

THE THERAPEUTIC PROPERTIES OF MEDICINAL PLANTS

Health-Rejuvenating Bioactive
Compounds of Native Flora



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Innovations in Plant Science for Better Health: From Soil to Fork

THE THERAPEUTIC PROPERTIES OF MEDICINAL PLANTS

Health-Rejuvenating Bioactive
Compounds of Native Flora

Edited by

Megh R. Goyal

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Apple Academic Press Inc.
4164 Lakeshore Road
Burlington ON L7L 1A4, Canada

Apple Academic Press Inc.
1265 Goldenrod Circle NE
Palm Bay, Florida 32905, USA

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Exclusive worldwide distribution by CRC Press, a member of Taylor & Francis Group

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International Standard Book Number-13: 978-1-77188-803-5 (Hardcover)

International Standard Book Number-13: 978-0-42926-520-4 (eBook)

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Library and Archives Canada Cataloguing in Publication

Title: The therapeutic properties of medicinal plants : health-rejuvenating bioactive compounds of native flora/edited by Megh R. Goyal, Hafiz Ansar Rasul Suleria, Ademola Olabode Ayeleso, T. Jesse Joel, Sujogya Kumar Panda.

Names: Goyal, Megh Raj, editor. | Suleria, Hafiz, editor. | Ayeleso, Ademola Olabode, editor. | Joel, T. Jesse, editor. | Panda, Sujogya Kumar, editor.

Series: Innovations in plant science for better health.

Description: Series statement: Innovations in plant science for better health: from soil to fork | Includes bibliographical references and index.

Identifiers: Canadiana (print) 20190169990 | Canadiana (ebook) 20190170018 | ISBN 9781771888035 (hardcover) | ISBN 9780429265204 (ebook)

Subjects: LCSH: Materia medica, Vegetable. | LCSH: Medicinal plants—Research. | LCSH: Botanical chemistry—Research.

Classification: LCC RS164 .T54 2020 | DDC 615.3/21--dc23

Library of Congress Cataloging-in-Publication Data

Names: Goyal, Megh Raj, editor. | Suleria, Hafiz, editor. | Ayeleso, Ademola Olabode, editor. | Joel, T. Jesse, editor. | Panda, Sujogya Kumar, editor.

Title: The therapeutic properties of medicinal plants : health-rejuvenating bioactive compounds of native flora/edited by Megh R. Goyal, Hafiz Ansar Rasul Suleria, Ademola Olabode Ayeleso, T. Jesse Joel, Sujogya Kumar Panda.

Other titles: Innovations in plant science for better health

Description: Oakville, ON : Palm Bay, Florida : Apple Academic Press, [2020] | Series: Innovations in plant science for better health: from soil to fork | Includes bibliographical references and index. | Summary: "This volume, The Therapeutic Properties of Medicinal Plants, provides some informative research on the scientific evidence of the health benefits that can be derived from medicinal plants and how their efficacies can be improved. The volume is divided into three sections covering the phytochemistry of medicinal plants, disease management with medicinal plants, and novel research techniques in medicinal plants. The pharmacological benefits of several specific plants, such as basil, fig, garlic, palm tree, etc., are discussed, addressing health issues including metabolic and mental disorders, acute mountain sickness, polycystic ovarian syndrome, and specific diseases such as Huntington's. It also looks at the role of antioxidants in disease management. Additionally, the book covers recent problems of drug resistance and how medicinal plants can serve as antibiotic, anthelmintic, and antiparasitic drugs that will be helpful for human and animals. Furthermore, it also covers novel approaches for the screening of plant-based medicines, extraction, toxicity and safety issues of essential oils, and nanoparticle-based delivery of plant metabolites. The findings reported in this book will be useful in health policy decisions and will help to motivate the development of new health care products from plants for health benefits. The book will further encourage the preservation of traditional medical knowledge of plants with therapeutic qualities. It will be a valuable reference for researchers, scientists, students, growers, traders, processors, industry professionals, dieticians, medical practitioners, and others"-- Provided by publisher.

Identifiers: LCCN 2019036017 (print) | LCCN 2019036018 (ebook) | ISBN 9781771888035 (hardcover) | ISBN 9780429265204 (ebook)

Subjects: LCSH: Materia medica, Vegetable. | Medicinal plants--Research. | Botanical chemistry--Research.

Classification: LCC RS164 .T522 2019 (print) | LCC RS164 (ebook) | DDC 615.3/21--dc23

LC record available at <https://lcn.loc.gov/2019036017>

LC ebook record available at <https://lcn.loc.gov/2019036018>

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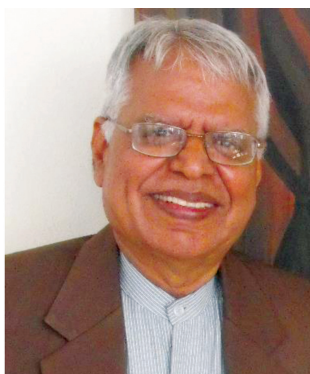


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ABBREVIATIONS

2-AAF	2-acetylaminofluorene
ABC	ATP binding cassette
ABPP	activity-based probe profiling
ABTS	2,2'-azino-bis (3-ethylbenzothiazoline-6-sulphonic acid)
ACE	angiotensin converting enzymes
AD	Alzheimer's disease
ADH	antidiuretic hormone
AFB1	aflatoxin B1
AGEs	advanced glycation end-products
AIDS	acquired immune deficiency syndrome
ALAD	δ -aminolevulinic acid dehydratase
AlCl ₃	aluminum chloride
ALP	alkaline phosphatase
ALT	alanine aminotransferase
ALT	aspartate aminotransferase
AMI	age-induced memory impairment
AMP	adenosine monophosphate
AMS	acute mountain sickness
ANN	artificial neural network
ANP	atrial natriuretic peptide
ARE	antioxidant response element
ASOs	antisense oligonucleotides
ATBC	alpha-tocopherol beta-carotene
ATP	adenosine triphosphate
AVP	average precision
BaP	benzo[a]pyrene
BBB	blood-brain-barrier
BCA	bicinchoninic acid
BDNF	brain-derived neurotrophic factor
BHT	butylated hydroxytoluene
bZIP	basic-leucine zippe
C2C12	immortalized mouse myoblast cell line
CAGE	cap analysis of gene expression
CAM	complementary and alternative medicines

CAMD	compacted air microwave distillation
CAMs	cell adhesion molecules
CAT	catalase
CCCP	compound centric chemical proteomics
CDRI	Central Drug Research Institute
CLMT	Chaihu-Jia-Longgu-Muli Tan
CNC	Cap'n'Collar
CNS	central nervous system
COMT	catechol-O-methyl transferase
COX-2	cyclooxygenase 2
CP	crude protein
CUS	chronic unpredictable stress
CV	computer vision
CVD	cardiovascular disease
CVS	chronic variable stress
CYP1A2	cytochrome P450 1A2 enzyme
DDS	drug delivery system
DETAPAC	diethylenetriaminepentaacetic acid
DMAD	dimethylallyl diphosphate
DMBA	7,12-dimethylbenz(a)anthracene
DMSO	dimethyl sulfoxide
DNA	deoxyribonucleic acid
DPPH	diphenyl picrylhydrazyl
DSS	dextran sodium sulfate
DUFARMS	director of university farms
EDTA	ethylenediaminetetraacetic acid
EGCG	epigallocatechin gallate
EOs	essential oils
EPG	egg count per gram
ERE	electrophile response element
ERK	extracellular regulated kinases
ESBL	extended spectrum beta lactamase
ESC	embryonic stem cell
EST	embryonic stem cell test
FAB-MS	fast atom bombardment mass spectrometry
FAO	Food and Agriculture Organization
FB ₁	fumonisin B ₁
FDA	Food and Drug Administration
FEC	faecal egg count
FeSO ₄	ferrous sulfate

FLBP	fuzzy local binary patterns
FRAP	ferric reducing antioxidant power
FSH	follicle stimulating hormone
FST	forced swim test
G1 phase	growth 1 phase
GABA	gamma amino butyric acid
GAD	generalized anxiety disorders
GAE	gallic acid equivalent
GC	gas chromatography
GCLC	glutamate cysteine ligase catalytic subunit
GDNF	glial cell derived neurotrophic factor
GGT	γ -glutamyl transferase
GIT	gastrointestinal tract
GLCM	gray level co-occurrence matrix
GLUT	glucose transporter
GPCRs	G-protein coupled receptors
GPx	glutathione peroxidase
GR	glutathione reductase
GUI	graphical user interface
H ₂ O ₂	hydrogen peroxide
HbAA	hemoglobin A
HbSS	hemoglobin S
HBV	hepatitis B virus
HCl	hydrochloric acid
HD	Huntington's disease
HD	hydrodistillation
HDAC	histone deacetylases
HDL-C	high-density lipoprotein cholesterol
HDL	high-density lipoprotein
HHA axis	hypothalamus-adrenocortical axis
HHT	homoharringtonine
HIV	human immunodeficiency virus
HMF	hydroxymethylfurfurals
HPA	hypothalamus-pituitary-adrenocortical
HPBMDM	human peripheral blood monocyte derived macrophages
HPLC	high performance liquid chromatography
HPV	human papilloma virus
HS-CRP	high sensitive C reactive protein
HUVEC	human umbilical vein endothelial cells
IC	inhibition concentration

IC ₅₀	50% median inhibition concentration
ICT	information communication technologies
IgM	immunoglobulin M
IL	interleukin
IL-10	interleukin 10
IL-6	interleukin 6
iNOS	inducible nitric oxide synthase
IPD	isopentenyl diphosphate
IPP	isopentenyl pyrophosphate
ISO	International Standard Organization
IVF	<i>in vitro</i> fertilization
kDa	Kilo Dalton
KH ₂ PO ₄	potassium phosphate
KV	kolaviron
LBP	local binary pattern
LBPV	local binary pattern variance
LC	liquid chromatography
LDH	lactate dehydrogenase
LDL-C	low density lipoprotein cholesterol
LH	luteinizing hormone
LLS	Lake Louise Score
LPO	lipid peroxidation
LPS	lipopolysaccharide
MAE	microwave-assisted extraction
MATLAB	matrix laboratory
MCV	mean cell volume
MDA	malondialdehyde
MDG	microwave dry-diffusion and gravity
MDR	multidrug resistant
MDR1	multi drug resistance gene-1
MedLeaf	medicinal leaf
mg/KgBW	milligram per kilogram body weight
MGDG	monogalactosyldiacyl glycerol
MHG	microwave hydrodiffusion and gravity
MIC	minimum inhibitory concentration
MILDA	microplate luminescence automated digital analyzer
MMC	mitomycin C
MNRET	micronucleated reticulocyte
MPA	metaphosphoric acid
MTT	3-(4,5-dimethylthiazol-2-yl)-2,5-diphenyltetrazolium bromide

MudPIT	multidimensional protein identification technology
MUFAs	mono-unsaturated fatty acids
MySQL	my structured query language
NAC	N-acetylcysteine
NAD	nicotinamide adenine dinucleotide
NAD(P)H	nicotinamide adenine dinucleotide phosphate
NADH	nicotinamide adenine dehydrogenase
NADP(H)	nicotinamide adenine dinucleotide phosphate
NaOH	sodium hydroxide
<i>NB</i>	Naivy Bayes
NCCIH	National Center for Complementary and Integrated Health
NCI	National Cancer Institute
NEMPS	nuclear-encoded mitochondrial proteins
NEUT	neutrophils
NF- $\kappa\beta$	nuclear factor- $\kappa\beta$
NIRS	near infrared spectroscopy
NMR	nuclear magnetic resonance
NO	nitric oxide
NPs	nanoparticles
NRF	nuclear receptor factor
O-PA	orthophosphoric acid
OPG	oocyst per gram
ORAC	oxygen radical absorbance capacity
OSRC	Oxidative Stress Research Centre
PCA	principal component analysis
PCOS	polycystic ovarian syndrome
PD	Parkinson Disease
PDR	pan drug resistant
PDR	product decision rule
PGC-1	peroxisome proliferator-activated receptor gamma coactivator-1
PKA	protein kinase A
PLE	pressurized liquid extraction
PLGA	poly(lactic-co-glycolic) acid
PNN	probabilistic neural network
PNPs	polymeric nanoparticles
PPI	prepulse inhibition
PQQ	pyroloquinoline quinone
Prx-1	peroxiredoxin-1
RAAS	renin-angiotensin-aldosterone system

RAGE	receptor the advanced glycation endproducts
<i>RF</i>	random forest
RIN-5F	rat islet tumor cell line
RNA	ribonucleic acid
RNAi	RNA interference
ROS	reactive oxygen species
RPMI	Roswell Park Memorial Institute
S phase	synthesis phase
SAGE	serial analysis of gene expression
SCA	sickle cell anemia
SCD	sickle cell disease
SCF	supercritical fluid technology
SCT	sickle cell trait
SD	sprague dawley
SDH	succinate dehydrogenase
SFE	supercritical fluid extraction
SFME	solvent-free microwave extraction
<i>SL</i>	simple <i>logistic</i>
SLNs	solid lipid nanoparticles
SMEDDS	self-microemulsifying drug delivery system
SOD	superoxide dismutase
SRC	standard rat chow
SSI	Sense of Smell Institute
SSRIs	selective serotonin reuptake inhibitors
STAT3	signal transduction as well as transcription 3
STZ	streptozotocin
SVMRFE	support vector machine recursive feature elimination
TBA	thiobarbituric acid
TBAR	thiobarbituric acid reactive
TBARS	thiobarbituric acid reactive substances
t-BHP	tertiary-butyl hydroperoxide
TC	total cholesterol
TFA	trifluoroacetic acid
Tfam	mitochondrial transcription factor A
TG	triglycerides
THPs	traditional health practitioners
TLC	thin layer chromatography
TNF- α	tumor necrosis factor- α
TPA	12-O-tetradecanoyl phorbol-13-acetate
TRAP	free radical trapping abilities

Txn-1	thioredoxin-1
UAE	ultrasound-assisted extraction
URF	University Research Fund
VEGF	vascular endothelial growth factor
VMHD	vacuum microwave hydro-distillation
VOCs	volatile organic compounds
WAD	West African Dwarf
WBC	white blood cells
WHCO5	human oesophageal cancer cells
WHO	World Health Organization
XDR	extensive drug resistant
YGS	Yi-gan san



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SYMBOLS

μM	micromole
$\alpha\text{-T}$	alpha-tocopherol
$\beta\text{-T}$	beta-tocopherol
$\gamma\text{-GT}$	γ -glutamyl transferase
$\gamma\text{-T}$	gamma-tocopherol
$\delta\text{-T}$	delta-tocopherol



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PREFACE

*To be healthy is our moral responsibility;
Towards Almighty God, ourselves, our family, and our society;
Eating fruits and vegetables makes us healthy;
Believe and have faith;
Reduction of food waste can reduce the world hunger
and can make our planet ecofriendly.*
—**Megh R. Goyal, PhD**
Senior Editor-in-Chief

Medicinal plants contain certain chemicals in their organs, such as leaves, stem, root, and fruits, which can provide therapeutic benefits against different kinds of diseases. These chemicals are often referred to as “phytochemicals.” The word “phyto” is a Greek word, that means “plant.” Phytochemicals are natural non-essential bioactive compounds found in plants/plant foods. Thousands of phytochemicals have already been identified, and more are still being discovered year by year. Plants that are used for medicinal purposes are often considered to be less toxic and induce fewer side effects than synthetic medicine. In our world today, many commercially available drugs have plant-based origins, with more than 30% of modern medicines directly or indirectly derived from medicinal plants. Indeed, plants can be a major source of pharmaceutical agents in the treatment of many life-threatening diseases.

The use of medicinal plants has largely increased because they are locally accessible, economical, as well as vital in promoting health. However, scientific data and information regarding the safety and efficacy of these medicinal plants are inadequate.

We introduce this book volume under the book series Innovations in Plant Science for Better Health: From Soil to Fork. This book mainly covers the current scenario of the research and case studies and contains scientific evidence on the health benefits that can be derived from medicinal plants and how their efficacies can be improved. The findings reported in this book can be useful in health policy decisions. It will also motivate the development of health care products from plants for health benefits. The book will further encourage the preservation of traditional medical knowledge of medicinal

plants. Therefore, these plant products are drawing the attention of researchers and policymakers because of their demonstrated beneficial effects against diseases with high global burdens such as diabetes, hypertension, cancer, and neurodegenerative diseases.

This book volume is a treasure house of information and an excellent reference for researchers, scientists, students, growers, traders, processors, industries, dieticians, medical practitioners, and others. We hope that this compendium will be useful for the students and researchers as well as those working in the food, nutraceutical, and herbal industries.

The contributions by the cooperating authors to this book volume have been most valuable in the compilation. Their names are mentioned in each chapter and in the list of contributors. We appreciate you all for having patience with our editorial skills. This book would not have been written without the valuable cooperation of these investigators, many of whom are renowned scientists who have worked in the field of plant science and food science throughout their professional career.

The goal of this book volume is to guide the world science community on how plant-based secondary metabolites can alleviate us from various conditions and diseases.

We will like to thank editorial and production staff, and Ashish Kumar, Publisher and President at Apple Academic Press, Inc., for making every effort to publish this book when all are concerned with health issues.

We request the reader to offer your constructive suggestions that may help to improve the next edition.

We express our admiration to our families and colleagues for their understanding and collaboration during the preparation of this book volume. As an educator, We give a piece of advice to one and all in the world: *“Permit that our almighty God, our Creator, provider of all and excellent Teacher, feed our life with Healthy Food Products and His Grace—; and Get married to your profession.”*

—Editors

PART I
Phytochemistry of Medicinal Plants



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CHAPTER 1

ANTIOXIDANT AGENTS FROM GREEN LEAFY VEGETABLES: A REVIEW

ABIOLA FATIMAH ADENOWO, MUHIBAH FOLASHADE ILORI, and
MUTIU IDOWU KAZEEM

ABSTRACT

Despite various reports on the biochemical importance of several green leafy vegetables abundant in Nigeria and other countries of sub-Saharan Africa, not enough research has been done on the utilization of these vegetables as nutraceuticals and drugs. It is highly recommended that further studies be carried out for:

1. Identification and isolation of the bioactive compounds responsible for the observed antioxidant properties of green leafy vegetables;
2. Utilization of isolated bioactive compound in the development of nutraceuticals and drugs to alleviate oxidative stress-related diseases and complications;
3. Utilization of bioactive component by the food manufacturing industry to retard oxidative degradation in foods in order to improve food quality and improve shelf-life.

1.1 INTRODUCTION

Green vegetables are herbaceous plant species, whose parts are eaten as auxiliary food or core dishes. The consumption of green leafy vegetables is a major aspect of cultural heritage that plays vital functions in the customs, traditions, and food culture of traditional households. Nigeria is bestowed with a diversity of traditional vegetables; and different ethnic groups consume different vegetables for various beneficial reasons [58].

Vegetables are reaped at different phases of growth and consumed either in fresh, processed, or semi-processed state by humans, whereas they are generally given in the fresh form to livestock. The nutritional constituents of vegetables differ substantially, but generally, they are not main sources of carbohydrates in comparison with starchy foods. Nevertheless, vegetables are packed with ample quantities of crude fiber, vitamins, minerals, carotene, and essential amino acids [38, 79].

Green leafy vegetables are useful for the preservation of health and prevention of diseases, due to treasured food nutrients necessary for body build-up as well as repair [1, 35]. Aside from rich nutritional values of vegetables, they are the cheapest and most abundant source of proteins due to their ability to produce amino acids from simple materials like water, carbon dioxide as well as atmospheric nitrogen [13, 20]. Aside from their low methionine content, most species of green leafy vegetables have amino acid profiles comparable with those of egg, fish, meat, and soybean; and their amino acid profile exceeds the Food and Agriculture Organization (FAO) stipulated pattern of essential amino acids [13].

Vegetables have low calories as well as insignificant amounts of utilizable energy; and therefore, these are ideal for obese individuals who can gratify their appetite without fear of accumulating calories. Additionally, vegetables are valuable in conserving alkaline reserve of the human body by acting as buffering mediators for acidic substances in the GI tract [4].

This chapter presents an overview of the antioxidant potential of green leafy vegetables and their role in the prevention and/or mitigation of oxidative stress-related diseases.

1.2 GREEN LEAFY VEGETABLES VERSUS OXIDATIVE STRESS-RELATED DISEASES

Numerous human degenerative conditions (such as cancer, atherosclerosis, *diabetes mellitus*, heart disease, stroke, ulcers, rheumatoid arthritis, osteoporosis, cataract, sunburn, and aging) have been documented as the outcome of damage by free radicals and reactive oxygen species (ROS) [3]. Several studies have also been undertaken on how to prevent or avoid the onset of such diseases. Nevertheless, the most possible and practical approach for combating degenerative ailments is by increasing the body antioxidant status, which can be realized by more intake of fruits and vegetables. Green leafy vegetables typically contain high amounts of natural antioxidants, which are able to scavenge free radicals [24, 25, 84].

The antioxidant action of leafy vegetables may be due to the presence of biocompounds like flavones, isoflavone, flavonoids, catechin, isocatechin, and anthocyanin rather than only vitamins C, E, and β -carotene [65, 70]. Dietary antioxidants are essential to manage ROS, which cause damage to the DNA, RNA, alter proteins, and affect lipid peroxidation (LPO) in the cells. Antioxidants can inhibit the commencement or proliferation of oxidative stress in organisms [64]. Researchers have shown a keen interest in the study of antioxidants for combating the deleterious consequence of free radicals; and natural products such as fruits and vegetables are in the limelight of such studies. Green leafy vegetables with reported antioxidant properties are discussed in this section.

1.2.1 *TALINUM TRIANGULARE* (WATER LEAF)

Talinum triangulare belongs to the family *Portulacaceae*, and originated from the tropical region of Africa, but it is extensively cultivated in Asia, South America and West Africa (especially in Nigeria) as a food crop [55]. It is greatly distributed in many ecological zones of Nigeria, where it is called ‘*Gbure*’ in Yoruba, ‘*Nte-oka*’ in Igbo and ‘*Alenyruwa*’ in Hausa [64]. *T. triangulare* is a perennial plant and is popularly called ‘waterleaf’ due to its high moisture content of approximately 90.8mg/100mg of edible leaf [86]. It is an herbaceous plant with succulent stem and pink flowers, which is used as a sauce, flavoring as well as condiment in foods. Furthermore, it is utilized in folk medicine to alleviate diuretic ailments, gastrointestinal disorder, and edema [9, 19, 58].

This plant is rich in protein, essential oils (EOs), total lipids, cardiac glycosides, flavonoids as well as polyphenols. Phytochemical studies have revealed the occurrence of omega-3-fatty acids and copious amounts of essential minerals such as calcium, potassium as well as magnesium. It also contains soluble fibers like pectin as well as vitamins C; α -tocopherol, β -tocopherols, and β -carotene that are required for growth and development [54, 68].

Anyasor et al., [19] investigated the *in-vitro* antioxidant activity of aqueous and methanol extracts of *T. triangulare*. They showed that both extracts tested positive to rapid thin layer chromatography (TLC) screening for the presence of antioxidant activity. The color change upon spraying with diphenyl picrylhydrazyl (DPPH) (deep violet to yellow spots) suggested the presence of free radical scavengers. Notably, the free radical scavenging ability of the extracts was increased with increase in the concentration of the extract. Additionally, thiobarbituric acid reactive (TBAR) substances assays

indicated that the extract prevented LPO at a concentration of 100 µg/ml. A similar study by Amorim et al., [16] showed that the stems of this vegetable have phenolic compounds possessing high antioxidant activity based on the results of 1,1-diphenyl-2-picrylhydrazyl (DPPH) scavenging activity.

Liang et al., [55] evaluated the antioxidant activity of polysaccharides from *T. triangulare*. The polysaccharides were extracted with boiling water and deproteinized using the Savage method. The polysaccharides demonstrated varied degrees of antioxidant activities in a dose-dependent fashion, which necessitated further studies towards its utilization for the management of oxidative stress-induced diseases. Methanol and hydro-ethanol extracts of *T. triangulare* were also studied for their Fe³⁺ reducing ability and free radical scavenging activity. Results showed that both extracts demonstrated antioxidant activity, though the hydro-ethanol extract had stronger antioxidant properties [71].

1.2.2 TELFAIRIA OCCIDENTALIS (FLUTED PUMPKIN)

Telfairia occidentalis (fluted pumpkin) belongs to the family *Cucurbitaceae*. Once harvested, the leaves are cautiously detached from the stems because the stems are regarded as poisonous and are thus thrown away as waste [31]. Fluted pumpkins occur chiefly in the forest zones of Central and West Africa, mostly in Cameroun, Republic of Benin and Nigeria. The leaves of this plant are widely consumed due to its acknowledged nutritional and medicinal benefits. In Nigeria, it is called by various traditional names such as ‘Ugu’ in Igbo, ‘Aporoko/Iroko’ in Yoruba, ‘Ubong’ in Efik and ‘Umeke’ in Edo language [51, 67].

The seeds are cooked and used as a protein supplement in a wide range of local foods. The seeds are also utilized as composite flours for the manufacture of bakery produce like bread and cookies [39]. Mineral elements in this plant include calcium, magnesium, phosphorus, sodium, and iron. Interestingly, the high iron content (approximately 700 ppm) scientifically authenticate the folk tradition of administering aqueous extract of the leaves as a blood tonic to convalescing individuals [51, 67]. *T. occidentalis* also contain phenols, flavonoids, alkaloids, oxalates, saponins, resins, and glycosides [67, 81]. Additionally, it is enriched with amino acids, including alanine, aspartate, cysteine, methionine, and phenylalanine [20, 51].

Oboh et al., [67] investigated the antioxidant activity of aqueous and ethanol extracts of *T. occidentalis*. Both extracts showed antioxidant activity, but the aqueous extract exhibited significantly higher reducing

power (1.9 O.D.₇₀₀) and free radical scavenging power (92%) than the ethanol extract. Several other researchers have equally established, with both *in vivo* and *in vitro* experiments, the ability of different extracts of *T. occidentalis* to prevent production of free radicals, scavenge free radicals, lower LPO as well as raised levels of antioxidant enzymes like catalase (CAT) and superoxide dismutase (SOD) [2, 46, 50, 63].

The recorded antioxidant potential of *T. occidentalis* might be attributed to the presence of bioactive constituents such as polyphenols, flavonoids, and vitamin C. An *in vivo* study showed the ability of aqueous leaf extract to enhance the oxidative status of the reproductive system of male Sprague-Dawley rats. The experimental rats were orally administered with 200, 400, and 800 mg/kg/day of freshly prepared aqueous extracts for 56 days (for spermatogenesis to take place). Thereafter, the animals were sacrificed, and the testicular oxidative status was assessed by measuring CAT activity, SOD activity, and glutathione peroxidase (GPx) activity. Results showed a dose-dependent (200 mg/kg/day gave best results) testiculo-protective ability of the aqueous extract of *T. occidentalis* [81].

1.2.3 GONGRONEMA LATIFOLIUM (UTAZI)

Gongronema latifolium (called ‘Utazi’ and ‘Arokeke’ in Igbo and Yoruba languages respectively in Nigeria) is a tropical rain forest plant predominantly used as a vegetable and spice. It is a member of the *Asclepiadaceae* family [32, 34]. This bitter-tasting green leafy vegetable is used traditionally in the management of anorexia, malaria, and nausea. Furthermore, the liquor of *G. latifolium* obtained after slicing the plant and boiling with lime extract or steeping in water for about three days is commonly taken as a purgative for colic, stomach upset and to treat signs associated with worm infections [35, 87].

Reports showed that this plant contains appreciable amounts of protein, fiber, saponins, EOs, flavonoids, and minerals (calcium, potassium, sodium, phosphorus, and cobalt). The amino acid profile revealed the plant is made up of both essential and non-essential amino acids and its pattern of amino acids is comparable with the World Health Organization (WHO) standards. Noteworthy is also the occurrence of high amounts of aspartic acid, glycine, and glutamic acid (13.8%, 10.3%, and 11.9%, respectively) [32, 33].

Fasakin et al., [37] studied the antioxidant activity of polyphenol extracts of *Gongronema latifolium* *in vitro*. Various assays (DPPH radical scavenging, chelation of metal ion, hydroxyl radical scavenging assay,

superoxide scavenging assay, and ferric reducing activity) confirmed the antioxidant property of various extracts of *G. latifolium*. The authors suggested that the leaf extract can function as a prospective source of natural antioxidants, while further research is needed for identification and isolation of the bioactive compounds for utilization in the food industry [37]. Similarly, other studies also confirmed the antioxidant activity of aqueous and ethanol extracts of *G. latifolium* [14, 87].

1.2.4 VERNONIA AMYGDALINA (BITTER LEAF)

Vernonia amygdalina belongs to the *Asteraceae* family, which grows in many African countries, including Cameroun, Zimbabwe, and Nigeria. It is commonly referred to as bitter leaf due to its bitter taste, which can be eliminated by continuous soaking in water and cooking. It is consumed as a vegetable food and in herbal preparations. Traditionally, it is used as a tonic, for tick control and in the treatment of constipation, fever, dysentery, hypertension, cough, and sexually transmitted diseases [36, 44]. Organic extracts of *V. amygdalina* have been established to possess cytotoxic activity against human carcinoma cells of nasopharynx as well as having antiparasitic and antimicrobial activities [36]. Numerous compounds with potent biological activities have been isolated from *V. amygdalina*, such as:

- Flavonoids;
- Anthraquinone;
- Steroids;
- Alkaloids;
- Glycosides;
- Luteolin;
- Luteolin 7-0-glucuronide;
- Luteolin 7-0-glucoside;
- Vernonioside [36, 47].

Johnson et al., [48] investigated the *in vitro* antioxidant activity of methanolic extract of *V. amygdalina*. The extract was established to possess DPPH scavenging activity of 96.65 µg/ml and Ferric reducing antioxidant power (FRAP) of 0.708. The results indicated that the extract possesses strong antioxidant activity [44]. Atangwho et al., [21] also reported a dose-dependent *in vitro* antioxidant activity of aqueous, methanol, chloroform as well as petroleum ester extracts of *V. amygdalina*.

Erasto et al., [34] also investigated the antioxidant effects (using reducing power assay and DPPH radical scavenging activity) of ethanol extract and two formerly isolated sesquiterpene lactones (vernodalol and vernolide) were reported. They showed that vernolide exhibited a higher reducing power than ethanol extract and vernodalol, while the ethanol extract showed higher radical scavenging activity. However, all three samples from *V. amygdalina* demonstrated appreciable antioxidant properties. Similarly, the antioxidant activity of boiled, cold, and methanolic extracts of *V. amygdalina* was estimated using DPPH free radical assay. Results confirmed the antioxidant action of this green leafy vegetable at various concentrations studied [43].

1.2.5 *OCIMUM GRATISSIMUM* (CLOVE BASIL OR SCENT LEAF)

Ocimum gratissimum is a perennial plant that is broadly distributed in the tropics of Africa as well as warm temperature regions. *O. gratissimum* (popularly called ‘Scent leaf’) is a traditional vegetable condiment used in Nigeria and some other countries to increase food flavor, as well as oral care products. This aromatic plant is widely used as expectorant and carminative, and traditionally employed in the treatment of epilepsy, rheumatism, paralysis, gonorrhea, diarrhea, influenza, and mental illness in India and Africa [22, 78]. It is used in south-eastern Nigeria in the management of umbilical cord of neonates to sustain sterility of the wound surfaces. Its roots are also used by Brazilian tropical forest inhabitants as part of the constituents of a decoction as a sedative for children [77]. Volatile aromatic oil from the leaves comprises principally of thymol and eugenol as well as xanthenes, terpenes, and lactones, which possess antiseptics, antibacterial, and antifungal activities as well as an insect repellent [10].

The free radical scavenging ability of methanol extract of the leaves of *O. gratissimum* was evaluated by assessing its ability to scavenge 2,2-diphenyl-1-picrylhydrazyl, superoxide anion, hydroxyl, nitric oxide (NO) radicals, and its capability to inhibit LPO. It was concluded that the plant is a potential source of natural antioxidants that would be valuable in food manufacturing to retard oxidative degradation of lipids and thus increase food quality [22].

Akinmoladun et al., [10] showed that methanol extract of *O. gratissimum* had a DPPH scavenging activity of 84.6% at 250 µg/ml and a reductive potential of 0.77 at 100 µg/ml. These values were comparable with values obtained for standards, gallic acid, and ascorbic acid. These results affirm

to the wide traditional use of the plant extract in the management of various human maladies. The essential oil of *O. gratissimum* has also been confirmed to possess good antioxidant activity through the DPPH radical scavenging activity and β -carotene-linoleic acid bleaching assay in a dose-dependent fashion [78]. This is an indication that the essential oil will be valuable in food processing and preservation.

1.2.6 *CORCORUS OLITORIUS* (JUTE)

C. olitorius (Jute) belongs to the *Tiliaceae* family. This annual herb is native to Africa. It is also cultivated in several parts of India and Bangladesh [40, 65]. Jute plant is a common traditional vegetable, which is prepared into a slimy soup or sauce in many of West African culinary traditions. In Western Nigeria, it is known as ‘ewedu’ while the Songhay people of Mali refer to it as ‘fakohoy.’ Furthermore, it is a common meal ingredient in the Northern province of the Philippines, where it is known as ‘saluyot’ and in Taiwan where it is boiled with sweet potatoes into a nourishing meal. The plant is rich in flavonoids, α -tocopherol, carotenoids, polyphenols, as well as vitamin C, iron, and calcium [65, 91]. It is popularly used in folk medicine for treating ailments like dysentery, bone pains, fever, gastroenteritis, and diabetes. The seeds are also used traditionally as a contraceptive [26, 40, 92].

Azuma et al., [23] identified six phenolic compounds from the leaves of *C. olitorus* using nuclear magnetic resonance (NMR) and Fast atom bombardment mass spectrometry (FAB-MS). The compounds are 5-caffeoylquinic acid (chlorogenic acid), 3,5-dicaffeoylquinic acid, quercetin 3-galactoside, quercetin 3-glucoside, and quercetin 3-(6-malonylglucoside). Thereafter, the content of these phenolic acids, α -tocopherol, and ascorbic acid was assessed, and their antioxidant potential was measured with radical generator-initiated peroxidation of linoleic acid. The results revealed that 5-caffeoylquinic acid was the major phenolic antioxidant in the leaves of *C. olitorus* [23].

Similarly, Oboh et al., [69] assessed the antioxidant activity of hydrophilic (water) and lipophilic (hexane) extracts of *C. olitorus* *in vitro*. Their study showed that both extracts exhibited significant antioxidant properties. However, the hydrophilic extract showed higher free radical scavenging ability, reducing power and trolox equivalent antioxidant capacity than the lipophilic extract, due to the higher content of total phenol, total flavonoid, and ascorbic acid compared with the lipophilic extract [69].

1.2.7 *GNETUM AFRICANUM* (UKAZI)

Gnetum africanum belongs to the family *Gnetaceae*, which is extensively distributed in Cameroun, Equatorial Guinea, Central Africa Republic, Gabon, and Nigeria. It is widely consumed in the South East region of Nigeria, where it is known as ‘Afang’ or ‘Ukazi.’ It is often cooked with water-leaves and also consumed as vegetable salad [15, 30].

Traditionally, *G. africanum* is used in the management of several illnesses including nausea, boils, enlarged spleen, and neutralization of some poisons and to reduce the pain during childbirth. It is also used for the treatment of hemorrhoids, diabetes, and as worm expeller [46, 76]. The leaves have been reported to have great nutritional value constituting a significant source of protein, minerals, and essential amino acids. Numerous molecular compounds associated to the families of stilbenes, flavonostilbenes, and glycosylflavones have been isolated and identified in the leaf extract of this plant. These compounds may be responsible for the interesting properties and biological activities of the plant [15, 29].

The antioxidant potential of both raw and cooked leaf extracts of *G. africanum* was investigated by Ogbonnaya and Chinedum [72]. The results for vitamin assay showed that both the raw and cooked leaf extracts had a significant content of vitamin C and E, and their concentrations were not affected by cooking. Analysis of the scavenging activities of both extracts using 2,2-diphenyl-1-picrylhydrazyl radical showed that both extracts had significant scavenging activity. The results obtained can be attributed to their polyphenolic contents, which possess varying levels of antioxidant activity and the hydrogen donating capacity of the OH groups of the phenolic compounds. Similar results were also reported by other researchers [6, 11].

1.2.8 *PIPER GUINEENSE* (AFRICAN BLACK PEPPER)

Piper guineense belongs to the family *Piperaceae*, and there are about 700 species spread across tropical and subtropical regions of the world. It is generally called African black pepper or hot leaf in various parts of Africa such as Ghana, Cameroon, and Nigeria, where it is consumed as a spice due to its nutritional and therapeutic properties. In Nigeria, it is known by various local names such as ‘Uziza’ in Igbo and ‘Iyere’ in Yoruba [18, 60]. In Nigeria, the seeds are used by women after childbirth to increase uterine contraction in order to ease the ejection of the placenta as well as other residues from the womb. The leaves are also used in the management of

cough, male infertility, respiratory, and intestinal diseases. They also have carminative, appetitive, and eupeptic properties [7]. The antimicrobial, antiparasitic, antifungal, and insecticidal activities of the leaves and seeds have also been reported. Its phytoconstituents include:

- Alkaloids;
- Sterols;
- Flavonoids;
- Saponins;
- Glycosides [18, 60].

The hydro-ethanolic and ethanolic extracts of leaves and stems of the *P. guineense* were investigated for their free radical scavenging activities as well as antioxidant potential. The results showed that the extracts significantly inhibited the DPPH, ABTS⁺, NO, and OH⁻ radicals in a concentration-dependent manner. A significant ferrous ion chelating ability was also observed through FRAP and phosphomolybdenum antioxidant potential assays. The study also revealed that the polyphenol content of this plant differs subject to the form of extracts and the solvent used. However, the hydro-ethanolic and ethanolic extracts of leaves presented a higher level of phenolic compounds [60]. Hence further studies need to be done to isolate and characterize the bioactive compounds responsible for the observed action. Similarly, Agbor et al., [7] investigated the *in vitro* antioxidant potential of three Piper species including *P. guineense*, and their results showed that methanolic leaf extract of all species studied possess antioxidant activity in a dose-dependent manner [7].

1.2.9 CELOSIA ARGENTIA (PLUMED COCKSCOMB)

Celosia argentea is a tropical herbaceous plant from the family *Amaranthaceae*, known for its attractive and distinctive brightly-colored flowers. It is commonly referred to as plumed cockscomb or the feathery amaranth. It is widely distributed in Southern Asia, China, and some parts of Africa. It is popularly consumed as a vegetable in the Western part of Nigeria, where it is known as ‘red soko’ due to red pigmentation on its leaves [49, 57, 90]. The seeds are very small and are used traditionally for the management of jaundice, fever, gonorrhea, and wounds; and for the treatment of itching, sores, ulcers, and fever. The plant has been studied for antibacterial, anti-inflammatory, diuretic, antipyretic, antidiarrheal and antidiabetic properties. Amino acids,

carbohydrates, flavonoids, tannins, saponins, phytosterols, and glycosides are some of the reported phytochemicals present in the plant [54, 82].

Mahadik et al., [56] investigated the antioxidant activity of ethanolic extract of *C. argentea* *in vitro* via the reducing power assessment. They showed that the ethanolic extracts of *C. argentea* compared significantly with that of the standard antioxidant agent ascorbic acid. Combination of the extract and ascorbic acid also produced a synergistic effect. These results indicated that ethanolic extract of the plant can be utilized as natural antioxidant either alone or in combination with ascorbic acid. A similar study with methanolic leaf extracts of *Celosia argentea* using DPPH, NO, and hydrogen peroxide (H₂O₂) radical showed a concentration-dependent free radical scavenging property [89]. *In vivo* and *in vitro* studies revealed the antioxidant ability of various extracts of *C. argentea*, and the observed activity was attributed to the presence of phenolic compounds in the plant [41, 80].

1.2.10 *SOLANUM MACROCARPON* (EGGPLANT)

Solanum macrocarpon is a flowering herbaceous plant usually used for its nutritional content and medicinal properties. It is commonly cultivated in Western Africa countries, where it functions as a source of fruit and leafy vegetable [27, 52]. It is found in the southwestern region of Nigeria, where it is called igbagba or 'Igbo' [62, 66]. The leaves are packed with fat, protein, calcium, zinc, crude fiber, as well as appreciable amounts of the amino acid and methionine [52].

Besides the use of the leaves for culinary purpose, different parts of the plants are used traditionally for diverse purposes. In Sierra Leone, the leaves are heated and masticated to treat throat complications; while in Kenya, the liquor from boiled roots is drinking to purge out hookworms, whereas the crushed leaves are taken to alleviate stomach disorders [27, 83]. The fruits are eaten in Nigeria as purgatives and are also used in the management of heart disease, while the flowers are chewed to maintain oral hygiene. The roots are also used for treating wounds, body aches, bronchitis, itch, and asthma while the seeds are used to cure toothache. Several reports on aqueous extract of its fruits have shown its hemolytic, hepatoprotective, and hypolipidemic activities [27, 52, 62, 66].

Study on the free radical scavenging activity of the ethanolic leaves extract using the DPPH assay showed a dose-dependent activity, which may be attributed to its high phenolic content [52]. Similarly, Olajire and Azeez [74] investigated the total antioxidant activity of various commonly

consumed vegetables in Nigeria. Their findings showed that *S. macrocarpon* has the best antioxidant properties, compared to other vegetables and standard antioxidant agent tocopherol. This was revealed by its lowest IC₅₀ value (6.21 mg ml⁻¹), compared to α -tocopherol (IC₅₀: 13.20 mg mL⁻¹). It also possesses the highest total phenolic, flavonoid as well as ascorbic acid contents compared with the other vegetables studied. Similar works also confirmed the ability of various extracts of *S. macrocarpon* leaves to alleviate oxidative stress due to its antioxidant activity [52, 59, 73].

1.2.11 *BASELLA ALBA (VINE SPINACH)*

Basella alba is a perennial plant, which belongs to the *Basellaceae* family. It is commonly called Malabar spinach, Red vine spinach or creeping spinach. It is generally found in the tropical countries of the world, including Nigeria, where it is common in the south-west region. It is composed of fat, proteins, vitamins A, B (folic acid), C, E, and K, as well as riboflavin, niacin, thiamine. It also has minerals, including iron, calcium, and magnesium. Studies showed that it has some unique constituents like betalain, basellasaponins, and kaempferol [5, 42, 75].

Various parts of the plant are used for several purposes in traditional medicines. In Nepal, the leaf extract is used in the treatment of cold and cough while the pastes are used externally in treating boils. The boiled leaves and stems are administered as laxatives, and the flowers are useful as an antidote for poisons [88]. The leaves and stem are also used in the management of diarrhea, dysentery, anemia, cough, cold, headaches, and ringworm. It also aids in the removal of placenta after childbirth as well as increase the flow of milk in nursing mother [5, 85].

Sridevi et al., [85] investigated the antioxidant potential of ethanolic leaf extract of *Basella alba* using *in vitro* assays and using 2,2-diphenyl-1-picrylhydrazyl (DPPH), reducing power assay, and phosphomolybdenum assay. The results showed that the extract exhibited significant free radical scavenging activity compared with the standard, gallic acid.

In an effort to study the antioxidant potential of ethyl acetate and chloroform extracts of *Basella alba* leaves, the phenol and flavonoid content of the extracts were measured by aluminum chloride assays and Folin Ciocalteu assay, respectively. Results showed that the total phenolic contents of the ethyl acetate and chloroform extracts were 0.029 mg/g and 0.030 mg/g, respectively, whereas the total flavonoids content were 0.045 mg/g and 0.085 mg/g, respectively.

Furthermore, both extracts showed radical scavenging activity in a concentration-dependent manner. Hence, *Basella alba* contains ample antioxidant agents and may be valuable for the formulation of drugs/nutraceuticals to combat oxidative stress-related disorders [75].

1.3 SUMMARY

This review summarizes some reported findings on the antioxidant potential of some commonly consumed green leafy vegetables in Nigeria. It focuses on the reported ability of various extracts of the studied green leafy vegetables to alleviate conditions associated with oxidative stress both *in vivo* and *in vitro*. Hence, these green vegetables are recommended for further study towards the development of nutraceutical and drugs for combating oxidative stress associated conditions.

KEYWORDS

- antioxidants
- bitter leaf
- clove basil
- epilepsy
- fluted pumpkin
- free radicals
- kaempferol
- omega-3 fatty acid
- oxidative stress
- plumed cockscomb
- polyphenols
- saponins
- superoxide dismutase
- nuclear magnetic resonance
- vernodalol
- waterleaf

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