(69), 336.
- Kaur, G., Kumar, V., Goyal, A., Tanwar, B., & Kaur, J. (2018). Optimization of Nutritional Beverage Developed From Radish, Sugarcane And Herbal Extract Using Response Surface Methodology. *Nutrition and Food Science*. 48(5), 733–743.
- King'ori, A. M. (2011). A Review Of The Uses Of Poultry Eggshells And Shell Membranes. *International Journal of Poultry Science*. 10(11), 908–912.

- Kishimoto, Y., Saito, N., Kurita, K., Shimokado, K., Maruyama, N., & Ishigami, A. (2013). Ascorbic Acid Enhances The Expression Of Type 1 And Type 4 Collagen And SVCT2 In Cultured Human Skin Fibroblasts. *Biochemical and Biophysical Research Communications*. 430(2), 579–584.
- Kok, H. T., Chia, F. C., & Alan, H. K. (2014). Impact of swiftlet's moult season on the value of edible bird nests. *International Proceedings of Chemical*, *Biological & Environmental Engineering*. 63, 17.
- Kwatra, B. (2020). Collagen Supplementation: Therapy For Skin Disorders: A Review Antimicrobial Discovery (Project Admis) View Project Collagen Supplementation: Therapy For Skin Disorders: A Review. Bharat et Al. World Journal of Pharmaceutical Research. 9, 2504.
- Kwatra, B., Solanki, A., Pal, M., Jasdanwala, S. S., & Pathak, T. (2021). Reviewing Effects of Rosehip, Curcumin, Piperine and Chondroitin Sulfate on Collagen. *International Journal of Pharmaceutical Sciences Review and Research*. 68(1).
- Larder, C. E., Iskandar, M. M., & Kubow, S. (2021). Assessment of Bioavailability after In Vitro Digestion and First Pass Metabolism of Bioactive Peptides from Collagen Hydrolysates. *Current Issues in Molecular Biology*. 43(3), 1592– 1605.
- Lee, T. H., Wani, W. A., Lee, C. H., Cheng, K. K., Shreaz, S., Wong, S., Hamdan, N., & Azmi, N. A. (2021). Edible Bird's Nest: The Functional Values of the Prized Animal-Based Bioproduct From Southeast Asia–A Review. *Frontiers in Pharmacology*. 12, 871.
- Lee, T.H., Lee, C.H., Azmi, N.A., Kavita, S., Wong, S., Znati, M., & Ben Jannet, H. (2020). Characterization of Polar and Non-Polar Compounds of House Edible Bird's Nest (EBN) from Johor, Malaysia. *Chemistry & Biodiversity*. 17(1).
- León-López, A., Pérez-Marroquín, X. A., Campos-Lozada, G., Campos-Montiel, R.
 G., & Aguirre-Álvarez, G. (2020). Characterization of Whey-Based
 Fermented Beverages Supplemented with Hydrolyzed Collagen: Antioxidant
 Activity and Bioavailability. *Foods*. 9(8).
- Lephart, E. D. (2018). Equol's Anti-Aging Effects Protect against Environmental Assaults by Increasing Skin Antioxidant Defense and ECM Proteins While Decreasing Oxidative Stress and Inflammation. *Cosmetics 2018*. 16.

Liaudanskas, M., Noreikienė, I., Zymonė, K., Juodytė, R., Žvikas, V., & Janulis, V.

(2021). Composition And Antioxidant Activity Of Phenolic Compounds In Fruit Of The Genus Rosa L. *Antioxidants*.10(4).

- Lin, P., Alexander, R. A., Liang, C. H., Liu, C., Lin, Y. H., Lin, Y. H., Chan, L. P., & Kuan, C. M. (2021). Collagen Formula With Djulis For Improvement Of Skin Hydration, Brightness, Texture, Crow's Feet, And Collagen Content: A Double-Blind, Randomized, Placebo-Controlled Trial. *Journal of Cosmetic Dermatology*. 188–194.
- Liu, H., Cao, J., & Jiang, W. (2015a). Evaluation Of Physiochemical And Antioxidant Activity Changes During Fruit On-Tree Ripening For The Potential Values Of Unripe Peaches. *Scientia Horticulturae*, *Complete*. 193, 32–39.
- Liu, H., Cao, J., & Jiang, W. (2015b). Evaluation And Comparison Of Vitamin C, Phenolic Compounds, Antioxidant Properties And Metal Chelating Activity Of Pulp And Peel From Selected Peach Cultivars. *LWT - Food Science and Technology*. 2(63), 1042–1048.
- Ma, F., & Liu, D. (2012). Sketch Of The Edible Bird's Nest And Its Important Bioactivities. *Food Research International*. 48(2), 559–567.
- Maehata, Y., Takamizawa, S., Ozawa, S., Izukuri, K., Kato, Y., Sato, S., Lee, M. C.
 il, Kimura, A., & Hata, R. I. (2007). Type III Collagen Is Essential For
 Growth Acceleration Of Human Osteoblastic Cells By Ascorbic Acid 2Phosphate, A Long-Acting Vitamin C Derivative. *Matrix Biology : Journal of the International Society for Matrix Biology*. 26(5), 371–381.
- Malo, C., & Wilson, J. X. (2000). Glucose Modulates Vitamin C Transport In Adult Human Small Intestinal Brush Border Membrane Vesicles. *The Journal of Nutrition*. 130(1), 63–69.
- Mardani, M., Yeganehzad, S., Ptichkina, N., Kodatsky, Y., Kliukina, O., Nepovinnykh, N., & Naji-Tabasi, S. (2019). Study On Foaming, Rheological And Thermal Properties Of Gelatin-Free Marshmallow. Undefined. 93, 335– 341.
- Mármol, I., Sánchez-De-Diego, C., Jiménez-Moreno, N., Ancín-Azpilicueta, C., & Rodríguez-Yoldi, M. (2017). Therapeutic Applications of Rose Hips from Different Rosa Species. *International Journal of Molecular Sciences*. 1137.
- Matthews, D. E. (2020). Review of Lysine Metabolism with a Focus on Humans. *The Journal of Nutrition*. 150

- May, J. M., & Qu, Z. C. (2005). Transport And Intracellular Accumulation Of Vitamin C In Endothelial Cells: Relevance To Collagen Synthesis. Archives of Biochemistry and Biophysics. 434(1), 178–186.
- Ménard, O., Cattenoz, T., Guillemin, H., Souchon, I., Deglaire, A., Dupont, D., & Picque, D. (2014). Validation of a new in vitro dynamic system to simulate infant digestion. *Food Chemistry*. 145, 1039–1045.
- Minekus, M., Alminger, M., Alvito, P., Ballance, S., Bohn, T., Bourlieu, C., Carrière, F., Boutrou, R., Corredig, M., Dupont, D., Dufour, C., Egger, L., Golding, M., Karakaya, S., Kirkhus, B., Le Feunteun, S., Lesmes, U., MacIerzanka, A., MacKie, A., Brodkorb, A. (2014). A Standardised Static In Vitro Digestion Method Suitable For Food An International Consensus. *Food & Function*. 1113–1124.
- Mokrani, A., Krisa, S., Cluzet, S., Da Costa, G., Temsamani, H., Renouf, E., Mérillon, J. M., Madani, K., Mesnil, M., Monvoisin, A., & Richard, T. (2016). Phenolic Contents And Bioactive Potential Of Peach Fruit Extracts. *Food Chemistry*. 202, 212–220.
- Neklyudov, A. D. (2003). Nutritive Fibers of Animal Origin: Collagen and Its Fractions as Essential Components of New and Useful Food Products. *Applied Biochemistry and Microbiology 2003 39:3*, 39(3), 229–238.
- Nicoletti, M. (2012). Nutraceuticals And Botanicals: *Overview And Perspectives*. 2–6.
- Njus, D., Kelley, P. M., Tu, Y. J., & Schlegel, H. B. (2020). Ascorbic Acid: The Chemistry Underlying Its Antioxidant Properties. *Free Radical Biology and Medicine*. 159, 37–43.
- Nurfatin, M. H., Etty Syarmila, I. K., Nur 'Aliah, D., Zalifah, M. K., Babji, A. S., & Ayob, M. K. (2016). Effect Of Enzymatic Hydrolysis On Angiotensin Converting Enzyme (ACE) Inhibitory Activity In Swiftlet Saliva. *International Food Research Journal*. 23(1), 141–146.
- Nuti, E., Casalini, F., Avramova, S. I., Santamaria, S., Cercignani, G., Marinelli, L., La Pietra, V., Novellino, E., Orlandini, E., Nencetti, S., Tuccinardi, T., Martinelli, A., Lim, N. H., Visse, R., Nagase, H., & Rossello, A. (2009). N-O-Isopropyl Sulfonamido-Based Hydroxamates: Design, Synthesis And Biological Evaluation Of Selective Matrix Metalloproteinase-13 Inhibitors As Potential Therapeutic Agents For Osteoarthritis. *Journal of Medicinal*

Chemistry. 52(15), 4757–4773.

- Pérez-Sánchez, A., Barrajón-Catalán, E., Herranz-López, M., & Micol, V. (2018). Nutraceuticals for Skin Care: A Comprehensive Review of Human Clinical Studies. *Nutrients 2018*. 10(4), 403.
- Quek, M. C., Chin, N. L., Tan, S. W., Yusof, Y. A., & Law, C. L. (2018). Molecular Identification Of Species And Production Origins Of Edible Bird's Nest Using FINS And SYBR Green I Based Real-Time PCR. *Food Control*. 118– 127.
- Rovná, K., Ivanišová, E., Žiarovská, J., Ferus, P., Terentjeva, M., Kowalczewski, P. Ł., & Kačániová, M. (2020). Characterization of Rosa canina Fruits Collected in Urban Areas of Slovakia. Genome Size, iPBS Profiles and Antioxidant and Antimicrobial Activities. *Molecules (Basel, Switzerland)*. 25(8).
- Saengkrajang, W., Matan, N., & Matan, N. (2013). Nutritional Composition Of The Farmed Edible Bird's Nest (Collocalia Fuciphaga) In Thailand. *Journal of Food Composition and Analysis*. 41–45.
- Seow, E. K., Ibrahim, B., Muhammad, S. A., Lee, L. H., & Cheng, L. H. (2016). Differentiation Between House And Cave Edible Bird's Nests By Chemometric Analysis Of Amino Acid Composition Data. LWT - Food Science and Technology, Complete. 428–435.
- Singh, M., Pande, S., & Battu, S. (2011). Medicinal Uses of L-Lysine: Past And Future. *Int. J. Res. Pharm. Sci.* 2(4), 637-642.
- Sionkowska, A., Adamiak, K., Musial, K., & Gadomska, M. (2020). Collagen Based Materials in Cosmetic Applications: *A Review Materials 2020*. 13(19), 4217.
- Skov, K., Oxfeldt, M., Thøgersen, R., Hansen, M., & Bertram, H. C. (2019). Enzymatic Hydrolysis of a Collagen Hydrolysate Enhances Postprandial Absorption Rate-A Randomized Controlled Trial. *Nutrients*. 11(5).
- Song, H., Tian, Q., & Li, B. (2020). Novel Hyp-Gly-Containing Antiplatelet Peptides From Collagen Hydrolysate After Simulated Gastrointestinal Digestion And Intestinal Absorption. *Food & Function*. 5553–5564.
- Sovrani, V., de Jesus, L. I., Simas-Tosin, F. F., Smiderle, F. R., & Iacomini, M. (2017). Structural Characterization And Rheological Properties Of A Gel-Like B-D-Glucan From Pholiota Nameko. *Carbohydrate Polymers*. 169, 1–8.
- Tan, H.Y., Babji, A. S., Lim, S. J., & Sarbini, S. R. (2021). A Systematic Review of Edible Swiftlet's Nest (ESN): Nutritional Bioactive Compounds, Health

Benefits As Functional Food, And Recent Development As Bioactive ESN Glycopeptide Hydrolysate. *Trends in Food Science & Technology*. 115, 117–132.

- Tan, K. H., Chia, F. C., & Alan, H. K. (2014). Impact of swiftlet's moult season on the value of edible bird nests. International Proceedings of Chemical, Biological & Environmental Engineering, 63, 17.
- Thulin, N. K., & Wang, T. T. (2018). The Role of Fc Gamma Receptors in Broad Protection against Influenza Viruses. *Vaccines* 2018. 36.
- Venkatesan, J., Anil, S., Kim, S. K., & Shim, M. S. (2017). Marine Fish Proteins and Peptides for Cosmeceuticals: A Review. *Marine Drugs*. 15(5).
- Wang, L., An, X., Xin, Z., Zhao, L., & Hu, Q. (2007). Isolation And Characterization Of Collagen From The Skin Of Deep-Sea Redfish (Sebastes Mentella). Journal of Food Science. 72(8).
- Wang, Lin, Wang, Q., Qian, J., Liang, Q., Wang, Z., Xu, J., He, S., & Ma, H. (2015). Bioavailability and Bioavailable Forms of Collagen after Oral Administration to Rats. *Journal of Agricultural and Food Chemistry*. 63(14), 3752–3756.
- Wickham, M., Faulks, R., & Mills, C. (2009). In Vitro Digestion Methods For Assessing The Effect Of Food Structure On Allergen Breakdown. *Molecular Nutrition & Food Research*. 952–958.
- Wong, C.-F., Chan, G. K.-L., Zhang, M.-L., Yao, P., Lin, H.Q., Dong, T. T.-X., Li, G., Lai, X.-P., Tsim, K. W.-K., Wong, C.-F., Chan, G. K.-L., Zhang, M.-L., Yao, P., Lin, H.-Q., Dong, T. T.-X., Li, G., Lai, X.-P., & Tsim, K. W.-K. (2017). Characterization Of Edible Bird's Nest By Peptide Fingerprinting With Principal Component Analysis. *Food Quality and Safety*, 83–92.
- Wong, F. C., Xiao, J., Wang, S., Ee, K. Y., & Chai, T. T. (2020). Advances On The Antioxidant Peptides From Edible Plant Sources. *Trends in Food Science and Technology*. 99, 44–57.
- Wong, Z. C. F., Chan, G. K. L., Wu, K. Q. Y., Poon, K. K. M., Chen, Y., Dong, T. T. X., & Tsim, K. W. K. (2018). Complete Digestion Of Edible Bird's Nest Releases Free N-Acetylneuraminic Acid And Small Peptides: An Efficient Method To Improve Functional Properties. *Food & Function*. 5139–5149.
- Woo, T., Lau, L., Cheng, N., Chan, P., Tan, K., & Gardner, A. (2017). Efficacy OfOral Collagen In Joint Pain-Osteoarthritis And Rheumatoid Arthritis. *Journal*

of Arthritis. 1-4.

Yang, Q., Zhao, D., & Liu, Q. (2020). Connections Between Amino Acid Metabolisms in Plants: Lysine as an Example. *Frontiers in Plant Science*. 11, 928.