



Editorial

Plants and Diabetes: Description, Role, Comprehension and Exploitation

Alessandra Durazzo ^{1,*} , Massimo Lucarini ^{1,*} and Antonello Santini ^{2,*}

¹ CREA-Research Centre for Food and Nutrition, via Ardeatina 546, 00178 Rome, Italy

² Department of Pharmacy, University of Napoli Federico II, via D. Montesano 49, 80131 Napoli, Italy

* Correspondence: alessandra.durazzo@crea.gov.it (A.D.); massimo.lucarini@crea.gov.it (M.L.); asantini@unina.it (A.S.)

Abstract: Many plants have been known for centuries to have medicinal importance with potential beneficial effects on health. Phytotherapeutic compounds are well known to play a globally significant role, in particular in the management and treatment of various chronic diseases. Among these, diabetes can cause long term damage to the body other than having a relevant economic burden on society being among the costliest chronic diseases. This motivated the focus of the proposed Special Issue, intended to develop and exploit the potential role of plants in the management and treatment of diabetes. The main topics included are: (i) description and use of medicinal plants for diabetes management; (ii) the elucidation and delineation of their main components, properties (anti-hyperglycaemic, hypoglycaemic, anti-inflammatory, apoptotic agents, etc.), (iii) the mechanism of action (in vitro and in vivo studies); (iv) formulation of nutraceuticals, botanicals, and dietary supplements useful as tools as an alternative or support to anti-diabetic pharmacological therapies; (v) development of new markers.

Keywords: diabetes; medicinal plants; nutraceuticals; mechanism of action; markers



Citation: Durazzo, A.; Lucarini, M.; Santini, A. Plants and Diabetes: Description, Role, Comprehension and Exploitation. *Int. J. Mol. Sci.* **2021**, *22*, 3938. <https://doi.org/10.3390/ijms22083938>

Received: 6 April 2021

Accepted: 7 April 2021

Published: 11 April 2021

Publisher's Note: MDPI stays neutral with regard to jurisdictional claims in published maps and institutional affiliations.



Copyright: © 2021 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (<https://creativecommons.org/licenses/by/4.0/>).

Introduction

Many plants have been known for a long time to have medicinal values with potential beneficial effects on health [1–8]. There is a great research interest in better understanding diabetes and finding better treatment options, including the identification of natural products with anti-diabetic effects [9–12].

It is worth mentioning the recent work of Yeung et al. [13] which investigates the natural products in diabetes based on quantitative literature analysis and giving the main directions of natural product research in diabetes up to now and useful hints on promising avenues for future research. Some highly cited natural products or compound classes including curcumin, flavanone, resveratrol, carotenoid, polyphenols, flavonol, flavone and berberine have been considered.

Diabetes is a lifelong condition that causes a person's blood sugar level to become too high. In recent years, diabetes prevalence increased in virtually all regions of the world, with millions of people worldwide now living with diabetes. This is a concern with deep effects on quality of life, demand on health services and economic costs. This health condition is related to a cascade of events which may include coronary heart disease, stroke and peripheral vascular disease, microvascular complications, renal disease, retinopathy and neuropathy, along with lower-extremity amputations [14].

With the growing rise in obesity and diabetes has come an increasing awareness of their impacts on infectious diseases, post-infection complications and mortality from critical infections implying also increased risk due to the coronavirus outbreak disease (COVID-19) that has become an evolving worldwide health crisis [15,16].

The focus of the proposed Special Issue is the potential role of plants in the management and treatment of diabetes. The main topics included are: (i) description and

use of medicinal plants for diabetes management; (ii) the elucidation and delineation of their main components properties (anti-hyperglycaemic, hypoglycaemic, anti-inflammatory, apoptotic agents, etc.); (iii) their mechanism of action (in vitro and in vivo studies); (iv) the formulation of nutraceuticals, botanicals and dietary supplements; (v) the develop and novel anti-diabetic drugs; (vi) the development of new markers for this health condition.

At the interface between food and medicine, Jeremic et al. [17] showed the potential effects on metabolic syndrome of diallyl trisulfide derived from garlic and the effect on myocardial function in rats affected by metabolic syndrome. The topic was previously studied by the same authors [18] for its positive effects on the cardiovascular system in rats with chemically induced Type 1 diabetes mellitus.

Lu et al. [19] showed how a novel dipeptidyl peptidase IV inhibitory tea peptide improves pancreatic β -cell function and reduces α -cell proliferation in streptozotocin-induced diabetic mice.

In the scenario of the re-use of agro-food waste in the perspective of biorefinery and circular economy [20,21], the work of Giacometti et al. [22] showed a beneficial role of phenolic compounds from the olive leaf in the regulation of glucose homeostasis in the skeletal muscle, by improving glucose translocation and influence glucose uptake as the treatment for streptozotocin (SZT)-induced diabetes in rats.

It is worth mentioning the reviews on: (i) the role of isoflavones in type 2 diabetes prevention and treatment [23]; (ii) the therapeutic potential of apigenin [24]; (iii) multiple cell signaling pathways of human proinsulin C-peptide in vasculopathy protection [25]; (iv) the role of algae-derived components on Type 2 diabetes mellitus prevention and treatment [26].

This Special Issue's end points have been to contribute to the growth of this area of research, trigger research interest on medicinal plants and diabetes and its implications, and reference their impact and use, by adding information scientifically substantiated with new data. We would like to thank all the Authors and the Reviewers of the papers published in this Special Issue for their great contributions and efforts. We are also grateful to the Editorial Board Members and to the Staff of the Journal for their kind support in the all the steps in the realization of this Special Issue.

Author Contributions: All authors listed (A.D., M.L., and A.S.) have made a substantial contribution to the work and approved it for publication. All authors have read and agreed to the published version of the manuscript.

Funding: This research received no external funding.

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

References

1. Santini, A.; Novellino, E. Nutraceuticals: Beyond the Diet before the Drugs. *Curr. Bioact. Compd.* **2014**, *10*, 1–12. [[CrossRef](#)]
2. Durazzo, A.; Lucarini, M.; Souto, E.B.; Cicala, C.; Caiazzo, E.; Izzo, A.A.; Novellino, E.; Santini, A. Polyphenols: A concise overview on the chemistry, occurrence, and human health. *Phytother. Res.* **2019**, *33*, 2221–2243. [[CrossRef](#)]
3. Durazzo, A. Study approach of antioxidant properties in foods: Update and considerations. *Foods* **2017**, *6*, 17. [[CrossRef](#)]
4. Santini, A. Nutraceuticals: Redefining a Concept. *Ann. Pharmacol. Pharm.* **2018**, *3*, 1147.
5. Santini, A.; Novellino, E. To Nutraceuticals and Back: Rethinking a Concept. *Foods* **2017**, *6*, 74. [[CrossRef](#)] [[PubMed](#)]
6. Daliu, P.; Santini, A.; Novellino, E. A decade of nutraceutical patents: Where are we now in 2018? *Expert Opin. Ther. Pat.* **2018**, *28*, 875–882. [[CrossRef](#)] [[PubMed](#)]
7. Santini, A.; Novellino, E. Nutraceuticals—shedding light on the grey area between pharmaceuticals and food. *Expert Rev. Clin. Pharmacol.* **2018**, *11*, 545–547. [[CrossRef](#)]
8. Daliu, P.; Santini, A.; Novellino, E. From pharmaceuticals to nutraceuticals: Bridging disease prevention and management. *Expert Rev. Clin. Pharmacol.* **2019**, *12*, 1–7. [[CrossRef](#)] [[PubMed](#)]
9. Salehi, B.; Ata, A.; Anil Kumar, N.V.; Sharopov, F.; Ramirez-Alarcon, K.; Ruiz-Ortega, A.; Abdulmajid Ayatollahi, S.; Valere Tsouh Fokou, P.; Kobarfard, F.; Amiruddin Zakaria, Z.; et al. Antidiabetic potential of medicinal plants and their active components. *Biomolecules* **2019**, *9*, 551. [[CrossRef](#)]

10. Vieira, R.; Souto, S.B.; Sanchez-Lopez, E.; Lopez Machado, A.; Severino, P.; Jose, S.; Santini, A.; Fortuna, A.; Garcia, M.L.; Silva, A.M. Sugar-lowering drugs for type 2 diabetes mellitus and metabolic syndrome—Review of classical and new compounds: Part-I. *Pharmaceuticals* **2019**, *12*, 152. [[CrossRef](#)]
11. Vieira, R.; Souto, S.B.; Sanchez-Lopez, E.; Lopez Machado, A.; Severino, P.; Jose, S.; Santini, A.; Silva, A.M.; Fortuna, A.; Garcia, M.L. Sugar-lowering drugs for type 2 diabetes mellitus and metabolic syndrome—Strategies for in vivo administration: Part-II. *J. Clin. Med.* **2019**, *8*, 1332. [[CrossRef](#)] [[PubMed](#)]
12. Souto, E.B.; Souto, S.B.; Campos, J.R.; Severino, P.; Pashirova, T.N.; Zakharova, L.Y.; Silva, A.M.; Durazzo, A.; Lucarini, M.; Izzo, A.A.; et al. Nanoparticle delivery systems in the treatment of diabetes complications. *Molecules* **2019**, *24*, 4209. [[CrossRef](#)] [[PubMed](#)]
13. Yeung, A.W.K.; Tzvetkov, N.T.; Durazzo, A.; Lucarini, M.; Souto, E.B.; Santini, A.; Gan, R.; Jozwik, A.; Grzybek, W.; Horbańczuk, J.O.; et al. Natural products in diabetes research: Quantitative literature analysis. *Nat. Prod. Res.* **2020**, 1–15. [[CrossRef](#)]
14. Harding, J.L.; Pavkov, M.E.; Magliano, D.J.; Shaw, J.E.; Gregg, E.W. Global trends in diabetes complications: A review of current evidence. *Diabetologia* **2019**, *62*, 3–16. [[CrossRef](#)]
15. Rubino, F.; Amiel, S.A.; Zimmet, P.; Alberti, G.; Bornstein, S.; Eckel, R.H.; Mingrone, G.; Boehm, B.; Cooper, M.E.; Chai, Z.; et al. New-Onset Diabetes in Covid-19. *N. Engl. J. Med.* **2020**, *383*, 789–790. [[CrossRef](#)]
16. Bloomgarden, Z.T. Diabetes and COVID-19. *J. Diabetes* **2020**, *12*, 347–349. [[CrossRef](#)]
17. Jeremic, J.N.; Jakovljevic, V.L.; Zivkovic, V.I.; Srejovic, I.M.; Bradic, J.V.; Milosavljevic, I.M.; Mitrovic, S.L.; Jovicic, N.U.; Bolevich, S.B.; Svistunov, A.A.; et al. Garlic Derived Diallyl Trisulfide in Experimental Metabolic Syndrome: Metabolic Effects and Cardioprotective Role. *Int. J. Mol. Sci.* **2020**, *21*, 9100. [[CrossRef](#)]
18. Jeremic, J.N.; Jakovljevic, V.L.; Zivkovic, V.I.; Srejovic, I.M.; Bradic, J.V.; Bolevich, S.; Nikolic Turnic, T.R.; Mitrovic, S.L.; Jovicic, N.U.; Tyagi, S.C.; et al. The cardioprotective effects of diallyl trisulfide on diabetic rats with ex vivo induced ischemia/reperfusion injury. *Mol. Cell Biochem.* **2019**, *460*, 151–164. [[CrossRef](#)]
19. Lu, Y.; Lu, P.; Wang, Y.; Fang, X.; Wu, J.; Wang, X. A Novel Dipeptidyl Peptidase IV Inhibitory Tea Peptide Improves Pancreatic β -Cell Function and Reduces α -Cell Proliferation in Streptozotocin-Induced Diabetic Mice. *Int. J. Mol. Sci.* **2019**, *20*, 322. [[CrossRef](#)] [[PubMed](#)]
20. Lucarini, M.; Durazzo, A.; Romani, A.; Campo, M.; Lombardi-Boccia, G.; Cecchini, F. Bio-Based Compounds from Grape Seeds: A Biorefinery Approach. *Molecules* **2018**, *23*, 1888. [[CrossRef](#)]
21. Durazzo, A.; D'Addezio, L.; Camilli, E.; Piccinelli, R.; Turrini, A.; Marletta, L.; Marconi, S.; Lucarini, M.; Lisciani, S.; Gabrielli, P.; et al. From Plant Compounds to Botanicals and Back: A Current Snapshot. *Molecules* **2018**, *23*, 1844. [[CrossRef](#)] [[PubMed](#)]
22. Giacometti, J.; Muhvić, D.; Grubić-Kezele, T.; Nikolić, M.; Šoić-Vranić, T.; Bajek, S. Olive Leaf Polyphenols (OLPs) Stimulate GLUT4 Expression and Translocation in the Skeletal Muscle of Diabetic Rats. *Int. J. Mol. Sci.* **2020**, *21*, 8981. [[CrossRef](#)]
23. Kuryłowicz, A. The Role of Isoflavones in Type 2 Diabetes Prevention and Treatment—A Narrative Review. *Int. J. Mol. Sci.* **2021**, *22*, 218. [[CrossRef](#)]
24. Salehi, B.; Venditti, A.; Sharifi-Rad, M.; Kręgiel, D.; Sharifi-Rad, J.; Durazzo, A.; Lucarini, M.; Santini, A.; Souto, E.B.; Novellino, E.; et al. The Therapeutic Potential of Apigenin. *Int. J. Mol. Sci.* **2019**, *20*, 1305. [[CrossRef](#)]
25. Souto, S.B.; Campos, J.R.; Fangueiro, J.F.; Silva, A.M.; Cicero, N.; Lucarini, M.; Durazzo, A.; Santini, A.; Souto, E.B. Multiple Cell Signalling Pathways of Human Proinsulin C-Peptide in Vasculopathy Protection. *Int. J. Mol. Sci.* **2020**, *21*, 645. [[CrossRef](#)] [[PubMed](#)]
26. Bocanegra, A.; Macho-González, A.; Garcimartín, A.; Benedí, J.; Sánchez-Muniz, F.J. Whole Alga, Algal Extracts, and Compounds as Ingredients of Functional Foods: Composition and Action Mechanism Relationships in the Prevention and Treatment of Type-2 Diabetes Mellitus. *Int. J. Mol. Sci.* **2021**, *22*, 3816. [[CrossRef](#)]