



OPEN ACCESS

EDITED BY
Javier Echeverria,
University of Santiago, Chile

REVIEWED BY
Luca Rastrelli,
University of Salerno, Italy

*CORRESPONDENCE
Prabhakar Semwal,
✉ semwal.prabhakar@gmail.com,
✉ prabhakarsemwal.ls@geu.ac.in

RECEIVED 11 July 2023
ACCEPTED 07 August 2023
PUBLISHED 11 August 2023

CITATION
Semwal P, Rauf A and Simal-Gandara J
(2023), Editorial: High altitude medicinal
plants and their bioactive compounds for
the prevention of oxidative stress-
induced diseases and disorders.
Front. Pharmacol. 14:1257018.
doi: 10.3389/fphar.2023.1257018

COPYRIGHT
© 2023 Semwal, Rauf and Simal-
Gandara. This is an open-access article
distributed under the terms of the
[Creative Commons Attribution License
\(CC BY\)](https://creativecommons.org/licenses/by/4.0/). The use, distribution or
reproduction in other forums is
permitted, provided the original author(s)
and the copyright owner(s) are credited
and that the original publication in this
journal is cited, in accordance with
accepted academic practice. No use,
distribution or reproduction is permitted
which does not comply with these terms.

Editorial: High altitude medicinal plants and their bioactive compounds for the prevention of oxidative stress-induced diseases and disorders

Prabhakar Semwal^{1,2*}, Abdur Rauf³ and Jesus Simal-Gandara⁴

¹Department of Biotechnology, Graphic Era (Deemed to be University), Dehradun, India, ²Research and Development Cell, Graphic Era Hill University, Dehradun, India, ³Department of Chemistry, University of Swabi, Khyber Pakhtunkhwa, Pakistan, ⁴Analytical Chemistry and Food Science Department, Faculty of Science, Universidade de Vigo, Nutrition and Bromatology Group, Vigo, Spain

KEYWORDS

high altitude, Himalaya, bioactive compounds, health promoting effects, oxidative stress, biological applications

Editorial on the Research Topic

High altitude medicinal plants and their bioactive compounds for the prevention of oxidative stress-induced diseases and disorders

Oxidative stress is a biochemical process occurring in cells and tissues due to the abnormal accumulation and production of reactive oxygen species (ROS)/free radicals, in which the endogenous antioxidants are unable to neutralize these free radicals and cause oxidative damage to proteins, lipids, DNA, etc. of the cells (Pizzino et al., 2017; Sharifi-Rad et al., 2020). If not managed properly it can accelerate the aging process and can be responsible for causing acute pathologies like trauma, stroke, etc., as well as many chronic and degenerative diseases and disorders (Vona et al., 2021).

Bioactive compounds are extra-nutritional constituents, generally found in different types of foods, fruits, vegetables, and grains, which provide health benefits beyond their basic and traditional nutritional value (Zhao et al., 2015). A large number of bioactive constituents have been isolated from botanicals, and these botanicals have been utilized from antiquity for their therapeutic abilities.

High-altitude medicinal plants are known for their ethnobotanical importance and have been used by various communities around the world for the treatment of various diseases and disorders due to their healing properties. The diverse geographical and climatic condition including biotic and abiotic factors leads to the production of novel bioactive metabolites in these plants (Heinrich et al., 2021; Semwal et al., 2022). In the past few years, many potential metabolites such as phenolic acids, alkaloids, flavonoids, volatile oils, glycosides, etc. have been discovered from natural sources for their antimicrobial, cardioprotective, anti-inflammatory, anticancer and chemoprotective activities. There is considerable evidence to suggest that these phytochemicals are beneficial to health on an epidemiological and clinical level (Chan et al., 2023; Du et al., 2023; Rahaman et al., 2023; Xu et al., 2023; Zhou et al., 2023; Zhu et al., 2023).

The present Research Topic aims to provide a platform for current research and evidences available about the role of High-altitude medicinal plants, their chemical components, and its role in reducing the risk of diseases and disorders related to oxidative stress. Within this dedicated edition, the guest editorial board was overwhelmed to receive a remarkable series of scientific reports from numerous regions around the world. In all 16 original research and review articles were submitted and after the critical evaluation and peer review process total 6 articles (02 Original research article, 01 Systematic review and 03 Review articles) have been approved for the publication in this special edition.

Limonin, a triterpenoid molecule that is typically present in citrus fruits, exhibits a diverse range of pharmacological properties, including antioxidant, antiviral, anti-inflammatory, anti-cancer, and liver protecting qualities. The first research article by Li et al. describes the protective effects of limonin compound and their molecular mechanism against non-alcoholic fatty liver disease by means of *in-vitro* and *in-vivo* models. In this investigation, zebrafish larvae were treated with thioacetamide to create a model of NAFLD, and the larvae received limonin treatment parallelly for 72 h. According to the findings, limonin dramatically decreased the formation of lipid bodies in liver and downregulated the levels of sterol regulatory element binding protein 1 (SREBP-1), two lipogenic transcription factors associated with NAFLD and fatty acid synthase. The study also showed that limonin inhibited the invasion of macrophages and reduced the expression levels of the pro-inflammatory mediators including IL-1beta, IL-6 and TNF-alpha, which are released by macrophages. Furthermore, limonene can reverse glutathione depletion and reactive oxygen species accumulation by regulating the NRF2/HO-1 signalling pathway in the liver. In conclusion, this study revealed that limonin has pharmacological effects in the treatment of non-alcoholic fatty liver disease by reducing oxidative stress, lipid accumulation, and inflammation induced by pro-inflammatory chemokines and macrophage infiltration Li et al.

Paeonol, a phenolic compound present in various plant species, is known to have several pharmacological activities, including antitumor, anti-inflammatory, neuroprotective, cardioprotective, nephroprotective activities. The second research article demonstrated the *in-vivo* anti-diabetic activity of paeonol against diabetic retinopathy. In this study, diabetes was induced using streptozotocin (55 mg/kg, i.p.) in male Sprague Dawley rats, and diabetic rats were administered with paeonol at doses between 50 and 200 mg/kg per day for the next 4 weeks after the initial 4-week treatment period. Different parameters such as retinal physiology, histopathology, biochemical and oxidative stress were recorded. The electroretinogram recording of paeonol-treated rats exhibited a substantial improvement in the a-wave amplitude, b-wave amplitude, a-wave latency, and b-wave latency ($p < 0.001$) at 15 cd s/m² when compared with control (diabetic animals). Additionally, paeonol treated animals exhibited a substantial drop in the plasma glucose level, and aldose reductase, sorbitol dehydrogenase and lactate dehydrogenase level compared to diabetic control animals. According to the study's findings, paeonol can be used as a management strategy for diabetic retinopathy (Adki and Kulkarni).

Worldwide, cardiovascular diseases (CVD) are among the leading causes of death. In a systematic review, Zeng et al.

reported the cardioprotective effects of curcumin on myocardial ischemia/reperfusion injury using preclinical meta-analysis in animal studies. This meta-analysis comprised 37 studies with a total of 771 animals, with technique quality values ranging from 4 to 7. The study suggests that curcumin treatment significantly reduced the extent of myocardial infarction, and also decreased the serum inflammatory cytokines and the myocardial apoptotic index Zeng et al.

The first review report on protective effects of salidroside's against different ischemia-related diseases with its possible molecular mechanism has been reported by Han et al. Salidroside, one of the main active components of *Rhodiola* species has been suggested to treat ischemia and ischemic damage by increasing the rate of cell survival and angiogenesis and reducing oxidative stress and inflammation. This article describes recent advances and research progress on salidroside for the treatment of ischemic diseases, including ischemic heart disease, ischemic acute kidney injury, cerebral ischemia, liver ischemia, etc., Han et al.

Gynaecological malignancies, such as ovarian, cervical, and endometrial cancers, have had a significant negative impact on women's health due to concealed illnesses, incorrect diagnosis, and high recurrence rates. Li et al. have investigated the molecular mechanisms of the beneficial effects of natural polysaccharides in treating gynecological cancer. In this review article, the role of natural polysaccharides, preparation of new dosages, available scientific evidences have been included Li et al.

Fordjour et al. explore the therapeutic potential of *Cannabis sativa*. In this articles, scientific information has been collected on different aspects such as ecology, chemical compounds, ethnomedicinal applications, biological applications, industrial applications and toxicological studies.

Overall, taken together the data available in this special edition clearly indicated the significant pharmacological effects of naturally-occurring bioactive constituents in multiple affections, with promissory data being increasingly published. However, despite these advances, it is still necessary to clarify the effects of single and multiple doses of drugs, pharmacokinetics, efficacy, their toxicity, pharmacodynamics, and safety profiles, as well as their mechanism of action, for a clearer understanding of their therapeutic properties and to establish stronger evidence-based medicine.

Author contributions

PS: Conceptualization, Investigation, Writing—original draft, Writing—review and editing. AR: Writing—review and editing. JS-G: Writing—review and editing.

Acknowledgments

The editors would like express their gratitude to the authors and reviewers who contributed to and participated in this Research Topic for their efforts, timely responses, and passion. We also thank the Frontiers Editorial Office for their assistance and support.

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

The author(s) declared that they were an editorial board member of Frontiers, at the time of submission. This had no impact on the peer review process and the final decision.

References

- Chan, W.-J. J., Adiwidjaja, J., McLachlan, A. J., Boddy, A. V., and Harnett, J. E. (2023). Interactions between natural products and cancer treatments: underlying mechanisms and clinical importance. *Cancer Chemother. Pharmacol.* 91 (2), 103–119. doi:10.1007/s00280-023-04504-z
- Du, Y.-X., Mamun, A. A., Lyu, A.-P., and Zhang, H.-J. (2023). Natural compounds targeting the autophagy pathway in the treatment of colorectal cancer. *Int. J. Mol. Sci.* 24 (8), 7310. doi:10.3390/ijms24087310
- Heinrich, M., Jiang, H., Scotti, F., Booker, A., Walt, H., Weckerle, C., et al. (2021). Medicinal plants from the Himalayan region for potential novel antimicrobial and anti-inflammatory skin treatments. *J. Pharm. Pharmacol.* 73 (7), 956–967. doi:10.1093/jpp/rgab039
- Pizzino, G., Irrera, N., Cucinotta, M., Pallio, G., Mannino, F., Arcoraci, V., et al. (2017). Oxidative stress: harms and benefits for human health. *Oxid. Med. Cell Longev.* 2017, 8416763, doi:10.1155/2017/8416763
- Rahaman, M. M., Hossain, R., Herrera-Bravo, J., Islam, M. T., Atolani, O., Adeyemi, O. S., et al. (2023). Natural antioxidants from some fruits, seeds, foods, natural products, and associated health benefits: an update. *Food Sci. Nutr.* 11 (4), 1657–1670. doi:10.1002/fsn3.3217
- Semwal, P., Painuli, S., Jamloki, A., Rauf, A., Rahman, M. M., Olatunde, A., et al. (2022). Himalayan wild fruits as a strong source of nutraceuticals, therapeutics, food and nutrition security. *Food Rev. Int.*, 1–37. doi:10.1080/87559129.2022.2121407
- Sharifi-Rad, M., Anil Kumar, N. V., Zucca, P., Varoni, E. M., Dini, L., Panzarini, E., et al. (2020). Lifestyle, oxidative stress, and antioxidants: back and forth in the pathophysiology of chronic diseases. *Front. Physiol.* 11, 694. doi:10.3389/fphys.2020.00694
- Vona, R., Pallotta, L., Cappelletti, M., Severi, C., and Matarrese, P. (2021). The impact of oxidative stress in human pathology: focus on gastrointestinal disorders. *Antioxidants (Basel)* 10 (2), 201. doi:10.3390/antiox10020201
- Xu, X., Han, C., Wang, P., and Zhou, F. (2023). Natural products targeting cellular processes common in Parkinson's disease and multiple sclerosis. *Front. Neurology* 14, 1149963. doi:10.3389/fneur.2023.1149963
- Zhao, Y., Wu, Y., and Wang, M. (2015). *Bioactive substances of plant origin. Handbook of food chemistry* (Berlin, Heidelberg: Springer Berlin Heidelberg). P. C. K. Cheung, and B. M. Mehta. 967–1008.
- Zhou, Y., Wang, D., and Yan, W. (2023). Treatment effects of natural products on inflammatory bowel disease *in vivo* and their mechanisms: based on animal experiments. *Nutrients* 15 (4), 1031. doi:10.3390/nu15041031
- Zhu, L.-R., Li, S.-S., Zheng, W.-Q., Ni, W.-J., Cai, M., and Liu, H.-P. (2023). Targeted modulation of gut microbiota by traditional Chinese medicine and natural products for liver disease therapy. *Front. Immunol.* 14, 1086078. doi:10.3389/fimmu.2023.1086078

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.