



ST. XAVIER'S COLLEGE

(AUTONOMOUS)

**5, Mahapalika Marg, Mumbai - 400 001,
INDIA.**

☎ 2262 0661/65

3.4.4: BOOKS AND CHAPTERS IN EDITED VOLUMES/BOOKS PUBLISHED BY TEACHERS

❖ 2017-18 (8)



Advanced Oxidation Processes for Wastewater Treatment

Emerging Green Chemical Technology

Edited by
Suresh C. Ameta
Rakshit Ameta



ADVANCED OXIDATION PROCESSES FOR WASTEWATER TREATMENT

Emerging Green Chemical Technology

Edited by

SURESH C. AMETA

PAHER University, Udaipur, Rajasthan, India

RAKSHIT AMETA

J. R. N. Rajasthan Vidyapeeth (Deemed-to-be University), Udaipur, Rajasthan, India



ACADEMIC PRESS

An imprint of Elsevier

Academic Press is an imprint of Elsevier
125 London Wall, London EC2Y 5AS, United Kingdom
525 B Street, Suite 1800, San Diego, CA 92101-4495, United States
50 Hampshire Street, 5th Floor, Cambridge, MA 02139, United States
The Boulevard, Langford Lane, Kidlington, Oxford OX5 1GB, United Kingdom

Copyright © 2018 Elsevier Inc. All rights reserved.

No part of this publication may be reproduced or transmitted in any form or by any means, electronic or mechanical, including photocopying, recording, or any information storage and retrieval system, without permission in writing from the publisher. Details on how to seek permission, further information about the Publisher's permissions policies and our arrangements with organizations such as the Copyright Clearance Center and the Copyright Licensing Agency, can be found at our website: www.elsevier.com/permissions.

This book and the individual contributions contained in it are protected under copyright by the Publisher (other than as may be noted herein).

Notices

Knowledge and best practice in this field are constantly changing. As new research and experience broaden our understanding, changes in research methods, professional practices, or medical treatment may become necessary.

Practitioners and researchers must always rely on their own experience and knowledge in evaluating and using any information, methods, compounds, or experiments described herein. In using such information or methods they should be mindful of their own safety and the safety of others, including parties for whom they have a professional responsibility.

To the fullest extent of the law, neither the Publisher nor the authors, contributors, or editors, assume any liability for any injury and/or damage to persons or property as a matter of products liability, negligence or otherwise, or from any use or operation of any methods, products, instructions, or ideas contained in the material herein.

British Library Cataloguing-in-Publication Data

A catalogue record for this book is available from the British Library

Library of Congress Cataloging-in-Publication Data

A catalog record for this book is available from the Library of Congress

ISBN: 978-0-12-810499-6

For Information on all Academic Press publications
visit our website at <https://www.elsevier.com/books-and-journals>

		Working together to grow libraries in developing countries
www.elsevier.com • www.bookaid.org		

Publisher: Candice Janco
Acquisition Editor: Louisa Hutchins
Editorial Project Manager: Emily Thomson
Production Project Manager: Swapna Srinivasan
Cover Designer: Miles Hitchen

Typeset by MPS Limited, Chennai, India

Contents

List of Contributors xi

About the Authors xiii

Preface xv

1. Introduction

SURESH C. AMETA

- 1.1 Environment 1
- 1.2 Pollution 1
- 1.3 Water Pollution 2
- 1.4 Wastewater Treatment 3
- 1.5 Advanced Oxidation Processes 6
- 1.6 Advantages 9
- 1.7 Applications 10
- References 11

2. UV-Hydrogen Peroxide Processes

JOSÉ C. MIERZWA, RAPHAEL RODRIGUES
AND ANTONIO C.S.C. TEIXEIRA

- 2.1 Introduction 13
- 2.2 Fundamentals 13
- 2.3 Kinetics 20
- 2.4 A Simplified Model for Performance
Evaluation 25
- 2.5 UV/H₂O₂ Oxidation Process Design 30
- 2.6 Practical Applications 43
- References 46

3. Fenton and Photo-Fenton Processes

RAKSHIT AMETA, ANIL K. CHOHADIA,
ABHILASHA JAIN AND PINKI B. PUNJABI

- 3.1 Introduction 49
- 3.2 Types of Fenton Processes 50
- 3.3 Electro-Fenton Processes 57
- 3.4 Sono-Fenton and Sono-Photo-Fenton
Processes 58

3.5 Heterogeneous Fenton and Photo-Fenton
Processes 59

3.6 Combined (Hybrid) Fenton and Photo-Fenton
Processes 62

3.7 Applications 64

3.8 Recent Developments 74

References 76

4. Ferrioxalate-Mediated Processes

RUBEN VASQUEZ-MEDRANO, DORIAN PRATO-GARCIA
AND MICHEL VEDRENNE

- 4.1 Introduction 89
- 4.2 The Fenton and Photo-Fenton
Reactions 90
- 4.3 The Ferrioxalate-Mediated Fenton
Reaction 92
- 4.4 Applications 100
- 4.5 Future Trends 106
- References 108
- Further Reading 113

5. Ozone-Based Processes

KEISUKE IKEHATA AND YUAN LI

- 5.1 Introduction 115
- 5.2 Ozone-Based AOPs 116
- 5.3 Ozonation By-Products 119
- 5.4 Wastewater Ozonation and Ozone-Based
AOPs 120
- 5.5 Recent Studies 121
- 5.6 Existing Ozone-Based Advanced Water
Reclamation Facilities 126
- 5.7 Planned Ozone-Based Advanced Water
Reclamation Projects 129
- 5.8 Concluding Remarks 130
- References 131

6. Photocatalysis

RAKSHIT AMETA, MEENAKSHI S. SOLANKI, SURBHI
BENJAMIN AND SURESH C. AMETA

- 6.1 Introduction 135
- 6.2 Photocatalysis 136
- 6.3 Modifications 144
- 6.4 Wastewater Treatment 154
- 6.5 Immobilization 162
- 6.6 Effect of Morphology 164
- 6.7 Other Applications 165
- References 165

7. Sonolysis

RICARDO A. TORRES-PALMA AND EFRAIM
A. SERNA-GALVIS

- 7.1 Introduction 177
- 7.2 Principles of the Process 177
- 7.3 Types of Main Reactors (Reaction Systems) 180
- 7.4 The Effect of Sonochemical Operational Parameters 181
- 7.5 Effect of the Chemical Pollutant Nature and Its Transformations Upon Sonochemical Process 192
- 7.6 Influence of Water Matrix in the Pollutants Degradation 199
- 7.7 Combination of Sonochemistry With Other Processes 206
- References 209
- Further Reading 213

8. Microwave/Hydrogen Peroxide Processes

AFSANE CHAVOSHANI, MOHAMMAD M. AMIN,
GHORBAN ASGARI, ABDOLMOTALEB SEIDMOHAMMADI
AND MAJID HASHEMI

- 8.1 Introduction 215
- 8.2 Wastewater Treatment 219
- 8.3 Enhancement of Sludge Anaerobic Biodegradability 224
- References 251
- Further Reading 255

9. Gamma-ray, X-ray and Electron Beam Based Processes

MAREK TROJANOWICZ, KRZYSZTOF BOBROWSKI,
TOMASZ SZREDER AND ANNA BOJANOWSKA-CZAJKA

- 9.1 Introduction 257
- 9.2 Sources of Radiation—Technological Installations 263
- 9.3 Disinfection of Wastewaters 271
- 9.4 Radiolytic Decomposition of Individual Compounds 274
- 9.5 Chemical Enhancement of Radiolytic Processes 295
- 9.6 Purification of Wastewaters of Different Origin 299
- 9.7 Economic Aspects 314
- 9.8 Conclusions 318
- Acknowledgments 319
- References 319

10. Supercritical Water Oxidation

VIOLETA VADILLO, JEZABEL SÁNCHEZ-ONETO,
JUAN R. PORTELA AND ENRIQUE J. MARTÍNEZ
DE LA OSSA

- 10.1 Introduction 333
- 10.2 Development of SCWO 338
- 10.3 Detected Problems 341
- 10.4 Energy Recovery in SCWO Plants 348
- 10.5 Economic Aspects 350
- 10.6 Conclusions 351
- Acknowledgements 351
- References 351
- Further Reading 358

11. Electrochemical Oxidation Processes

KURAVAPPULLAM V. RADHA AND
KARUNAMOORTHY SIRISHA

- 11.1 Introduction 359
- 11.2 Electrochemical Oxidation Processes 360
- 11.3 Advantages 369
- 11.4 Disadvantages 369
- 11.5 Applications 369
- 11.6 Current Scenario 370
- 11.7 Future Prospects 371
- References 372

12. Catalytic Wet Peroxide Oxidation

ALI R. TEHRANI-BAGHA AND TAREK BALCHI

12.1 Introduction 375

12.2 Catalysts for CWPO 377

12.3 Efficiency of CWPO of Phenol 384

12.4 Effect of the Main Parameters 388

12.5 CWPO Performance 392

References 396

Index 403

Fenton and Photo-Fenton Processes

*Rakshit Ameta¹, Anil K. Chohadia², Abhilasha Jain³
and Pinki B. Punjabi⁴*

¹J. R. N. Rajasthan Vidyapeeth (Deemed-to-be University), Udaipur, Rajasthan, India

²M. P. Govt. P. G. College, Chittorgarh, Rajasthan, India ³St. Xavier's College, Mumbai, Maharashtra, India ⁴M. L. Sukhadia University, Udaipur, Rajasthan, India

3.1 INTRODUCTION

Global economic growth is increasing exponentially in the first century of the new millennium, but at the same time, rapid urbanization and industrialization release enormous volumes of wastewater imposing various adverse effects on human health and grading the quality of the environment as a whole. It has been revealed that generation of wastewaters with complex and recalcitrant molecules is increasing day by day. The presence of these organic compounds in water poses a serious threat to public health since most of them are toxic, endocrine disrupting, mutagenic, or potentially carcinogenic to humans, animals, and aquatic life. There is a pressing demand for newer technologies for the complete mineralization of wastewaters.

Several conventional treatment methods are available such as biological, adsorption, chemical treatment, filtration, flocculation, activated charcoal and ion exchange resins for wastewater remediation. It has been frequently observed that pollutants not amenable to biological treatments may also be characterized by high chemical stability and/or by strong difficulty to be completely mineralized. In this context, oxidation processes are preferred to degrade such biorefractory substances present in wastewater. However, pollution load, process limitations, and operating conditions are the key factors to be considered during the selection of the most appropriate oxidation process for the degradation of a particular compound. Apart from high degradation efficiency, direct oxidation processes demand specified operating conditions to degrade the target compounds, which will increase the operation cost of the process.

Floral Diversity of Nandur Madhameshwar Wildlife Sanctuary

A Pictorial guide



**By
Rajendra D. Shinde**

**Floral Diversity of Nandur Madhameshwar Wildlife Sanctuary
- A pictorial Guide**

Concept and Text:

Dr Rajendra D. Shinde

Head, Dept of Botany & Director, Blatter Herbarium,
St. Xavier's College, Mumbai 400 001.
E-Mail : rajendra.shinde@xaviers.edu

Photographs:

Dr Rajdeo Singh
E-Mail: rajdeo.1982@gmail.com

**Front Cover : *Ipomoea nil*
& Back Cover by :**

Mr Bharat Shinde
E-Mail: phoenix8585@gmail.com

Published by:

Nashik Wildlife Division,
Aranya Sankul,
Old Mumbai Agra Road,
Trambak Naka, Nashik - 422 002.
Tel.: 0253-2505114/15
E-Mail: acfnmsnashik@gmail.com

Layout and Design:

AP ADVERTISING NETWORK
202, 2nd Floor, Niti Apt.,
Off. S. V. Road, Malad (W),
Mumbai - 400 064.
Tel.: 022-2844 6989
E-Mail: apadvertising@gmail.com

Copy Right © Rajendra D. Shinde

Price: Rs. 150/-

Floral Diversity of Nandur Madhameshwar Wildlife Sanctuary

A Pictorial guide



By
Rajendra D. Shinde

PREFACE

The diverse flora of India has been documented mainly in formal "Floras" ranging from J.D.Hooker's Flora of British India (1875-1897) to district or even taluka-level floras. However, the rapidly growing interest amongst laypeople in the environment and in nature study in the last two to three decades has created a demand for literature which would help a layperson to identify and learn more about the flora of a region; Flowers of Sahyadri by Shrikant Ingalhalikar (2001) was a pioneering and very successful attempt to fill this lacuna for the much-visited mountains of the Sahyadris; Pradip Krishen's Trees of Delhi (2006) and his more recent, Jungle Trees of Central India - a Field Guide for Tree Spotters (2013) are other popular examples; incidentally both the above authors are non-botanists!

A visitor to a wildlife sanctuary is primarily interested in the fauna, in a wetland the primary interest is birds, yet in the intervals when the animals are not visible a secondary interest often arises in identifying the flora, especially flowers or plants with some peculiar morphology. This book is an attempt to help the visitor to Nandur Madhameshwar Wildlife Sanctuary identify and learn some uses of the plants most likely to be encountered and noticed by an amateur visiting the Sanctuary. We hope to stimulate an interest in the diversity and beauty of the flowering plants in a wetland and, thus, promote their appreciation and conservation.

Rajendra Dattatraya Shinde

ACKNOWLEDGEMENT

This book would not have been possible without the help of Late Dr. Marselin R. Almeida and Dr. (Ms.) Saramma Almeida who initiated the project and was involved in the project till its completion.

I thank the Maharashtra Forest Department for financing this book, especially Shri. M.K. Rao, Additional Principal Chief Conservator of Forests (Wildlife West), Borivali, Mumbai; Shri. N. R. Praveen, Conservator of Forests (Wildlife), Nashik, and Shri. Bharat Shinde, Assistant Conservator of Forest, Nandur Madhameshwar Wildlife Sanctuary, Nashik; it was Shri. B. Shinde who pushed me to write this book.

I am thankful to Dr. Rajdeo Singh who took the responsibility of taking photographs in the field as also in the compilation of the book & Ms. Candice Dcosta, who painstakingly edited the original manuscript of my thesis earlier.

I would like to acknowledge the help provided by Mr. V. K. Mohan, Retired IFS office, who was the DFO, Nashik during 1984-88, and Mr. Debi Goenka who accompanied us on field trips and taught me to identify a few common waders.

I am grateful to Dr. Agnelo Menezes, Principal, St. Xavier's College (Autonomous), Mumbai for constant support and encouragement, and to my colleagues in the Blatter Herbarium & Botany Department for helping me in sharing my responsibilities and giving me time to do this work.

And finally to my family who always support me in all my endeavors...

Rajendra Dattatraya Shinde
Mumbai
January 11, 2018.

Disclaimer

All the contents or information provided in this book is designated to provide helpful information on the subjects discussed. This book is not meant to be used, nor should it be used, to diagnose or treat any medical conditions. For diagnosis or treatment of any medical problem, consult your own physician. The publisher and author are not responsible for any specific health or allergy needs that may require medical supervision and are not liable for any damages or negative consequences from any treatment, action, application or preparation, to any person reading or following the information in this book.

Neither the publisher nor the author shall be liable for any physical, psychological, emotional, financial or commercial damages, including, but not limited to, special, incidental, consequential or other damages.

Rajendra D. Shinde

Nashik Wildlife Division

INTRODUCTION :

Nandur Madhameshwar Wildlife Sanctuary, also known as 'Mini-Bharatpur' of Maharashtra, is situated at 20°00.780' N and 74°10.4424' E in Niphad tehsil of Nashik district (Map 1). The primary routes to reach this area are from Nashik via, either Sayakheda (35 kms) or Sinnar (55 kms). Niphad railway station on the Central Railway is 12 kms from Nandur Madhameshwar and can be traversed by ST bus. A stone pick-up weir was constructed in 1907-13 across the river Godavari just below the confluence of Kadwa and Godavari rivers at Nandur Madhameshwar; the water level therefore is always fluctuating in Nandur Madhameshwar Lake. This reservoir is surrounded by grape vineyards and fields of sugarcane, onions, jowar, wheat. There are no forests in this area but it is rich in herbaceous flora and aquatic vegetation.

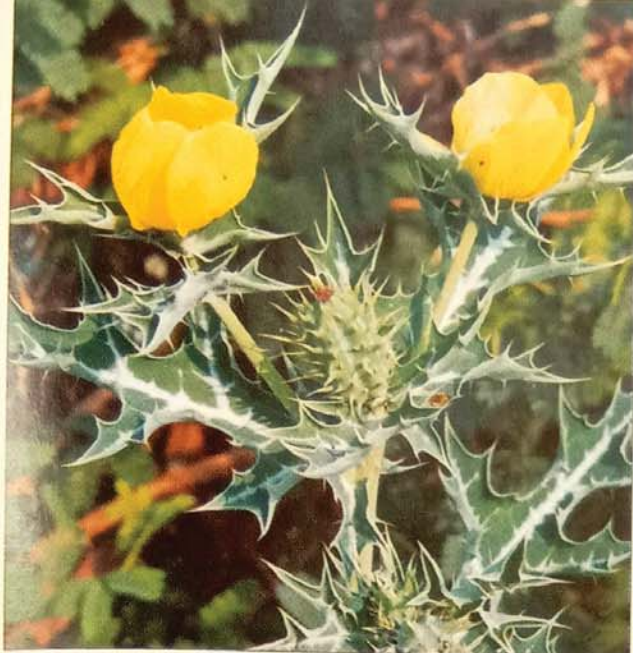
An irrigation reservoir which is known as 'Khangaon thadi' or Nandur Madhameshwar reservoir is situated near Khangaon thadi village, about 2 km away from Nandur Madhameshwar. Due to siltation, the reservoir is gradually becoming a shallow lake and it represents a sort of marshy ecosystem. It has three big islands in the middle and has an abundance of vegetation, fishes, mollusks and insects. Various species of *Cyperus*, *Typha*, *Amaranthus*, *Potamogeton*, *Ipomoea* and *Eichhornia* which are abundant on the islands provide excellent hiding and roosting places especially for different types of ducks. It is a paradise for many other birds; they feed upon plants and build their nests on them. Recent surveys have recorded 265 species of birds, 7 mammal species and 41 butterfly species in the area (Forest Dept., Nashik Div.). Due to constant efforts of local organizations, Bombay Natural History Society (BNHS), World Wide Fund for Nature-India (WWF-India) and the Forest Department, in 1978, this area was declared as a 'Protected Area' under the Wildlife (Protection) Act 1972 by the Maharashtra Government (Gaz. of Govt. of Maharashtra, 1978) and subsequently was declared as Nandur Madhameshwar Wildlife Sanctuary (Government of Maharashtra, Revenue and Forest Dept. Gazetteer, March 20, 1986, under the sub-sections (1) and (2) of Section 18 of the Wildlife (Protection) Act, 1972 (53 of 1972) (Appendix I page 29). The Nandur Madhameshwar Wildlife Sanctuary spans an area of 100.12 sq. km.

Two canals emanate from the Nandur Madhameshwar reservoir - the Godavari Left Bank Canal and the Godavari Right Bank Canal - with a total capacity of 7,763 m.c.ft. The reservoir irrigates a cultivable area of 88,000 Acres, which falls in Niphad and Yeola tehsils of Nashik district and Kopergaon tehsil of Ahmednagar district. The Godavari Right Bank Canal is approximately 111 km in length and irrigates a cultivable area of 1,36,380 Acres, falling in Niphad and Sinnar tehsils of Nashik district and Kopergaon and Shirampur tehsils of Ahmednagar districts.

Historical importance:

In the middle of the riverbed in between Khangaon thadi and Nandur Madhameshwar village, standing on a small rocky islet, is a 250 years old temple of 'Madhyameshwara' from which the village has derived its second half of its name. The lamp pillar near the temple bears an inscription dated 1738 with the name of an ascetic.

Agricultural importance : Niphad tehsil has an area of 1,05,228 ha, of which 90,631 ha is



Argemone mexicana L.

Family : Papavaraceae

Common Name : Pivala-Dhotra

Habitat : Common weed in waste land and in cultivated fields

Location : Khangaon thadi, Manjargaon

Fl. & Fr. : Throughout the year

Description : Annual, erect, prickly herb. Leaves are radical or cauline, variegated white, spiny on margins and veins, sessile. Flowers are yellow, axillary, solitary.

Uses : Entire plant is anti-fungal and also possess anti-leprotic activity.

Asparagus racemosus Willd.

Family : Liliaceae

Common Name : Shatavari

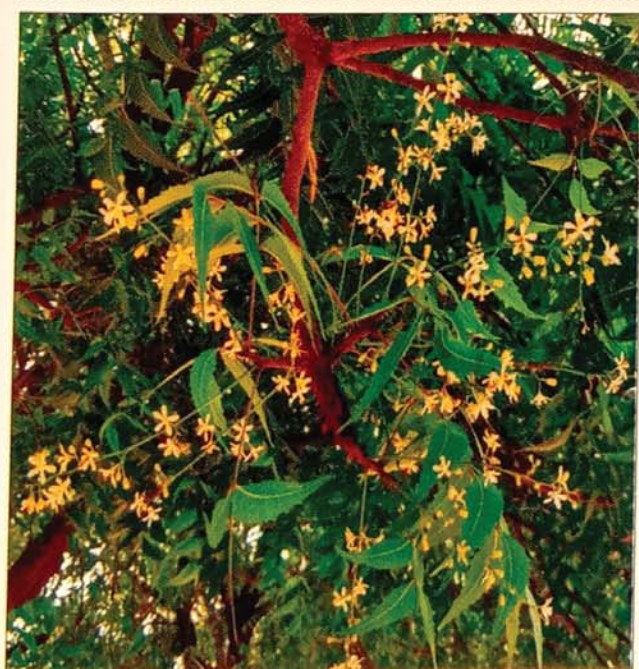
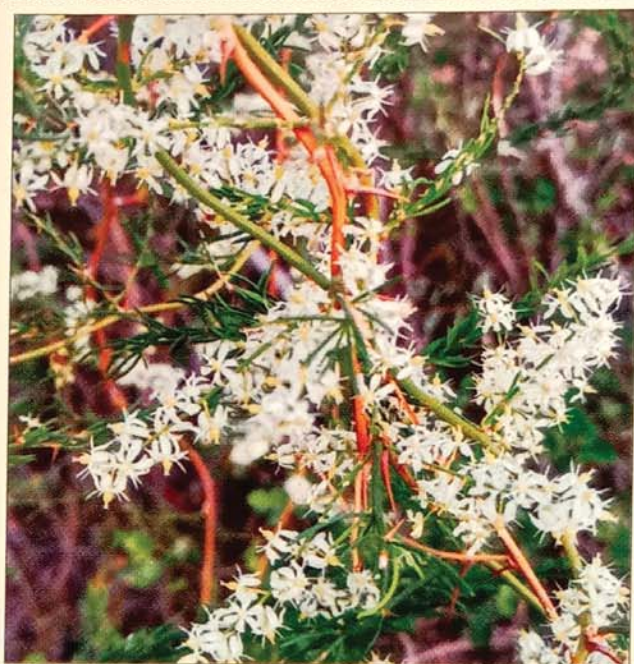
Habitat : Common along the hedges

Location : Khangaon thadi

Fl. & Fr. : June to October

Description : Shrubs with tuberous fascicled roots. Cladodes slender, glabrous. Flowers white in raceme.

Uses : Bark of plant show antibacterial activity and roots are used as galactagogue.



Azadirachta indica (L.) A. Juss.

Family : Meliaceae

Common Name : Kadunimb

Habitat : Commonly cultivated along the canal and road sides.

Location : Khangaon thadi, Manjargaon

Fl. & Fr. : March to June

Description : Trees with leaves crowded near end of branches. Leaflets ovate-lanceolate. Flowers white, in axillary panicles.

Uses : Flowers and Leaves are antibacterial and used as an analgesic.



Impatiens balsamina L.

Family : Balsaminaceae

Common Name : Terda

Habitat : Rare in horticultural land.

Location : Khangaon thadi

Fl. & Fr. : July to August

Description : Annual herbs with alternate, elliptic, acute leaves. Flowers pink, axillary, fascicled; lateral sepals ovate with short spur.

Uses : Plant possess anti-fungal and anti-cancer activity.

Indigofera cordifolia Heyne ex Roth

Family : Fabaceae

Common Name : Bechka

Habitat : Common along the river bed

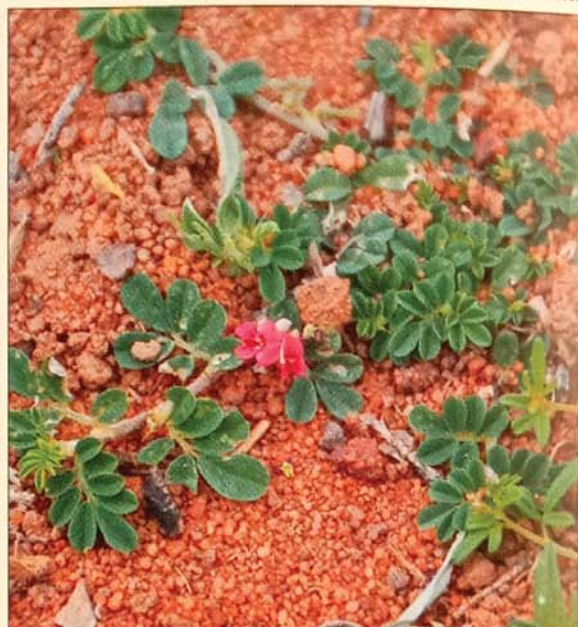
Location : Khangaon thadi

Fl. & Fr. : Throughout the year

Description : Herbs, prostrate, pilose. Leaves are ovate, acute, cordate, pilose on both surfaces. Flowers are red, 4-6 in number, in condensed racemes, tomentose.

Uses : Seeds are aphrodisiac and used as bitter tonic.

Photo : Dr. Mayur Nandikar



Indigofera linifolia (L.f.) Retz.

Family : Fabaceae

Common Name : Lal Godhadi

Habitat : Common along the river bed.

Location : Khangaon thadi, Manjargaon

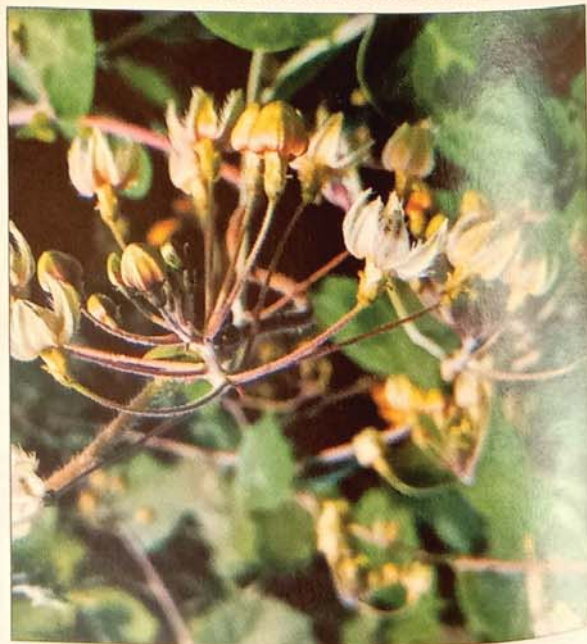
Fl. & Fr. : July to December

Description : Herbs, prostrate, much branched, pubescent. Leaves simple, linear, acute, pubescent on both sides. Flowers bright red, in sessile or shortly peduncled axillary racemes.

Uses : Seed oil is anti-microbial and nutritive tonic.

***Pergularia daemia* (Forssk.) Chiov**

Family: Asclepidaceae
Common Name: Utarn
Habitat: Common along the hedges
Location: Khangaon thadi, Manjargaon, Tarul-Khedale
Fl. & Fr.: March to December
Description: Herbs, twining, with milky sap. Leaves suborbicular with cordate base. Flowers in cymes, pubescent.
Uses: Leaf juice is emetic and used against snake bite.



***Persicaria glabra* (Willd.) M. Gomez**

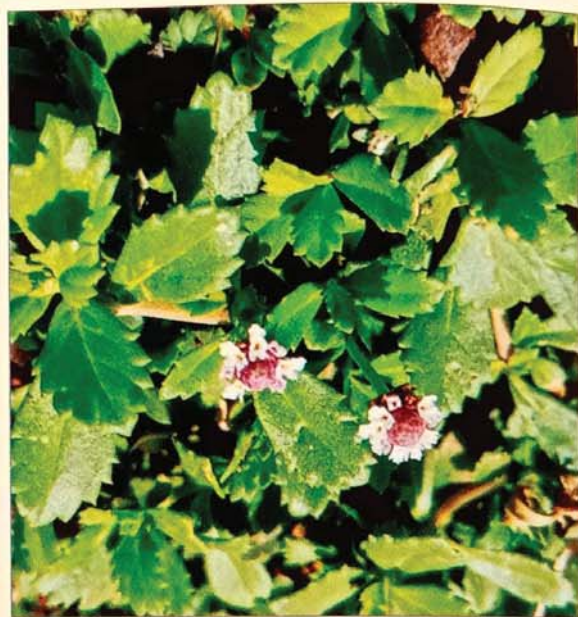
Family: Polygonaceae
Common Name: Sheral
Habitat: Common along the river banks
Location: Madhameshwar, Manjargaon
Fl. & Fr.: October to March
Description: Herbs, stem thick, reddish after drying. Leaves lanceolate, acute or acuminate at apex, entire, glabrous. Flowers pink, in terminal racemes.
Uses: Leaves are used in colic pain and as febrifuge.

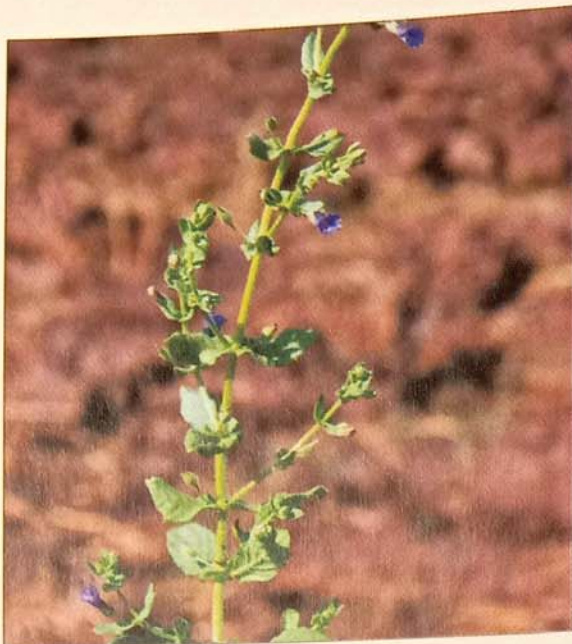


***Phyla nodiflora* (L.) Greene**

Family: Verbenaceae
Common Name: Jalpimpili
Habitat: Common in dried cultivated lands

Location: Khangaon thadi
Fl. & Fr.: Throughout the year
Description: Herbs, prostrate, with woody root stocks, rooting at the nodes, pubescent. Leaves oblanceolate-ovate, subsessile to sessile. Flowers purple or white, solitary, axillary or in spikes.
Uses: Infusion of leaves and stalks are given to women after delivery and also plant is used for joint pain.





Stemodia viscosa Roxb.

Family : Scrophulariaceae

Common Name : Satmodi

Habitat : Common along the river banks and in moist places

Location : Khangaon thadi, Manjargaon, Madhameshwar

Fl. & Fr. : November to January

Description : Herbs, erect, aromatic; stem and branches are viscidly pubescent. Leaves sessile, oblong, acute, serrate, pubescent. Flowers purple, solitary, axillary or in terminal racemes.

Uses : Infusion of plant is used as demulcent.

Striga angustifolia (D. Don) Saldanha

Family : Scrophulariaceae

Habitat : Rare in grasslands

Location : Madhameshwar

Fl. & Fr. : July to October

Description : Much branched stout herb. Leaves linear, sessile, scabrous. Flowers sessile or shortly pedicellate, in long, erect spikes or racemes.

Uses : Plant is used in diabetes



Syzygium cumini (L.) Skeels

Family : Myrtaceae

Common Name : Jambhul

Habitat : Common; cultivated in the farms and gardens for its edible fruits

Location : Khangaon thadi

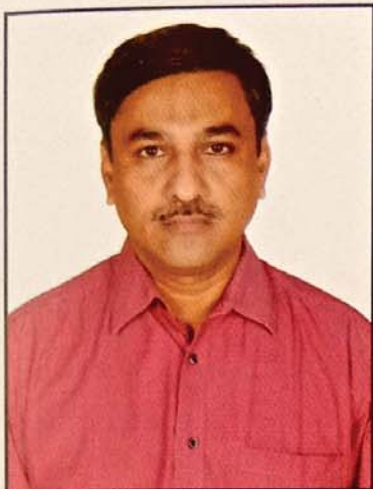
Fl. & Fr. : March to May

Description : Trees, Leaves lanceolate, elliptic-oblong or broadly ovate-elliptic. Flowers dirty white, in paniced cymes.

Uses : Fruit is used in diarrhoea and dysentery.



A red Coloured mat of *Azolla pinnata* R. Br., an aquatic free-floating fern on the water surface and *Typha angustifolia* L. in the background.



Dr Rajendra D. Shinde
M.Sc., Ph.D.

Dr Rajendra D. Shinde is the Head, Department of Botany & the Director, Blatter Herbarium at St. Xavier's College (Autonomous), Mumbai.

He is the member of the "Plant Biodiversity Expert Committee", Maharashtra State Biodiversity Board since 2017. He has served the St. Xavier's College in the capacity of Vice-Principal from 2010 to 2017. He is Elected Fellow of the Indian Association Angiosperm Taxonomy, Life Member of Society of Ethnobotany, Association of Plant Taxonomy, Indian Botanical Society, Bombay Natural History Society, Alumni of the Fulbright-Nehru Programme (2012), Rotary Foundation-Rotary International: GSE programme (2000), Nominated Member of Tree Authority - Thane Municipal

Corporation (2014-2017). Angiosperm Taxonomist by specialization, he has been teaching Botany at St. Xavier's College, Mumbai since 1991. He has also served as a Curator of the Blatter Herbarium from 1983 to 1991. During the year 2003-2004, Dr Shinde had an opportunity to serve as a senior lecturer at the Faculty of Natural Sciences, University of Guyana, Georgetown, Guyana (South America). Systematic and ecological studies on the Nandur Madhmeshwar, Nashik District, Maharashtra (1988), Arboreal Flora of Greater Bombay (1993), Tree Census of Greater Bombay (1998), Tree Census of Thane Municipal Corporation (2002), Digitized Inventory of Medicinal Plants Resources of Maharashtra (2009-2013) are some of the major projects completed by him along with five minor research projects from various funding agencies.

Besides several research papers in reputed and peer-reviewed journals, he has authored a book entitled "Ethno medicinal plants of Raigad District, Maharashtra (2016). He is a well-known research guide in the field of Angiosperm Taxonomy and Ethnobotany and got 4 PhD awarded under his guidance so far.

During his Master's study at St. Xavier's College, along with Late Dr. M.R. Almeida, he had visited Nandur Madhameshwar during 1984-1988 regularly to study the flora of the area. He updated this work in the year 2012-2015. He has reported 536 plant species from the Nandur Madhameshwar Wildlife Sanctuary during this study. This book is the outcome of this study.

A Study of Retention and Motivational Practices in Banks with Special Reference to Mumbai City

Editors

Dr. Suvaiba I. Shirshikar

Dr. Megha Somani



Himalaya Publishing House

ISO 9001:2008 CERTIFIED

A STUDY OF RETENTION AND MOTIVATIONAL PRACTICES IN BANKS WITH SPECIAL REFERENCE TO MUMBAI CITY

EDITORS

DR. SUVAIBA INAYAT SHIRSHIKAR

*M.Com., Ph.D., UGC-NET
Assistant Professor,
Department of Commerce,
St. Xavier's College – Autonomous
Mumbai - 400 001.*

DR. MEGHA SOMANI

*M.Com., M.M.M., Ph.D., UGC-SET
Assistant Professor,
Department of Commerce,
M.M.K. College of Commerce and Economics,
Mumbai - 400 050.*

First Edition : 2017



Himalaya Publishing House

ISO 9001:2008 CERTIFIED

© **Authors**

No part of this publication may be reproduced, stored in a retrieval system, or transmitted in any form or by any means, electronic, mechanical, photocopying, recording and/or otherwise without the prior written permission of the publisher.

First Edition : 2017

Published by : Mrs. Meena Pandey for **Himalaya Publishing House Pvt. Ltd.**,
"Ramdoot", Dr. Bhalerao Marg, Girgaon, Mumbai - 400 004.
Phone: 022-23860170/23863863, Fax: 022-23877178
E-mail: himpub@vsnl.com; Website: www.himpub.com

Branch Offices :

New Delhi : "Pooja Apartments", 4-B, Murari Lal Street, Ansari Road, Darya
Ganj, New Delhi - 110 002.
Phone: 011-23270392, 23278631; Fax: 011-23256286

Nagpur : Kundanlal Chandak Industrial Estate, Ghat Road,
Nagpur - 440 018.
Phone: 0712-2738731, 3296733; Telefax: 0712-2721216

Bengaluru : Plot No. 91-33, 2nd Main Road, Seshadripuram,
Behind Nataraja Theatre, Bengaluru - 560020.
Phone: 08041138821, Mobile: 09379847017, 09379847005.

Hyderabad : No. 3-4-184, Lingampally, Besides Raghavendra Swamy
Matham, Kachiguda, Hyderabad - 500 027.
Phone: 040-27560041, 27550139

Chennai : New No. 48/2, Old No. 28/2, Ground Floor, Sarangapani Street,
T. Nagar, Chennai - 600 012. Mobile: 09380460419

Pune : First Floor, "Laksha" Apartment, No. 527, Mehunpura,
Shaniwarpet (Near Prabhat Theatre), Pune - 411 030.
Phone: 020-24496323/24496333; Mobile: 09370579333

Lucknow : House No. 731, Shekhupura Colony, Near B.D. Convent
School, Aliganj, Lucknow - 226 022.
Phone: 0522-4012353; Mobile: 09307501549

Ahmedabad : 114, "SHAIL", 1st Floor, Opp. Madhu Sudan House, C.G. Road,
Navrang Pura, Ahmedabad - 380 009.
Phone: 079-26560126; Mobile: 09377088847

Ernakulam : 39/176 (New No.: 60/251) 1st Floor, Karikkamuri Road,
Ernakulam, Kochi - 682011.
Phone: 0484-2378012, 2378016 Mobile: 09387122121

Bhubaneswar : 5 Station Square, Bhubaneswar - 751 001 (Odisha).
Phone: 0674-2532129, Mobile: 09338746007

Kolkata : 108/4, Beliaghata Main Road, Near ID Hospital, Opp. SBI Bank,
Kolkata - 700 010,
Phone: 033-32449649, Mobile: 07439040301

DTP by : Prerana Enterprises, Mumbai.

Printed at : M/s. Aditya Offset Process (I) Pvt. Ltd., Hyderabad.
On behalf of HPH.

ABOUT THE AUTHORS



Dr. Suvaiba Shirshikar has done her Masters and Bachelors from Sydenham College of Commerce and Economics - University of Mumbai. She later cleared her NET and successfully completed Ph.D under the guideship of Dr. Megha Somani. This book is the part of her thesis work. She has 7 years of teaching experience and 2 years of corporate experience as well.



Dr. Megha Somani has done her double masters and has cleared SET. She is holding a Ph.D and is recognised guide from Mumbai University. She has given guidance to several students till now. She has more than 10 years of approved teaching experience in both graduate and post graduate level.

www.himpub.com

ISBN: 978-93-5273-492-4



9 789352 734924

ISBN: 978-93-5273-492-4

HBK 123

₹ 895/- \$ 36

शशि-भारती

पाठ्य-पुस्तक

(F. Y. B. A.)



संपादिका

डॉ. आशा नैथानी दायमा

शशि-भारती

पाठ्य-पुस्तक
(F.Y.B.A.)

संपादिका

डॉ. आशा नैथानी दायमा



परिदृश्य प्रकाशन

मुंबई

हिन्दी में किताबों का श्रेष्ठ चयन

ISBN : 81-86869-99-9

प्रथम संस्करण : 2017

© डॉ. आशा नैथानी दायमा

अक्षर संयोजन एवं आवरण
क्रिएटिव इमेज, मुंबई

E-mail : creativeimage24@gmail.com

प्रकाशक :

मूल्य : ₹ 125/-

परिदृश्य प्रकाशन

6, दादी संतुक लेन, धोबी तालाव,
मरीन लाईन्स, मुंबई-400002.

फोन : 022-6452 6072 / 2206 8040

E-mail : hindikitab@gmail.com

Shashi Bharti : Dr. Asha Naithani Dayama

eISSN: 2213-3585

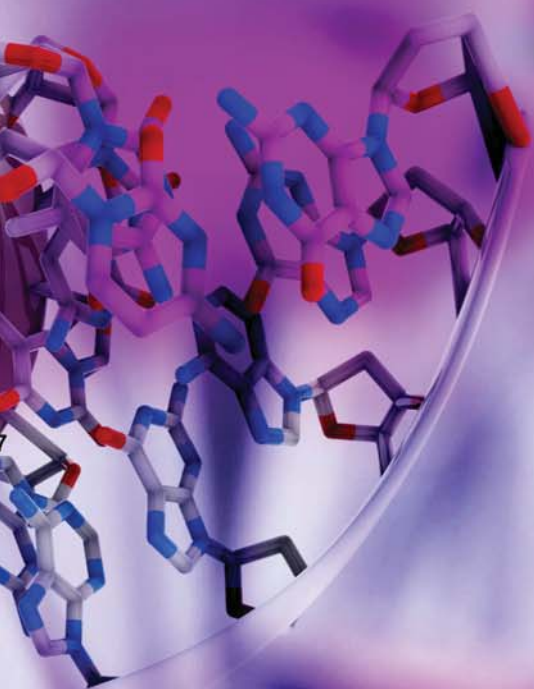
ISSN: 2468-5860

eISBN: 978-1-68108-455-8

ISBN: 978-1-68108-456-5

Volume 6

Topics in **Anti-Cancer Research**



Editors:

Atta-ur-Rahman, *FRS*

Khurshid Zaman

Bentham  Books

Patents eBook Series
“Topics in Anti-Cancer Research”

(Volume 6)

Edited by

Atta-ur-Rahman, *FRS*

Kings College University of Cambridge, Cambridge, UK

&

Khurshid Zaman

Bentham Science Publishers, USA

Topics in Anti-Cancer Research

Volume # 6

Editors: Atta-ur-Rahman, *FRS* and Khurshid Zaman

ISSN (Print): 2468-5860

ISSN (Online): 2213-3585

ISBN (Online): 978-1-68108-455-8

ISBN (Print): 978-1-68108-456-5

©2017, Bentham eBooks imprint.

Published by Bentham Science Publishers – Sharjah, UAE. All Rights Reserved.

BENTHAM SCIENCE PUBLISHERS LTD

End User License Agreement (for non-institutional, personal use)

This is an agreement between you and Bentham Science Publishers Ltd. Please read this License Agreement carefully before using the ebook/echapter/ejournal (“**Work**”). Your use of the Work constitutes your agreement to the terms and conditions set forth in this License Agreement. If you do not agree to these terms and conditions then you should not use the Work.

Bentham Science Publishers agrees to grant you a non-exclusive, non-transferable limited license to use the Work subject to and in accordance with the following terms and conditions. This License Agreement is for non-library, personal use only. For a library / institutional / multi user license in respect of the Work, please contact: permission@benthamscience.org.

Usage Rules:

1. All rights reserved: The Work is the subject of copyright and Bentham Science Publishers either owns the Work (and the copyright in it) or is licensed to distribute the Work. You shall not copy, reproduce, modify, remove, delete, augment, add to, publish, transmit, sell, resell, create derivative works from, or in any way exploit the Work or make the Work available for others to do any of the same, in any form or by any means, in whole or in part, in each case without the prior written permission of Bentham Science Publishers, unless stated otherwise in this License Agreement.
2. You may download a copy of the Work on one occasion to one personal computer (including tablet, laptop, desktop, or other such devices). You may make one back-up copy of the Work to avoid losing it. The following DRM (Digital Rights Management) policy may also be applicable to the Work at Bentham Science Publishers’ election, acting in its sole discretion:
 - 25 ‘copy’ commands can be executed every 7 days in respect of the Work. The text selected for copying cannot extend to more than a single page. Each time a text ‘copy’ command is executed, irrespective of whether the text selection is made from within one page or from separate pages, it will be considered as a separate / individual ‘copy’ command.
 - 25 pages only from the Work can be printed every 7 days.
3. The unauthorised use or distribution of copyrighted or other proprietary content is illegal and could subject you to liability for substantial money damages. You will be liable for any damage resulting from your misuse of the Work or any violation of this License Agreement, including any infringement by you of copyrights or proprietary rights.

Disclaimer:

Bentham Science Publishers does not guarantee that the information in the Work is error-free, or warrant that it will meet your requirements or that access to the Work will be uninterrupted or error-free. The Work is provided "as is" without warranty of any kind, either express or implied or statutory, including, without limitation, implied warranties of merchantability and fitness for a particular purpose. The entire risk as to the results and performance of the Work is assumed by you. No responsibility is assumed by Bentham Science Publishers, its staff, editors and/or authors for any injury and/or damage to persons or property as a matter of products liability, negligence or otherwise, or from any use or operation of any methods, products instruction, advertisements or ideas contained in the Work.

Limitation of Liability:

In no event will Bentham Science Publishers, its staff, editors and/or authors, be liable for any damages, including, without limitation, special, incidental and/or consequential damages and/or damages for lost data and/or profits arising out of (whether directly or indirectly) the use or inability to use the Work. The entire liability of Bentham Science Publishers shall be limited to the amount actually paid by you for the Work.

General:

1. Any dispute or claim arising out of or in connection with this License Agreement or the Work (including non-contractual disputes or claims) will be governed by and construed in accordance with the laws of the U.A.E. as applied in the Emirate of Dubai. Each party agrees that the courts of the Emirate of Dubai shall have exclusive jurisdiction to settle any dispute or claim arising out of or in connection with this License Agreement or the Work (including non-contractual disputes or claims).
2. Your rights under this License Agreement will automatically terminate without notice and without the need for a court order if at any point you breach any terms of this License Agreement. In no event will any delay or failure by Bentham Science Publishers in enforcing your compliance with this License Agreement constitute a waiver of any of its rights.
3. You acknowledge that you have read this License Agreement, and agree to be bound by its terms and conditions. To the extent that any other terms and conditions presented on any website of Bentham Science Publishers conflict with, or are inconsistent with, the terms and conditions set out in this License Agreement, you acknowledge that the terms and conditions set out in this License Agreement shall prevail.

Bentham Science Publishers Ltd.

Executive Suite Y - 2

PO Box 7917, Saif Zone

Sharjah, U.A.E.

Email: subscriptions@benthamscience.org



CONTENTS

FOREWORD	i
INTRODUCTION	ii
LIST OF CONTRIBUTORS	iii
CHAPTER 1 TARGETING POLYUNSATURATED FATTY ACID METABOLISM IN COLORECTAL CANCER THERAPY: A REVIEW OF RECENT PATENTS	1
<i>Arkadiusz Michalak, Paula Mosińska, and Jakub Fichna</i>	
1. INTRODUCTION	2
2. PUFAS AND THEIR METABOLITES	3
3. PRIMARY PREVENTION	12
4. TREATMENT	13
5. NOVEL TECHNOLOGIES IN CANCER THERAPY	18
6. SUPPLEMENTARY TREATMENT	19
7. ADJUVANT TREATMENT IN CHEMO- OR RADIOTHERAPY	20
8. FORMULATIONS OF PUFAS	21
CURRENT & FUTURE DEVELOPMENTS	22
CONFLICT OF INTEREST	23
AUTHOR CONTRIBUTIONS	23
ACKNOWLEDGEMENTS	24
LIST OF ABBREVIATIONS	24
REFERENCES	25
CHAPTER 2 MICROTUBULES AS ANTI-CANCER DRUG TARGETS	31
<i>Biswa P. Chatterji, Needa N. Bagban and Riddhi I. Bhavsar</i>	
1. INTRODUCTION	32
1.1. Cancer	32
1.2. Hallmarks of Cancer	33
2. METHODS TO TREAT CANCER	34
2.1. Chemotherapy	34
2.2. Hormonal Therapy	35
2.3. Radiation Therapy	35
2.4. Adjuvant Therapy	35
2.5. Immunotherapy	35
2.6. Targeted Cancer Treatments	36
2.6.1. Growth Signal Inhibitors	36
2.6.2. Apoptosis Inducing Drugs	36
2.6.3. Endogenous Angioinhibitors	36
3. MICROTUBULES AS DRUG TARGETS	36
3.1. Microtubule Dynamics in Mitosis	39
4. MICROTUBULE DESTABILIZING AGENTS	39
4.1. Vinca Site-Binding Agents	39
5. AGENTS THAT BIND AT THE COLCHICINE SITE	44
6. MICROTUBULE STABILIZERS	46
6.1. Agents that Bind at the Taxane Site	46
6.2. Epothilones	49
6.3. Taccalonolides	50
7. PATENTS ON NOVEL AGENTS THAT TARGET MICROTUBULES	50
7.1. Aurora A	50
7.1.1. US Patent 9012475 by Hirai and Sootome (2015)	50

7.2. Synergistic Drug Combinations for Cancer Therapy	51
7.2.1. US Patent Application 20150140125 by Zaid et al. (2015)	51
7.3. Pentoxifylline	52
7.3.1. US Patent Application 20150374702 by Kuh and Kim (2015)	52
7.4. Benzimidazole Derivatives	53
7.4.1. US Patent Application 20160002214 by Ahmed et al. (2016)	53
7.5. Combination of Casein Kinase 2 (CK2) Peptide Inhibitor (Termed P15) with Standard Chemotherapeutic Agents	54
7.5.1. US Patent 9278118 by Rodriguez et al. (2016)	54
7.6. Curcumin	56
7.6.1. WO Patent Application WO2015067282 by Fawzy (2015)	56
7.7. Anticancer Drug and/ or Therapeutically Active Molecule	57
7.7.1. WO Patent Application WO2015193702 by Lavitrano et al. (2015)	57
7.8. Pharmaceutical Compositions Comprising Polymeric Nanoparticles of Anticancer Drugs	59
7.8.1. WO Patent Application WO2016020697 by Shrikhande et al. (2016)	59
7.9. Tubulin inhibitors	60
7.9.1. US Patent 8883793 by Chen et al. (2014)	60
7.10. Bicyclic and Tricyclic Pyrimidine Tyrosine Kinase Inhibitors with Antitubulin Activity	61
7.10.1. US Patent 9139590 by Gangjee (2015)	61
7.11. Combretastatin A-1 Phosphate and Combretastatin B-1 Phosphate Prodrugs.	62
7.11.1. US Patent 7078552 by Pettit et al. (2006)	62
7.12. Combretastatin Analogs	64
7.12.1. US Patent 7429681 by Pinney et al. (2008)	64
CONCLUSION	66
FUTURE OF CANCER TREATMENT	66
Anti-Angiogenic Chemotherapy	67
Other Targeted Treatments	67
Nanotechnology	67
RNA Expression Profiling	67
LIST OF ABBREVIATIONS	67
CONFLICT OF INTEREST	68
ACKNOWLEDGEMENTS	68
REFERENCES	68
CHAPTER 3 EFFECTS OF INFLAMMATION, MITOCHONDRIA AND ENERGY METABOLISM IN THE HEART DUE TO CANCER	75
Thomas G. Ikonmidis, Timothy C. Tan and Patsie Polly	
1. INTRODUCTION	76
2. INFLAMMATORY CYTOKINES AND MITOCHONDRIAL DYSFUNCTION IN CARDIAC TISSUE	76
2.1. Tumour Necrosis Factor- α	76
2.2. Interleukin-6 and STAT3 Effects	79
3. MITOCHONDRIAL DYSFUNCTION AND CANCER CACHEXIA	80
4. CANCER CACHEXIA-INDUCED CARDIOMYOCYTE AUTOPHAGY AND MITOPHAGY	84
CURRENT & FUTURE DEVELOPMENTS	86
CONFLICT OF INTEREST	86
ACKNOWLEDGEMENTS	86
REFERENCES	86

CHAPTER 4 CHALCONE AND THEIR DERIVATIVES AS ANTICANCER AGENTS	93
<i>Zahoor A. Wani and Suaib Luqman</i>	
1. INTRODUCTION	94
2. CONJUGATES AND DERIVATIVES OF CHALCONES	94
3. CHALCONE AND CHALCONE DERIVATIVES AS TARGETED ANTICANCER AGENTS	99
3.1. Chalcones as Inhibitors of Nuclear Factor κ B (NF κ B) Signalling Pathway	99
3.2. Chalcones as Inhibitors of Histone Deacetylases (HDAC)	100
3.3. Chalcones as Angiogenesis Inhibitors	103
3.4. Chalcones as Inhibitors of Proteasome	104
3.5. Chalcones as Inhibitors of p53 Degradation	104
4. STRUCTURE ACTIVITY RELATIONSHIP (SAR) OF CHALCONES	104
CURRENT & FUTURE DEVELOPMENTS	106
CONFLICT OF INTEREST	107
ACKNOWLEDGEMENTS	107
REFERENCES	107
CHAPTER 5 REGULATION/INHIBITION OF HUMAN LACTATE DEHYDROGENASE A: AN INNOVATIVE AND POTENTIAL APPROACH FOR ANTI-CANCER DRUGS DEVELOPMENT	114
<i>Vinit Kumar, Atul Kumar and Reshma Rani</i>	
1. INTRODUCTION	115
2. CANCER CELL METABOLISM	115
3. GLYCOLYSIS AND LDHA	116
4. ISOFORM OF LDH	118
5. LDH ACTIVE SITE	118
5. LDH CATALYTIC MECHANISM	120
6. HLDH-A AND ITS LINK WITH TUMORIGENESIS	122
7. REGULATION/INHIBITION OF LACTATE DEHYDROGENASE	124
8. SMALL MOLECULES AS HLDHA INHIBITORS	125
9. SYNTHETIC LDHA INHIBITORS	126
10. NANOPARTICLE BASED DELIVERY FOR HLDH-A INHIBITION	131
11. LDH IN OTHER DISEASES	131
12. DETERMINATION AND QUANTIFICATION OF LDH	131
CURRENT AND FUTURE DEVELOPMENTS	135
CONFLICT OF INTEREST	136
ACKNOWLEDGEMENTS	136
LIST OF ABBREVIATIONS	136
REFERENCES	137
CHAPTER 6 CANCER CHEMO-IMMUNOTHERAPEUTICS	143
<i>Muzammal Hussain and Jiancun Zhang</i>	
1. INTRODUCTION	143
2. CURRENT CHEMO-IMMUNOTHERAPEUTIC DRUG COMBINATIONS IN CLINICAL STUDIES	144
2.1. Chemotherapy-mAbs Combinations	147
2.2. Chemotherapy-Immune Checkpoint Inhibitor Combinations	148
2.3. Chemotherapy-Vaccine Combinations	149
2.4. Chemotherapy-Cytokine Combinations	151
2.5. Chemotherapy-CIK Cells Combinations	152
2.6. Chemotherapy-Oncolytic Immunotherapy Combinations	152

3. TUMOURAL INHIBITION OF HOST-TUMOUR CELL INTERACTIONS: MAPPING THE ROAD AHEAD TO ‘TWO HIT’ CHEMO-IMMUNOTHERAPEUTICS	153
3.1. ‘Two Hit’ Chemo-Immunotherapeutic Potential of Oncogenic Kinase Signaling Inhibition	154
3.2. Does Tumoural Inhibition of ARG and IDO Possess ‘Two Hit’ Chemo-Immunotherapeutic Potential?	157
4. PATENTS ISSUED AND RECENT PATENT APPLICATIONS IN CANCER CHEMO-IMMUNOTHERAPEUTICS	159
CURRENT AND FUTURE DEVELOPMENTS	163
CONFLICT OF INTEREST	164
ACKNOWLEDGEMENTS	164
REFERENCES	164
CHAPTER 7 RECENT ADVANCES AND CHALLENGES IN MICRORNA-BASED CANCER THERAPEUTICS	178
<i>Amjad Ali, Faryal M. Awan, Aqsa Ikram □□Shifa T. Ashraf</i>	
1. INTRODUCTION	179
2. miRNA BIOGENESIS	181
3. STRATEGIES FOR miRNA MANIPULATION	184
3.1. Sandwich RNAi Inhibition Strategy	184
3.2. Multiplex RNAi Inhibition Strategy	184
3.3. Small Molecules Inhibitors of miRNAs	185
4. THE POTENTIAL OF miRNAs IN CANCER THERAPY: SAFETY AND TOXICITY	186
5. miRNAs IN TUMOR GROWTH AND METASTASIS	186
6. SIGNIFICANCE AND CONTRIBUTION OF BIOINFORMATICS PREDICTION IN miRNA-BASED CANCER THERAPEUTICS	187
7. APPROACHES USED IN miRNA BASED THERAPIES	189
8. miRNA MIMIC THERAPY	190
9. ANTI-miRNA THERAPY	193
10. RECENT PATENTS AND CLINICAL TRIAL STATUS OF miRNA THERAPEUTICS IN VARIOUS CANCERS	196
10.1. Targeting miRNAs miR-409-5p, miR-379 and miR-154* to Treat Prostate Cancer Bone Metastasis and Drug Resistant Lung Cancer	197
10.2. Use of Catenin- Beta 1-Targeting miRNAs for Treating Liver Cancer	198
10.3. Production and Extraction of MiRNA Precursor as Drug for Cancer Therapy	198
10.4. Targeting miRNAs for the Treatment of Liver Cancer	199
10.5. Pharmaceutical Composition for Treating Liver Diseases	200
10.6. Use of Glypican-3-Targeting miRNAs for Treating Liver Cancer	201
10.7. Sorafenib-miRNA Combination Therapy for Liver Cancer	202
CHALLENGES & FUTURE PERSPECTIVES	203
CURRENT & FUTURE DEVELOPMENTS	203
CONSENT FOR PUBLICATION	204
CONFLICT OF INTEREST	204
ACKNOWLEDGEMENTS	204
REFERENCES	204
AUTHOR INDEX	211
SUBJECT INDEX	21□

FOREWORD

The sixth volume of Topics in Anti-Cancer Research presents some exciting contributions in frontier areas of anti-cancer research. These include the role of microtubules for the treatment of various cancers, novel chemoimmunotherapy drug combinations & methods in clinical studies/trial and current studies in targeting polyunsaturated fatty acids (PUFAs) in the treatment of colorectal cancer. Natural and synthetic chalcones and their derivatives that have shown potent anticancer activity against a number of cancer cell lines and murine tumor models are discussed. The discovery of selective small-molecule hLDH-A inhibitors and LDH-based approaches in the progress of anticancer therapy are also presented. Recent advances in microRNA-based cancer therapeutics for the treatment of cancer are presented. The role of inflammation in chemotherapy-induced neuromuscular effects and the side effects and recent relevant patents for beneficial approaches to improve heart failure cases due to inflammation, mitochondria and energy metabolism in cancer cachexia are also covered. It is hoped that the present volume will be found useful by a large number of scientists working in this field.

The editors are thankful to the authors for their excellent contributions and to the reviewers for their in -depth comprehensive comments for the improvement of chapters. We are also grateful to Mr. Mahmood Alam, Mrs. Rafia Rehan and other colleagues for their support and assistance in the finalization of this volume.

Atta-ur-Rahman, FRS

International Center for Chemical and Biological Sciences
University of Karachi
Karachi 75270
Pakistan

Khurshid Zaman

Honorary Editor
Bentham Science Publishers

INTRODUCTION

Topics in Anti-Cancer Research covers important advances on both experimental (preclinical) and clinical cancer research in drug development. The book series offers readers an insight into current and future therapeutic approaches for the prevention of different types of cancers, synthesizing new anti-cancer agents, new patented compounds, targets and agents for cancer therapy as well as recent molecular and gene therapy research.

The comprehensive range of themes covered in each volume will be beneficial to clinicians, immunologists, and R&D experts looking for new anti-cancer targets and patents for the treatment of neoplasms, as well as varied approaches for cancer therapy.

The topics covered in the sixth volume of this series include:

- The role of microtubules for the cure of various untreated cancers
- Novel chemoimmunotherapeutic drug combinations & methods in clinical studies/trials
- Targeting polyunsaturated fatty acids (PUFAs) in the treatment of colorectal cancer
- Anti-cancer activity of natural and synthetic chalcones and their derivatives
- Recent advances in microRNA-based cancer therapeutics
- Treatment of heart failure due to inflammation, mitochondria and energy metabolism in cancer cachexia
- Regulation/inhibition of human lactate dehydrogenase A for discovering anti-cancer drugs

List of Contributors

Ali, Amjad	Atta-ur-Rahman School of Applied Biosciences (ASAB), National University of Sciences and Technology (NUST), H-12 Islamabad, Pakistan Email: amjaduni@gmail.com; amjad.ali@asab.nust.edu.pk
Ashraf, Shifa T.	Atta-ur-Rahman School of Applied Biosciences (ASAB), National University of Sciences and Technology (NUST), H-12 Islamabad, Pakistan Email: stariq.asab@gmail.com
Awan, Faryal M.	Atta-ur-Rahman School of Applied Biosciences (ASAB), National University of Sciences and Technology (NUST), H-12 Islamabad, Pakistan Email: faryal_mehwish@yahoo.com
Bagban, Needa N.	Department of Biotechnology, St. Xavier's College (Autonomous), Mumbai-400001, India Email: neebagban@gmail.com
Bhavsar, Riddhi I.	Department of Biotechnology, St. Xavier's College (Autonomous), Mumbai-400001, India Email: riddhibhavsar2@gmail.com
Chatterji, Biswa P.	Department of Biotechnology, St. Xavier's College (Autonomous), Mumbai-400001, India Email: biswaprasun@gmail.com
Fichna, Jakub	Department of Biochemistry, Faculty of Medicine, Medical University of Lodz, Lodz, Poland Email: jakub.fichna@umed.lodz.pl
Muzammal, Hussain	Institutes of Biomedicine and Health, Chinese Academy of Sciences, 190 Kaiyuan Avenue, Science Park, Guangzhou, 510530, P.R. China Email: muzammal@gibh.ac.cn
Ikonomidis, Thomas G.	Mechanisms of Disease and Translational Research Group, Department of Pathology, School of Medical Sciences, University of New South Wales, Sydney 2052, Australia Email: thomasikono@gmail.com
Ikram, Aqsa	Atta-ur-Rahman School of Applied Biosciences (ASAB), National University of Sciences and Technology (NUST), H-12 Islamabad, Pakistan Email: aqsa_ikram@yahoo.com
Kumar, Vinit	Amity Institute of Molecular Medicine and Stem Cell Research, Amity University, Noida 201313, India Email: vkumar25@amity.edu
Kumar, Atul	Amity Institute of Engineering & Technology, Amity University, Greater Noida 201313, India Email: atul206@gmail.com
Luqman, Suaib	CSIR-Central Institute of Medicinal and Aromatic Plants, Lucknow-226015, Uttar Pradesh, India Email: s.luqman@cimap.res.in

Michalak, Arkadiusz	Department of Biochemistry, Faculty of Medicine, Medical University of Lodz, Lodz, Poland Email: arkadiusz.michalak.lek@gmail.com
Mosińska, Paula	Department of Biochemistry, Faculty of Medicine, Medical University of Lodz, Lodz, Poland Email: paula.mosinska@gmail.com
Polly, Patsie	Mechanisms of Disease and Translational Research Group, Department of Pathology, School of Medical Sciences, University of New South Wales, Sydney 2052, Australia Email: patsie.polly@unsw.edu.au
Rani, Reshma	Amity Institute of Biotechnology, Amity University, Noida 201313, India Email: reshudcy@gmail.com; rrani@amity.edu
Tan, Timothy C.	Western Clinical School and Westmead Hospital, Westmead, New South Wales, Australia Email: timothy.tan9@gmail.com
Wani, Zahoor A.	National Postdoctoral Fellow at CSIR-Central Institute of Medicinal and Aromatic Plants, Lucknow-226015, Uttar Pradesh, India Email: zahoorwani5@gmail.com
Zhang, Jiancun	Institutes of Biomedicine and Health, Chinese Academy of Sciences, 190 Kaiyuan Avenue, Science Park, Guangzhou, 510530, P.R. China Email: zhang_jiancun@gibh.ac.cn

CHAPTER 1

Targeting Polyunsaturated Fatty Acid Metabolism in Colorectal Cancer Therapy: A Review of Recent Patents

Arkadiusz Michalak[#], Paula Mosińska^{*,#} and Jakub Fichna

Department of Biochemistry, Faculty of Medicine, Medical University of Lodz, Lodz, Poland

Abstract: In the recent years, fatty acids (FAs) have been acknowledged not only as building materials for lipid membranes and carbon source for β -oxidation, but also as important signaling molecules. In this field, polyunsaturated fatty acids (PUFAs) have received special attention as modulators of inflammation. The enzymes that process PUFAs into bioactive metabolites (cyclooxygenases, lipoxygenases) have already been targeted by pharmaceutical agents. Given the fact that intense synthesis of FAs is a metabolic hallmark of cancer, it is expected that FAs play an important role in cancer development, progression and invasion, and could be targeted by modern therapies. In this chapter, we will discuss the possible use of FAs and drugs affecting their metabolism against colorectal cancer (CRC), which is strongly associated with environmental factors such as high-fat, high caloric diet and obesity. We will cover the role of n-3 PUFAs as dietary supplements in primary prevention of CRC based on the results obtained from clinical trials, and elaborate on the latest patents designed to improve the bioavailability of PUFAs concentrates as nutritional treatments for patients with CRC. We will also discuss the enzymes processing PUFAs and their role in tumorigenesis with focus on their potential as markers for “molecular staging” (fatty acid synthases and elongases) and targets in therapy (cyclooxygenase 2 and lipoxygenase 5). Finally, we will examine new drug formulations (e.g. liposomes) and their utility in CRC therapy. The chapter is based on the review of literature (PubMed Database) and patent documents.

Keywords: Adjuvant therapy, chemotherapy, colorectal cancer, cyclooxygenase, dietary supplementation, docosahexaenoic acid, eicosapentaenoic acid, fatty acids, gastrointestinal cancer, inflammation, lipoxygenase, liposomes, nutritional treatment, polyunsaturated fatty acids, prevention, patents.

^{*} **Corresponding author Paula Mosińska:** Department of Biochemistry, Faculty of Medicine, Medical University of Lodz, Mazowiecka 6/8, 92-215 Lodz, Poland Tel: ++48 42 272 57 07; Fax: ++48 42 272 56 94; E-mail: paula.mosinska@gmail.com

[#] Equal contribution

1. INTRODUCTION

Colorectal cancer (CRC) is the second most common cancer in women and third in men, responsible for 600,000 deaths annually worldwide [1 - 3]. It is the fourth cause of oncological deaths, which creates a substantial global burden [4]. Up to 50% of CRC risk is lifestyle-related - most prominent risk factors include obesity, sedentary behavior, alcohol consumption, tobacco smoking, high-meat / high-calorie intake, as well as fat-rich and fiber-deficient diet [5]. All of these disturb the metabolic balance and add to CRC development. A cause-effect relation has been proven for alcohol (which promotes folate deficiency and thus leads to DNA instability and carcinogenesis) and tobacco smoking (which spreads carcinogens from cigarettes to colorectal mucosa, stimulating carcinogenesis) [5]. In turn, dietary habits and sedentary lifestyle not only cause obesity but also lead to the development of metabolic syndrome highlighted by a range of abnormalities encompassing impaired glucose tolerance, elevated blood pressure and dyslipidemia. These metabolic disorders tip the cytokine balance toward chronic low-grade inflammation and further disturb the levels of adipokines e.g. adiponectin and leptin, and insulin growth factors which all affect cellular proliferation, adhesion and migration [6 - 8]. Moreover, unbalanced diet can directly promote carcinogenesis by modifying the intestinal microbiome and making alterations in the complexity of the colorectal mucosa - for details see [6].

Alterations in lifestyle patterns through higher intake of fish and fish oils, dietary fiber, vitamin D and calcium, regular use of aspirin and habitual physical exercise modulate the course of CRC, especially at the initial stage of its development, and improve the quality of life of patients [5]. The protective role of fish and fish oils is mainly attributed to the high content of polyunsaturated fatty acids (PUFAs). The fact that aspirin also acts on the metabolism of PUFAs further suggests that these fatty acids may play a significant role in CRC development and possible prevention.

PUFAs are organic acids comprising of a carbohydrate chain with more than one double (C=C) bond in their structure. Long-chain PUFAs are divided into n-6 PUFAs (first double bond at C6, counting from the methyl C) and n-3 PUFAs (first unsaturated bond at C3). The main representatives of these groups are linoleic acid (LA, 18:2) for n-6 PUFAs and α -linolenic acid (ALA, 18:3) for n-3 PUFAs, together called essential fatty acids (FAs). The term "essential" emphasizes their importance in maintaining the optimal health of humans and other animals, as they cannot be synthesized *de novo* but have to be supplemented in the diet. These FAs provide the carbon chain necessary for the synthesis of longer FAs: n-6 arachidonic acid (AA, 20:4), and n-3 eicosapentaenoic acid (EPA, 19:5) and docosahexaenoic acid (DHA, 22:6) in the reactions catalyzed by

elongases and desaturases. In humans, the efficacy of transforming ALA to longer n-3 PUFAs is low and personally variable [9] and thus its derivatives should also be supplemented in diet. Animal-derived products (meat, eggs, dairy) are the most common source of LA and its derivative AA, whereas fish, particularly salmon, provides mainly n-3 PUFAs.

This chapter will briefly describe the fundamental knowledge of PUFAs and their metabolism. A detailed section is devoted to reports from the *in vitro* and *in vivo* studies investigating links between PUFAs and CRC. The main body covers various ways in which PUFAs could be utilized to prevent or treat cancer, especially CRC, based on the already established patents and promising reports from the literature.

The review is based on literature search conducted in the following databases: PubMed (for original papers and reviews), ClinicalTrials.gov, EU Clinical Trials Register and UMIM (for clinical trials), and WIPO (for pertaining to patents). The keywords used to search for patents included: adjuvant therapy, chemotherapy, colorectal cancer, dietary supplementation, docosahexaenoic acid, eicosapentaenoic acid, endocannabinoids, fish oil, liposomes, polyunsaturated fatty acids and resolvins. The literature was searched in relation to relevant patents. Non-English articles were not included in the review. All patents and clinical trials mentioned in this paper are summarized in Tables 1 and 2, respectively.

2. PUFAS AND THEIR METABOLITES

PUFAs are important elements of cellular lipid membranes released into circulation by phospholipase A2. By undergoing various enzymatic and non-enzymatic pathways, PUFAs are converted into biologically active lipid metabolites and mediators (Fig. 1). The most prominent enzymes participating in the formation of bioactive metabolites of n-3 and n-6 PUFAs include:

- Cyclooxygenases (COXs) that produce prostaglandins (PGs), thromboxanes (TXs) and prostacyclins;
- Lipoxygenases (LOXs) which process AA into lipoxins (LXs) and leukotrienes (LTx), and n-3 PUFA into protectins, marensins and resolvins;
- Cytochrome 450 (Cyp 450) which converts PUFAs into hydroxyeicosatetraenoic acids (HETEs).

CHAPTER 2**Microtubules as Anti-Cancer Drug Targets****Biswa P. Chatterji*, Needa N. Bagban and Riddhi I. Bhavsar***Department of Biotechnology, St. Xavier's College (Autonomous), Mumbai- 400001, India*

Abstract: In developmental biology, all cellular events are suitably synchronized to ensure proper growth and development of any multicellular organism. A healthy adult tissue is often characterized by stem cells that can undergo orchestrated cell division and differentiation. Disruption of these events often leads to cancer resulting from the accretion or accumulation of the genetic and epigenetic changes that occur at the somatic as well as the germ line levels. In the recent years, significant progress has been made in the early detection, treatment and prevention of cancer. Targeted cancer therapies include the use of apoptosis inducing drugs and drugs that target microtubules among others. Over the past few years, drugs that inhibit microtubule dynamics have been successfully used as anticancer drugs. They can either be microtubule stabilizers (Vincristine, Vinblastine, Colchicine etc.) or microtubule destabilizers (Paclitaxel, Docetaxel, Etoposide, Taccalonolides etc.). Recently, new classes of compounds have been identified that interfere with cell growth and proliferation as a consequence of binding to tubulin $\alpha\beta$ - dimers. Natural compounds like Curcumin have shown to inhibit tubulin activity. Whereas some antimitotic agents like Aurora A/B, Pentoxifylline, benzimidazole derivatives, combrestatin, polymeric nanoparticles etc. have been reported to show significant effect in the treating several types of cancer which were previously deemed untreatable. The following chapter acknowledges the presence of these anti-tumor compounds and how they target microtubules and further aid in the treatment of various cancers afflicting human beings.

Keywords: Antimitotic agents, Aurora A, Apoptosis, Cancer, Combretastatin analogs, Colchicine, Curcumin, Docetaxel, Etoposide, Microtubules, Microtubule polymerizers, Microtubule depolymerizers, Paclitaxel, Pentoxifylline, Taccalonolides, Taxanes, Tubulin, Vinca alkaloids, Vinblastine, Etoposide, Taccalonolides, Microtubule destabilizers, Vinca alkaloids, Colchicine, Halichondrin B, Dolostatin 10, Aurora A, Curcumin, Pentoxifylline, Combretastatin, Benzimidazole derivatives, Casein K2 peptide inhibitor, Nanoparticles.

Corresponding author Biswa P. Chatterji: Department of Biotechnology, St. Xavier's College (Autonomous), Mumbai- 400001, India; Tel: +91-9920010371; E-mail: biswaprasun@gmail.com

Atta-ur-Rahman and Khurshid Zaman (Eds.)
All rights reserved-© 2017 Bentham Science Publishers

1. INTRODUCTION

1.1. Cancer

Cancer is the second leading cause of death in the world with the first position being conferred to cardiovascular diseases. The word cancer finds its origin from the Greek word *Karkinos* which was the term used by the Greek physician, Hippocrates (460-370 B.C) to describe carcinoma tumours. Within the developing embryo, events such as stem cell divisions, their fate determination, proliferation of cells and their migration followed by apoptosis need to be orchestrated. In an adult, the constant turnover of cells as well as their optimal functioning is ensured by the division and differentiation of the stem cells that are present in small numbers in the healthy tissues [1]. Cancer is thought to be the disruption of this critical organisation resulting from the accretion or accumulation of the genetic and epigenetic changes that occur at the somatic as well as the germ line levels [2]. This causes uncontrolled proliferation of normal cells and subsequently leads to the formation of cancerous cells which proceed to grow, proliferate and re-divide giving rise to abnormal cells instead of undergoing apoptosis. These cells have several features in common with stem cells. The observation of these similarities led to the advent of two alternative hypotheses, one stating that the stem cells might themselves be the targets of the mutations that transform them while the other suggesting that the dedifferentiation of those cells that are transformed and terminally differentiated gives rise to cancer stem cells (CSCs), thereby manifesting the disease [3]. Cells of different types of cancer migrate via the blood circulation or lymph vessels to the other regions of the body and begin to grow in that target region. This phenomenon is termed as “metastasis”. These cells are incapable of DNA repair and hence can be considered malignant (cancerous). However, some tumors do not possess the capacity to grow and migrate to the other parts of the body and are, therefore, categorized as benign (non-cancerous) [4].

Cancer is a multistage disease however; work carried out on cancer recognizes a link between chronic inflammation and cancer with Virchow (1863) hypothesising that cancer originates at the sites of chronic inflammation [4]. More advanced studies on cancer suggest that the inflammatory cells are actually involved in the promotion of cancer progression [5]. Most of the malignancies have been observed to be initiated by chronic inflammation or tissue injury, which can be associated to known parasitic, viral or bacterial infections [6].

Worldwide estimates about 15% of the malignancies (1.2 million/year) attributed to chronic infections instances of which include liver cancer caused due to Hepatitis B and C infection, cervical cancer caused due to human papilloma virus

and gastric cancer resulting from *Helicobacter pylori* (*H. pylori*) infection. Individuals susceptible to an increased risk of cancer exhibit increased polymorphisms in the genes encoding pro-inflammatory cytokines. Population-based studies have established that when tissues are chronically inflamed, the susceptibility to cancer increases and also the risk of many cancers reduces significantly as a consequence of long-term use of NSAIDs, thereby demonstrating the vital role of inflammation in the pathogenesis of cancer [7].

1.2. Hallmarks of Cancer

Advances in cancer research has gained new insights and it is thought to be a disease that involves dynamic changes in the genome; the basis of which has been built on the discovery of mutations that lead to the production of oncogenes and tumour suppressor genes. **Oncogenes** are the genes which have gained dominance over the function. They drive the normal cells towards unrestrained growth and develop into cancer cells. Proto-oncogenes are the normal genes of the cell that regulate the frequency of cell division as well as the extent of its differentiation. Oncogenes arise when mutations occur in the proto-oncogenes. **Tumor suppressor genes** are the normal genes that are involved in regulating cell division, DNA repair and signalling of apoptosis. Tumor suppressor genes experience recessive loss of function. Any dysfunction in a tumor suppressor gene results in an uncontrollable growth of cells thereby causing cancer [8].

Cancer cells possess impaired regulatory circuits that are responsible for normal cell proliferation and maintenance of homeostasis. More than 100 distinct types of cancer exist with many subtypes of tumors being found within specific organs. The vast catalog of cancer genotypes is thought to be manifested due to six of the crucial alterations in the physiology of cells which eventually lead to a malignant growth. The six alterations seen are as follows:

1. Loss of sensitivity to growth-inhibitory signals: Tissue homeostasis is maintained when cells respond to anti-proliferative signals during G1 phase. However cancerous cells become insensitive to such signals due to the disruption of retinoblastoma proteins which helps in filtering anti proliferative signals.
2. Evasion from apoptosis: Cancerous cells grow in number not only because they become proliferative but also because they tend to evade cell death mechanisms.
3. Self- sufficiency in growth signals: Once a normal cell gets transformed into a cancerous cell, their dependency on exogenous stimulatory growth factor is scaled down. This is due to the fact that the oncogenes tend to mimic the growth signals in one form or the other.

CHAPTER 3

Effects of Inflammation, Mitochondria and Energy Metabolism in the Heart due to Cancer

Thomas G. Ikonomidis¹, Timothy C. Tan^{1,2} and Patsie Polly^{1,*}

¹ Mechanisms of Disease and Translational Research Group, Department of Pathology, School of Medical Sciences, University of New South Wales, Sydney 2052, Australia

² Western Clinical School and Westmead Hospital, Westmead, New South Wales, Australia

Abstract: Cancer cachexia is a paraneoplastic syndrome characterised by significant skeletal muscle wasting and cardiac atrophy. It occurs in 50% of patients with cancer and approximately 20% of cancer deaths are attributed to cachexia. Heart failure due to cancer cachexia is suggested to contribute to the high mortality rate and currently there is limited therapeutic intervention. The relationship between inflammation and energy metabolism as well as mitochondrial dysfunction in the heart in the context of cancer cachexia will be discussed. This chapter provides an understanding of potential, novel molecular mechanisms that could be of interest when considering therapeutic interventions for heart failure due to cancer cachexia. In summary, several interrelated molecular effects should be considered in cancer-induced cachexia in cardiomyocytes. TNF- α induced mitochondrial dysfunction may be important for the generation of ROS. IL-6 may induce an autophagic/mitophagic response as a result of downregulation of mitochondrial STAT3 due to mTOR suppression. An imbalance in mitochondrial dynamics may contribute to insulin-resistance and atrophy. Decreased expression of ANT1 may contribute to MPTP dysfunction and an altered energetic profile from adult to fetal metabolism. The effects of ANT1 expression in cardiac muscle during cancer cachexia is worth investigating in mouse models as discussed with reference to an ANT1 patent in this chapter. Furthermore, patents that are relevant for therapeutic strategies to ameliorate heart failure in cancer cachexia have also been discussed. Patents addressing interventions that could be applied to cancer cachexia-induced cardiac atrophy include: sodium selenite treatment, inhibitory agents of NADPH oxidase such as phycobilin, an AMPK inhibitor, modulation of mitochondrial biogenesis and modulation of mTOR. Understanding the underlying molecular mechanisms of mitochondrial dysfunction in cardiomyocytes during cancer cachexia-induced cardiac atrophy may reveal novel molecular targets for therapeutic intervention.

Keywords: Cancer, cancer cachexia, cardiac atrophy, cardiomyocyte, energy metabolism, heart failure, inflammation, inflammatory cytokines, mitochondria,

* Corresponding author Patsie Polly: Mechanisms of Disease and Translational Research Group, Department of Pathology, School of Medical Sciences, University of New South Wales, Sydney, NSW 2052, Australia; Tel: +61 2 9385 2924; Fax: +61 2 9385 1389; E-mail: patsie.polly@unsw.edu.au

mitochondrial dynamics, mitochondrial dysfunction, mitophagy, muscle wasting.

1. INTRODUCTION

Cancer cachexia is a multifaceted paraneoplastic syndrome occurring in approximately 50% of patients with cancer [1, 2]. It is a significant contributor to cancer morbidity and mortality, with approximately 20% of cancer deaths due to this syndrome [1, 2]. The disease is defined as progressive skeletal muscle wasting with anorexia, increased catabolic drive and functional impairment that is not effectively reversed with nutritional supplementation [3]. Heart failure and cardiac atrophy have also been reported in cancer cachexia [4]. Recent investigations suggest that cardiac atrophy may contribute to the high mortality rates in patients with cancer induced cachexia [5, 6]. The pathogenic mechanisms of cancer cachexia-induced cardiac failure are not well established.

Current research on cancer cachexia in cardiac tissue has relied on the development of animal and cell culture models due to the reduced availability and accessibility of human clinical samples [4]. Given the significant metabolic derangement in patients with cancer-induced cachexia, mitochondrial dysfunction may play a role in the pathogenesis of cancer cachexia-induced heart failure [7 - 9]. This chapter will focus on how known pathological mechanisms and previously identified key molecules relate to mitochondrial dysfunction in the heart in the context of cancer cachexia.

2. INFLAMMATORY CYTOKINES AND MITOCHONDRIAL DYSFUNCTION IN CARDIAC TISSUE

2.1. Tumour Necrosis Factor- α

Elevated inflammatory cytokines may facilitate heart failure in cancer cachexia by affecting cardiomyocyte mitochondrial function. Tumour Necrosis Factor (TNF)- α induced mitochondrial dysfunction in cardiomyocytes involves the generation of increased reactive oxygen species (ROS) [10, 11]. ROS may either induce, or result from, mitochondrial dysfunction. Therefore, investigating the source of ROS in cardiomyocytes may determine if mitochondrial dysfunction is a primary or secondary process in the pathogenesis of cancer cachexia in the heart. The main mitochondrial source of ROS in the heart is the electron transport chain (ETC), whilst non-mitochondrial sources in the heart include NADPH oxidases (Nox) and uncoupled nitric oxide synthases (NOS) [12]. TNF- α administered to a rat cardiomyocyte cell culture model demonstrated that the ETC was the major source of ROS [10]. Furthermore, in a ventricular pacing-induced canine model of

Congestive Heart Failure (CHF), TNF- α inhibition partially and completely restored cardiomyocyte mitochondrial complex III and ATP synthase activities respectively and ameliorated oxidative stress (Table 1) [13]. Thus, demonstrating that ETC is a source of ROS. In another ventricular pacing-induced canine model of CHF, blocking the function of complex I in the ETC in cardiomyocytes increased ROS production 2.8-fold. Interestingly, complex I enzymatic activity was decreased in the context of heart failure, possibly contributing to uncoupling and ROS production in mitochondria [14]. Further evidence of an ETC source has also been observed in a fibrosarcoma cell culture model, where TNF- α primarily produced ROS at the ubiquinone site [15]. These studies suggest a potential role for TNF- α - induced alterations in mitochondrial function in cardiomyocytes. Future therapeutic strategies, such as sodium selenite treatment, could target mitochondrial dysfunction to ameliorate cancer cachexia-induced cardiac atrophy [16].

Table 1. Summary of the Effects of Inflammatory Cytokines on Mitochondria in the Heart.

Cytokine	Model	Effect of the Cytokine Studied	Mechanism or Signalling Pathway	Reference
TNF- α	Canine model of pacing-induced CHF	Mitochondrial production of ROS	Complex III and ATP synthase dysfunction	Moe <i>et al.</i> , 2004 [13]
	Adult human cardiomyocyte cell culture	Activation of NF- κ B	NADPH oxidase production of ROS	Moe <i>et al.</i> , 2014 [17]
	Adult male Sprague-Dawley rats	Decreased ANT protein levels Altered membrane permeability transition pore opening in mitochondria	Increased ROS Unknown - hypothesised to be due to down regulation of ANT1	Mariappan <i>et al.</i> , 2007 [19]
	Neonatal Wistar rat ventricular cell culture	ROS mediated mitochondrial DNA damage	Sphingomyelin-ceramide pathway	Suematsu <i>et al.</i> , 2003 [10]
IL-6	Simulated ischemia/reperfusion in neonatal Sprague-Dawley rat ventricular cell culture	Increase inner mitochondrial membrane polarisation and increase mitochondrial Ca ²⁺ loading	PI3-kinase/Akt pathway	Smart <i>et al.</i> , 2006 [32]

In addition to the ETC, there are other non-mitochondrial sources of ROS in the heart including Nox and NOS [12]. There is emerging evidence for TNF- α -induced ROS generation from NADPH oxidase in cardiomyocytes [17] and TNF-

CHAPTER 4**Chalcone and Their Derivatives as Anticancer Agents****Zahoor A. Wani and Suaib Luqman****Molecular Bioprospection Department, CSIR-Central institute of Medicinal and Aromatic Plants, Lucknow-226015, Uttar Pradesh, India*

Abstract: Cancer has eventually stepped into the molecular insights focussing on the development of new generation of anticancer drugs especially of natural origin and its analogues with less or no toxicity issues and targeting specific molecular signalling pathways. In various therapeutic areas, numerous natural products and their derivatives have been effectively used to treat many human diseases or disorders. Chalcones, as metabolic precursors of some flavonoids and isoflavonoids have a structure of open chain flavonoids (1,3-diaryl-2-propen-1-ones) present in fruits and vegetables, possessing a broad range of biological activities including cancer chemotherapeutic and chemopreventive property. The anticancer properties of chalcones have been improved by substituting aryl rings (e.g. methoxy substitution on both aryl rings A and B) and introducing heterocyclic moieties. Hybridization with other pharmacologically important moieties (benzodiazepines, benzothiazoles, imidazolones etc.) by taking the help of SAR (structure-activity relationship) studies with much ease in preparation and oral administration ultimately has made chalcone a safe therapeutic agent. Some clinical trials revealed that these compounds did not cause toxicity and are present in plasma at optimum concentrations. Nowadays several chalcones are also used in cosmetic formulations and in food additives which could further be utilized for its chemopreventive potential. This book chapter briefly summarizes the demanding efforts made in the development of novel anticancer chalcones recorded in recent literatures with focussed cancer targets as well as presents an outline of the patents published in recent decades.

Keywords: Angiogenesis, antiproliferative, apoptosis, cancer, cell cycle arrest, cell line, chalcone, chalcone derivatives, chemoprevention, cytotoxicity, heterocyclic chalcone derivatives, IC₅₀, metastasis, NFκB, p53, p21, p23, TRIAL, tumor.

* **Corresponding author Suaib Luqman:** CSIR-Central Institute of Medicinal and Aromatic Plants, Lucknow-226015, Uttar Pradesh, India; Tel: +91-522-2718635; Fax: +91-522-2716141; E-mail: s.luqman@cimap.res.in

1. INTRODUCTION

Chalcones are phytomolecules belonging to the largest group of secondary metabolites of plant system with chemical structure of 1, 3-diphenyl-2-propene-1-ones, in which the two aromatic rings are linked by three carbon α , β -unsaturated carbonyl system (Fig. 1). The name was derived from “*chalcos*” meaning bronze due to its variant colour and it was given by Kostanecki and Tambor [1]. These compounds have a conjugated system where p -electron systems are delocalized with conjugated double bonds on both the benzene rings [2]. Chalcones due to their Michael acceptor features and small structure, easily bind with different cellular metabolites resulting in profound molecular and cellular effect ultimately exhibiting a broad range of biological activities [3]. Chalcones in general have lower redox potential due to enone (alkene-ketone) system and therefore, it prefers more electron transfer reactions. They are assumed to be intermediate metabolites in the synthesis of flavonoids and isoflavonoids that serve mainly for the defense system in the plants and thus protecting them from ROS (Reactive Oxygen Species) and consequently minimizing molecular and environmental injury. Functionally, it serves to regulate cholesterol levels, maintaining blood glucose levels, decrease the blood pressure, remove joint and muscle pain, help in sleep, improves immune system, liver and kidney functions, and enhance vision, skin beauty, hair growth and memory [4].

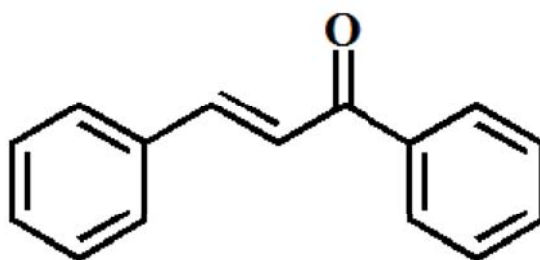


Fig. (1). Basic chalcone structure.

2. CONJUGATES AND DERIVATIVES OF CHALCONES

Chalcone has been widely used in organic synthesis to establish highly enantio-selective Michael adducts. The asymmetric catalytic conjugate adding a stabilized carbanion nucleophile to α , β -unsaturated carbonyl compounds shows one of the most essential carbon-carbon bond forming reactions in organic chemistry because the adducts are interesting intermediates for further optimization, such as amino-carbonyls, pyrrolidines and amino-alkanes. Many chalcone derivatives have also been prepared due to their convenient synthesis [3].

Madhavi *et al.* (2017) studied the synthesis of chalcones incorporating quinazoline derivatives as anticancer agents [5].

Bhale *et al.*, (2017) worked on the synthesis of protracted conjugated indolyl chalcones as the strong antioxidant, anti-inflammatory and anti-breast cancer agents [6]. Cai *et al.* (2017) studied the analogues of xanthon-chalcones and bis-chalcones as α -glucosidase inhibitors and anti-diabetes candidates [7]. Hawash and collaborators worked on the synthesis and bioactivity of novel pyrazolic chalcone derivatives as novel hepatocellular carcinoma therapeutics [8]. Ramaiah *et al.*, (2011) studied Chalcone-imidazolone conjugates and they found that these synthesized conjugates trigger DNA damage in the cells and show apoptosis [9]. Kamal *et al.*, (2015) studied about phenstatin/isocombretastatin-chalcone conjugates as effective tubulin polymerization inhibitors and mitochondrial apoptotic inducers [10]. Similarly, Khan (2009) published a patent on certain novel chalcones derivatives (in particular boronic chalcone derivatives) wherein he claimed that they possess anti-proliferative activity against cancer cells at micro molar concentrations. The invention provides the design and synthesis of novel boronic chalcone derivatives, and pharmaceutical compositions of chalcones derivatives. The invention also reveals the high activity and less toxicity of numerous compounds against breast cancer cell lines compared to normal MCF12A cells [11].

In glioblastoma cell lines, Indole chalcone has been recognized as a possible anticancer agent as it decreases the multiplication of cells under *in vitro* conditions. It was revealed that the indole chalcone battle for the binding site with colchicine and induces the inhibition of tubulin polymerization. Moreover, it distorted microtubule formation and triggered G2/M phase arrest and apoptosis. The molecule also worked as the dual inhibitor of Pgp and BCRP in glioblastoma cell line [12]. Chalcones are reported for both cancer chemotherapeutic action as well as for its chemopreventive mechanism as it possesses ability to inhibit carcinogenesis due to various modes like by increasing reduced glutathione levels and maintaining the optimum redox level [13].

TRAIL (Tumor necrosis factor related apoptosis inducing ligand) indicates programmed cell death specific to cancers and without any toxic effect to normal cells. TRAIL in association with the death receptor DR4 and/or DR5 mediates programmed cell death [14, 15]. However, decrease in the expression of pro-apoptotic proteins, TRAIL-R1 and TRAIL-R2 (death receptors) with concurrent upsurge in the level of anti-apoptotic proteins in tumor cells mediates TRAIL-resistance [16]. Szliszka *et al.* (2010) reported that the TRAIL-induced programmed cell death and cytotoxicity enhanced in prostate cancer cells by chalcones and dihydrochalcones such as phloretin. Their results show the

Regulation/Inhibition of Human Lactate Dehydrogenase A: An Innovative and Potential Approach for Anti-Cancer Drugs Development

Vinit Kumar¹, Atul Kumar² and Reshma Rani^{3,*}

¹ Amity Institute of Molecular Medicine and Stem Cell Research, Amity University, Noida 201313, India

² Amity Institute of Engineering & Technology, Amity University, Greater Noida 201313, India

³ Amity Institute of Biotechnology, Amity University, Noida 201313, India

Abstract: Human lactate dehydrogenase (*h*LDH-A), a glycolytic enzyme responsible for the conversion of pyruvate to lactate coupled with oxidation of NADH to NAD⁺, plays a crucial role in the promotion of glycolysis in invasive tumor cells. *h*LDH-A has been considered a vital therapeutic target for invasive cancers therefore, *h*LDH-A inhibition reflects a valuable attempt in the development of innovative anticancer strategies. Reagents that regulate or inhibit *h*LDH-A enzyme/ gene can play a role in the prevention and treatment of various cancers and related diseases. In fact, selective inhibition of *h*LDH-A using small molecules holds potential prospects for the treatment of cancer. Consequently, significant progress has been made in the discovery of small-molecules, the selective inhibitors of *h*LDH-A displaying remarkable inhibitory potency. The LDH-based approaches in the development of anticancer therapy and treatment of related diseases are worthwhile because of the existence of LDH enzyme at the end of glycolytic pathway. In this book chapter, 59 review and research articles, and 15 patents filed on LDH and its application are discussed. Latest contributions in regulation/inhibition of the LDH-A enzyme by various agents are summarized in this book chapter.

Keywords: Aerobic glycolysis, anaerobic glycolysis, anti-inflammatory activity, anti-proliferative activity, cancer cell metabolism, cancer cell proliferation, epileptic treatment, FDG-PET, FRET, glycolytic pathway, gossypol, human lactate dehydrogenase A, human lactate dehydrogenase B, isostere of pyruvate, metabolic switch, mitochondrial dysfunction, NADH/NAD⁺, nanosensor, *N*-hydroxy-indole, pyruvate dehydrogenase complex, selective *h*LDH5 inhibitors, tumor glycolysis, warburg effect.

* Corresponding author Reshma Rani: Amity Institute of Biotechnology, Amity University, Noida 201313, India; Tel: +91 1204392721; E-mails: reshudcy@gmail.com, rrani@amity.edu

1. INTRODUCTION

Cancer is one major cause among the leading causes of morbidity and mortality worldwide. According to the World Health Organization (WHO) approximately 14 million new cases of cancers and 8.2 million cancer related deaths were reported in 2012 worldwide and among them, most common causes of cancer death are cancers of lung (1.59 million deaths), liver (745 000 deaths), stomach (723 000 deaths), colorectal (694 000 deaths), breast (521 000 deaths) and oesophageal cancer (400 000 deaths) [1]. This data reflect the serious threats of cancer posed to human health, and the development of promising anticancer agents is, therefore, urgently required. WHO launched the global action plan in 2013, for the prevention, control and treatment of non-communicable diseases during 2013-2020 by aiming to reduce premature mortality from cancers. Common conventional approaches that are clinically used for the treatment of cancers, including surgery, chemo- and radiation therapy have some limitations due to serious side effects [2]. Various research studies confirm that cancer is a complex process and many enzymes including glycolytic enzymes are involved in initiation, maintenance and survival of various human cancers and some of them are thus considered as innovative targets for the development of anticancer agents [3]. Significant inhibition as well as reduction of enzyme activity involved in cancer by means of small molecules or other agents is a current and significant approach for drug development. Great efforts have been dedicated to design and develop ‘drug-like’ small molecules by following target-, structure- and fragment-based approaches for selective enzyme inhibition for the treatment of cancers [4].

2. CANCER CELL METABOLISM

The metabolic properties of cancer cells differ significantly from those of the normal cells. Unlike normal cells, most cancer cells rely on the enhanced rate of glycolysis that tends to ferment glucose into lactate, even under aerobic conditions. In fact, cancer cells are abnormally dependent on aerobic glycolysis for energy production at higher rate for maintenance of cancers [5]. It is hard to discuss the cancer cell metabolism without first mentioning the German scientist “Otto Warburg” who made a striking discovery in the 1920’s. For the first time, he observed that the cancer cells hold the metabolic switch from oxidative phosphorylation (OXPHOS) towards aerobic glycolysis (Warburg effect) and thus, established a link between cancer and the peculiar glucose metabolism in cancer cells [6]. Since then “Warburg effect” has been validated in various human tumors and the parallel increase in enhanced glucose uptake has been exploited clinically for the diagnosis, staging, and monitoring of various cancers, including nonHodgkins lymphoma (NHL) by using fluorodeoxyglucose positron emission tomography (FDG-PET). It is a useful and sensitive modality for assessing

disease activity in thyroid lymphoma and in cancer metastasis. In this technique, the biologically active molecule, an analogue of glucose i.e. fluorodeoxyglucose (FDG), is used for PET where the concentration of FDG tracer indicates metabolic activity of the tissue that corresponds to the regional glucose uptake [7]. Besides tumor imaging, Warburg effect can be exploited for drug designing to treat human cancers. Initially, Warburg hypothesized that the metabolic alteration specific to cancer cells is caused by a mitochondrial defect where complete oxidation of glucose is lost; however, it was later proven that this metabolic alteration is from oncogene-directed metabolic reprogramming, not from mitochondrial dysfunction [8 - 10]. The unique characteristic of tumor glycolysis of being highly functional is accompanied by high glucose consumption due to lower efficiency in energy production that ensures an adequate and rapid energy supply and biosynthetic intermediates for rapidly growing cancer cells [10 - 12]. In essence, cancer cells are hungrier for nutrients than normal cells are; thus, tumor glycolysis provides selective advantages to tumor cells for survival and proliferation. Succinctly, cancer is a metabolic disease and can be targeted by the following two facts (i) to produce enough energy to survive when supplies and waste disposal are limited, and (ii) to distract abundant metabolic intermediates to the biosynthetic pathways supporting cell proliferation. Recently, keen research interest in tumor glycolysis has emerged due to the strong metabolic dependencies of cancer cells. Despite various key factors (enzymes and transporters) that are intricate, tumor glycolysis is thus considered a promising target [10, 12].

3. GLYCOLYSIS AND LDHA

Glycolysis is a metabolic process, which comprises ten successive steps catalyzed by specific enzymes in the cytoplasm of the cells. At the end of glycolysis, two pyruvate molecules are formed by the catabolism of one glucose molecule with concurrent generation of two ATP molecules and two NADH (Nicotinamide Adenine Reduced Dinucleotide) molecules. Glycolysis can take place in both conditions; i) in the presence of oxygen i.e. aerobic condition and, ii) in the absence of oxygen i.e. anaerobic condition [13]. Although, in both the situations, the final products viz. two molecules of pyruvate, two molecules of ATP and two molecules of NADH are the same but depending upon the presence or absence of oxygen, further pyruvate can follow two different pathways. In aerobic (normoxia) condition, two molecules of pyruvate are transported into mitochondrial matrix where these molecules are decarboxylated and then enter into Krebs cycle (TCA; tricarboxylic acid) cycle to produce 36 molecules of ATP, carbon dioxide and water by oxidative phosphorylation (Fig. 1A). In normal cells under normoxia, glycolysis and oxidative phosphorylation are tightly coupled processes. In contrast, in an anaerobic (hypoxia) condition pyruvate is red-

CHAPTER 6

Cancer Chemo-Immunotherapeutics

Muzammal Hussain^{a,b,*} and Jiancun Zhang^{a,*}

^a State Key Laboratory of Respiratory Disease, Guangzhou Institutes of Biomedicine and Health, Chinese Academy of Sciences, 190 Kaiyuan Avenue, Science Park, Guangzhou, 510530, P.R. China

^b University of Chinese Academy of Sciences, Beijing, 100049, P.R. China

Abstract: Cancer chemo-immunotherapeutics has evolved with a strategic slogan - ‘marrying chemotherapy with immunotherapy’ - in order to optimize the chance for cure. The ultimate goal is to execute a ‘two hit’ impact, able on the one hand to mount a robust anti-tumour immune response, and on the other hand, selectively eradicate tumour growth and progression. Tremendous progress has been, and being, made in this regard by testing various ‘chemotherapy-immunotherapy’ drug combinations in the clinic, and also implementing multiple pharmacological and biological interventions against fundamental regulatory pathways involved in tumour development, progression, and tumour immune escape mechanisms. This chapter discusses the current ‘chemotherapy-immunotherapy’ combinations in clinical studies/trials, as well as the pharmacological manipulation of host-tumour cell interactions mapping the road ahead to a novel trend/concept of ‘two hit’ chemo-immunotherapeutics. At the end, we also discuss the patents issued and recent patent applications stating the novel chemo-immunotherapy methods with diverse interventional combinations, some of which produce synergistic anti-tumour effects, to treat multiple advanced cancers.

Keywords: ARG, Cancer, Chemotherapy, Chemo-immunotherapy, Cancer vaccines, CIK cells, CTLA4, Cytokine, FOLFOX, GVAX, IDO, Immune suppression, Immuno-oncology, Kinase inhibition, mAbs, mTOR, MAPK, PKA, R-CHOP, TroVax[®].

1. INTRODUCTION

Cancer chemo-immunotherapeutics implements treatment regimens that both maximize tumour regression and the anti-tumour immune response for the long

* **Corresponding author Muzammal Hussain:** Guangzhou Institutes of Biomedicine and Health, Chinese Academy of Sciences, 190 Kaiyuan Avenue, Science Park, Guangzhou, 510530, P.R. China; University of Chinese Academy of Sciences, Beijing, 100049, P.R. China; Tel: +86 020 32015323; E-mail: muzammal@gibh.ac.cn

* **Corresponding author Jiancun Zhang:** Guangzhou Institutes of Biomedicine and Health, Chinese Academy of Sciences, 190 Kaiyuan Avenue, Science Park, Guangzhou, 510530, P.R. China; Tel: +86 020 32015323; E-mail: zhang_jiancun@gibh.ac.cn

term clinical benefit of cancer patients [1]. Although cancer chemotherapy has historically been considered as immunosuppressive, emerging evidence indicates that certain chemotherapies can augment tumour immunity through multiple mechanisms including induction of immunogenic cell death and by disrupting strategies that tumours use to evade immune recognition [2 - 4]. This dual role, cytotoxicity and immune activation, from chemotherapy has prompted the scientific community to call for a radical but strategic shift in the way tumours are treated in order to achieve better clinical outcomes [1]. To this regard, cancer chemo-immunotherapeutics has evolved with a strategic slogan - 'marrying chemotherapy with immunotherapy' - in order to optimize the chance for cure [5, 6]. The ultimate goal is to execute a 'two hit' impact, able on the one hand to mount a robust anti-tumour immune response, and on the other hand, selectively eradicate tumour growth and progression. Tremendous progress has been, and being, made in this regard by implementing multiple chemotherapy-immunotherapy drug combinations with complimentary mechanisms of action to attain additive or synergistic anti-tumour effects [7, 8]. Apart from this, the complex interactions between host and tumour cells within the tumour microenvironment may lead to 'tumour-associated immune suppression', which is characterized by diverse mechanisms of oncogenic signaling pathways that play critical roles in tumour initiation, progression, and immunoescape [9]. Emerging data indicate that small-molecule based therapeutic interventions against such signaling pathways may have the potential to provide a 'two-hit' chemo-immunotherapeutic opportunity: direct killing of tumour cells, and the rescuing of endogenous anti-tumour immunity [10 - 12]. The purpose of this chapter is to discuss the clinical significance of current chemo- and immuno-therapy drug combinations, as well as highlight the 'two hit' chemo-immunotherapeutic potential of targeting the oncogenic signaling pathways that play crucial roles in tumour growth and progression, and in tumour-associated immune suppression. Moreover, we also discuss patents issued and recent patent applications demonstrating novel chemo-immunotherapeutic methods/formulations for the treatment of cancer.

2. CURRENT CHEMO-IMMUNOTHERAPEUTIC DRUG COMBINATIONS IN CLINICAL STUDIES

In recent decades, a general trend of harnessing endogenous anti-tumour immunity by modifying the diverse mechanisms of immunosuppressive tumour immune microenvironment has provided the knowledge and techniques to develop novel immunotherapeutic approaches for the treatment of cancer [7]. These include:

- a. Antibodies, such as monoclonal antibody drugs (mAbs) and the recent immune checkpoint-modulating antibodies;
- b. Vaccines, such as tumour cell-based autologous vaccines and dendritic cell (DC)-based vaccines; and
- c. Immunostimulatory cytokines, and cytokine-induced killer (CIK) cells.

Several combinations of these immunotherapeutics with traditional chemotherapy are in various phases of clinical trials (Table 1). In addition, superior clinical benefits of chemotherapy-immunotherapy combinations have also been convincingly demonstrated in the settings of large randomized trials.

Table 1. Summary of Currently Ongoing Clinical Trials Involving Cancer Chemo-Immunotherapy.

Chemo-Immunotherapy	Target of the Listed Antibody	Type of Cancer	Phase	Reference (NCT ID)
Chemotherapy-mAbs Combinations				
Doxorubicin, vincristine, cytarabine, etoposide, cyclophosphamide, methotrexate, leucovorin, filgrastim, epratuzumab	CD22	Acute lymphoblastic leukemia	Phase I/II	NCT00098839
Bendamustine, lenalidomide, rituximab	CD20	Chronic lymphocytic leukemia; Small lymphocytic lymphoma	Phase II	NCT01754857
Cyclophosphamide, fludarabine, ofatumumab	CD20	B-cell lymphoid leukemia	Phase II	NCT01762202
Bendamustine + ofatumumab	CD20	Mantle cell lymphoma	Phase II	NCT01437709
Carboplatin, paclitaxel, oregovomab	CA125	Ovarian neoplasms	Phase II	NCT01616303
Cisplatin, docetaxel, cetuximab	EGF receptor	Lung cancer	Phase II	NCT01059188
Fludarabine, cyclophosphamide, ofatumumab	CD20	Chronic lymphocytic leukemia; Small lymphocytic lymphoma	Phase II	NCT01145209
Paclitaxel, lapatinib, trastuzumab	HER2/neu	Breast carcinoma	Phase II	NCT01891357
Cyclophosphamide, fludarabine, rituximab	CD20	Multiple leukemias and lymphomas	Phase III	NCT02048813
CHOP chemotherapy (cyclophosphamide, hydroxydaunorubicin, vincristine, prednisone) plus G-CSF, combined with alemtuzumab	CD52	T-cell lymphoma	Phase III	NCT00646854
Fludarabine, rituximab	CD20	Chronic lymphocytic leukemia	Phase III	NCT00513747

Recent Advances and Challenges in microRNA-Based Cancer Therapeutics

Amjad Ali*, Faryal M. Awan, Aqsa Ikram and Shifa T. Ashraf

Atta-ur-Rahman School of Applied Biosciences (ASAB), National University of Sciences and Technology (NUST), H-12 Islamabad, Pakistan

Abstract: Despite significant advancements in understanding the cancer-associated signaling cascades, effective treatment strategies remain scarce. This intricacy of cancer enigma highlights a pressing need to develop novel therapeutics. The seminal discovery of microRNAs (miRNAs), a class of natural RNA-interfering agents, provides a new hope for accomplishing this task. Bolstered by a novel mode of action, the ability to function as tiny master regulators of cellular processes, ease of administration and sufficient uptake along with apparent lack of toxicity in normal tissues give miRNAs an extra edge and make them an ideal candidate for emerging therapeutic developments. Genome-wide investigations have shown more than half of the human miRNA genes being located on genomic regions or at fragile sites associated with cancer, unveiling the substantial significance of these small RNAs. Very soon after the discovery of the first miRNA, miRNA-based therapeutics has entered clinical trials and has shown fascinating results in preclinical development. This rapid progress through the discovery pipeline into clinical development imitates the significance of miRNAs as critical regulators in human diseases, and holds the pledge of yielding a novel class of therapeutics that could signify an attractive addition to the existing drug pipeline of Big Pharma. In this chapter, we will give an overview of the recent miRNA-based therapeutic approaches (patents: EP3110951, WO2017005771, EP3126496, EP3106168, WO2017005773, US9399773, EP2217248 and US9469854), and will discuss current translational challenges and further potential developments. These patents describe the potential of different miRNAs inhibitors/mimics for treating various types of cancers, these miRNAs include miR-34 mimic to treat hematologic malignancy/solid tumors; miR-409-5p, miR-379 and miR-154 inhibitors to treat prostate cancer and drug resistant lung cancer; miR-548z, miR-624-5p, let-7i-3p, miR-885-5p, miR-449b-3p to treat hepatoblastoma cancer; pre-miR-302 (an miRNA precursor) for cancer reversion; miR-21, miR-125a-5p, miR-191, miR-210, miR-222, miR-378, miR-423-3p, miR-638 inhibitors to treat hepatocellular carcinoma (HCC); hsa-miR-4510, hsa-miR-548aa, hsa-miR-548v and hsa-miR-37b-3p mimics to treat HCC; sorafenib- miR-34-mimic/ miR-215 mimic combination therapy for treating liver cancer and miR-21-3p mimic for treating liver diseases. The outcome of these patents may hopefully provide exciting opportunities and deeper

* **Corresponding author Amjad Ali:** Atta-ur-Rahman School of Applied Biosciences (ASAB), National University of Sciences and Technology (NUST), H-12 Islamabad, Pakistan;
Tel: +92-51-90856138/+92-333-9191903; E-mails: amjaduni@gmail.com, amjad.ali@asab.nust.edu.pk

insights into novel anti-cancer paradigms. Compared to conventional drug therapies, miRNA-based therapeutics appears to hold great promise to combat cancer, at least for those cancers where other treatment options have plateaued. Further developments in miRNA-based therapeutics are anticipated to translate miRNA-based therapeutic strategies into a clinical reality and may create a paradigm shift in medicine and pharmaceutical industry.

Keywords: Anti-miR oligonucleotides, cancer, cancer therapy, cancerous cells, clinical trials, drug resistance, drug target, *in vivo* studies, miRNA inhibitor, miRNA masking, miRNA mimic, miRNAs sponges, miRNA therapeutics, mouse models, mRNAs, Non-coding RNAs, oncogenic miRNAs, pre-clinical trials, siRNAs, tumor suppressor miRNAs.

1. INTRODUCTION

The genesis of the current non-coding RNAs' (ncRNAs) scientific paradigm can easily be traced to the influential discovery of a microRNA (miRNA) gene in 1993 by a group led by Ambros [1]. Extensive research on these evolutionarily conserved, small, regulatory RNAs in the last decade has shown fascinating breakthroughs of recent times [2]. Concomitantly, experimental confirmation of the results attained by the human genome project further revealed that several transcripts are actually non-coding transcripts, and that miRNAs signify the most important class of ncRNA molecules. The involvement of miRNAs in the development of cancer was initially reported in 2002: since then, the role of miRNAs has been intensively investigated in manifold human disorders [3, 4]. The noteworthy increment in the number of patent application filings over the last 10 years reflects a considerable amount of novelty in this area.

miRNAs are currently considered as master regulators of the human genome [5]. Clinical and functional implications of miRNAs in various disorders have hauled up these tiny cellular components to the ranks of ideal drug targets [6]. In eukaryotes, they serve as significant modulators of gene expression. They influence the transcriptome and proteome by targeting protein coding transcripts, hence aiding in cell fate determination. Furthermore, miRNAs have loomed as vital molecules in cancer research and they have proved to hold potential in cancer. They have the ability to repress stability of protein-coding transcripts and cellular translation by targeting the 3' untranslated regions (UTRs) in a sequential manner [7, 8]. This nature of selective silencing of gene expression by miRNAs is known to have a significant effect on human health and disease [9]. The therapeutic functioning of miRNA is based upon the catalytic process of the naturally occurring 15-22 nucleotide single stranded RNA, that enters the cytoplasmic multiprotein complex RNA-Induced Silencing Complex (RISC) to pair with mRNAs carrying complementary sequences and, as a result, repress

gene expression. Over 500 miRNAs of distinct nature have been identified in humans and more than 1000 have been predicted in total [10, 11]. Furthermore, exploration of the human genome sequences facilitated the discovery of the fact that miRNA genes frequently reside in fragile sites and genomic regions which are hot spots for chromosomal abnormalities [12]. Chromosomal abnormalities have been shown to be important for the etiology of various cancers. Various studies based on genome-wide approaches have reported strong association between various cytogenetic and molecular abnormalities and the location of miRNA genes [13 - 15]. Wey *et al.* investigated the genome-wide miRNA expression profiling in Intraductal papillary mucinous neoplasms (IPMNs) tissue and discovered six miRNAs (miR-100, miR-99b, miR-99a, miR-342-3p, miR-126, miR-130a) that may differentiate 'high-risk' IPMNs from 'low-risk' IPMNs [14]. Faltejskova *et al.* studied genome-wide miRNA expression profiling in colorectal cancer patients in order to discover miRNA signatures (miR-122, miR-122*, miR-885-5p, miR-10b, miR-143, and miR-28-5p) that would enable differentiation between primary tumors and their corresponding matched liver metastases [15, 16].

The ability of miRNAs to target multiple genes may hold the key to therapeutic accomplishment in cancer which is a heterogeneous disease and cannot be effectively treated by targeting a single gene of interest. Calin *et al.* first reported the role of miRNAs in cancer through the characterization of chromosome 13q14 done on chronic lymphocytic leukemia [3]. Successively, numerous studies gave strong evidences of deregulated miRNA expression in the hallmarks of cancer [17, 18]. In cancer, various cellular mechanisms are involved in the miRNA dysregulation such as genetic mutations [19], aberrant DNA methylation [20] and histone acetylation [21] along with alternative splicing, changes in the miRNA processing machinery and polyadenylation may cause hindrance in the maturation of miRNA [9, 22]. Abnormal gain or loss of miRNAs plays a role in initiation, progression, and metastasis and drug resistance in a variety of cancers. They can act as either oncogenes or tumor suppressors, depending upon the pathway or genes they impact. For instance, miRNAs of the let-7 family are a class of tumor suppressors. Let-7 expression has been reported to be downregulated in breast, head, neck, ovarian, lung and prostate cancers [23]. Additional miRNAs, namely miR-17/92 cluster, miR-221, miR-222, miR-21, miR-155 and miR-9 are upregulated in various cancers [24]. The therapeutic and diagnostic significance of miRNAs is remarkable since they have unique profiles and high stability in biological samples. For cancer therapy, miRNA expression modulation is under investigation but therapeutic tempering is attained by oncogenic inhibition of miRNAs, or by altering tumor suppressor miRNAs [9]. Furthermore, Phytonutrients that modulate expression and action of miRNA which are functionally involved in cancer pathobiology may have a potential to consider as a

Author Index

Amjad Ali	Patsie Polly
Aqsa Ikram	Paula Mosińska
Arkadiusz Michalak	Reshma Rani
Atul Kumar	Riddhi I. Bhavsar
Biswa P. Chatterji	Shifa T. Ashraf
Faryal M. Awan	Suaib Luqman
Hussain Muzammal	Thomas G Ikonmidis
Jakub Fichna	Timothy C. Tan
Jiancun Zhang	Vinit Kumar
Needa N. Bagban	Zahoor A. Wani

SUBJECT INDEX

A

- Abraxane 48
 Acids 1, 2, 3, 6, 7, 9, 17, 51, 131, 132
 bile 17
 docosaheptaenoic 1, 2, 3, 7, 9
 eicosapentaenoic 1, 3, 7, 9
 linoleic 2, 6, 9
 linolenic 51
 pyruvic 131, 132
 Activity 4, 6, 12, 41, 50, 52, 56, 65, 79, 82, 84,
 100-103, 118, 119, 133, 158, 186, 189,
 194
 anti cancer 12, 56
 catalytic 119
 Adenine nucleotide translocator (ANT) 78
 Adenosine 82, 86, 154
 S-Adenosyl methionine (SAM) 200
 Adipocytes 18
 Adriamycin 161
 Advanced testicular cancer 40
 Aerobic glycolysis 114, 115, 135
 Agents 31, 40, 53, 59, 61, 62, 64, 95, 103, 104,
 162
 anti-breast cancer 95
 antimicrotubule 59
 antimitotic 31, 62, 103
 antitubulin 61
 microtubule-stabilizing 162
 potent cancer cytotoxic 104
 potent microtubule destabilizing 40
 tubulin binding 53, 64
 Aggressive 11, 122
 lipogenic cancer phenotype 11
 phenotypes 122
 Angiogenesis aids 36
 Angioinhibitors 36, 67
 Anticancer 52, 53, 102, 104, 114, 147, 191
 adjuvant 52, 53
 antibiotics 53
 benzimidazoles 53
 mAbs 147
 medicines 102
 therapeutics 191
 therapy 104, 114
 Anticancer agents 53, 115, 126
 antibody 53
 promising 115
 Anti-miRNA 193, 194
 oligonucleotides 194
 therapy 193
 Anti-miR oligonucleotides (AMOs) 179, 182,
 193, 194, 195
 Anti-proliferative activity 95, 105, 114
 Antisense oligonucleotide 124, 125, 193, 194
 stranded 124, 125
 Anti-tumour 143, 144, 150-153, 155, 157, 160,
 164
 activity 150, 151
 robust 143, 144, 155, 160, 164
 Anti-tumour immunity 144, 149, 155
 endogenous 144, 149
 Apoptosis 11, 14-17, 19, 31-33, 36, 39, 48, 50,
 55-58, 79-81, 93, 95, 97, 104, 125, 158,
 190, 191, 192, 194, 201
 cancer cell 56
 cellular 125
 Aryl benzimidazoles 53, 54
 Aspirin 9, 14
 action 14
 placebo 9
 ATP synthase 85
 Autophagy 81, 84, 85
 in cancer cachexia 84
 induction 81, 84, 85

B

- Bendamustine 145
 Benzimidazole derivatives 31, 53
 Biosynthesis-related genes 201
 Breast cancer 16, 41, 43, 45, 48, 49, 53, 58,
 60, 126, 127, 146, 147, 152, 155, 157,
 195, 196, 197

human 196
 metastatic 41, 49
 negative 146
 resistant 49
 Burkitt's lymphoma cells 123

C

Cachexia 75, 76, 81
 cancer-induced 75, 76
 Cachexia-induced cardiac failure 76, 78, 79
 Caco-2 colon cancer cells 17
 Cancer 4, 5, 8, 9, 13, 16, 18, 21, 33, 53, 58, 76,
 77, 104, 107, 114-116, 122, 127, 131,
 144, 148, 151, 152, 158, 161, 178, 191,
 198
 adipocyte 18
 associated signaling cascades 178
 advanced metastatic 151
 ameliorate 77
 anal 21
 bladder 53, 191
 brain 53, 127
 chemotherapy 53, 144
 gallbladder 53
 gastric 33, 122, 152
 genotypes 33
 glycolytic 131
 hepatoblastoma 178, 198
 hepatocellular 8, 9
 invasion 131
 invasive 114
 immunotherapy 148, 158
 neck 122
 metabolism 18
 metastasis 116
 morbidity 76
 non-melanoma skin 16
 oesophageal 115
 preventing 4, 5, 13, 104
 rectal 53
 stomach 53, 58, 107
 treating drug-resistant 161
 uterine 53

Cancer cachexia 75, 76, 78, 79, 80, 81, 82, 83,
 84, 85, 86
 context of 75, 76
 induced cardiomyocyte autophagy 84
 skeletal muscle of 83
 treatment of 80, 83
 Cancer cell(s) 10, 13, 15-18, 20, 33-36, 52, 57,
 58, 67, 83, 95, 96, 98, 99, 100, 106, 115-
 118, 125, 128, 133, 135, 136, 147, 155,
 158, 159, 186-188, 199, 200, 201, 202
 aids 67
 arrest 57
 breast 35
 chemo-sensitivity 158
 conditioned medium (CCCM) 83
 liver 199, 200, 201, 202
 lung 15, 187
 ovarian 18
 pancreatic 15, 125, 128
 proliferation/progression 159
 prostate 95, 100
 remnant 35
 resistant 58
 trastuzumab-resistant breast 125
 Cancer chemo 154, 155, 159
 immunotherapeutics, two hit 154, 155
 immunotherapy 159
 Cancer chemotherapeutic 93, 95 98, 107
 action 95
 effective 107
 Cancerous cells 32, 33, 34, 66, 96, 104, 179,
 181, 190, 192
 gefitinib-resistant 181
 sensitive 96
 Cancer stem cells (CSCs) 32, 197
 Cancer-targeting vehicles 23
 Cancer therapeutics 189, 191, 196
 miRNA Based 196
 Cancer therapy 4, 11, 15, 18, 20, 31, 51, 66,
 152, 162, 179, 180, 186, 187, 193, 195,
 196, 198
 regimens 66
 Cancer treatment 65, 66, 134, 135, 160, 162,
 181
 colorectal 160
 improving colorectal 181

- Cancer types 16, 152, 184, 188
multiple 152
- Cancer vaccines 143, 149, 150
five-peptide 150
personalized therapeutic 149
- Carboplatin 43, 55, 59, 145, 146, 148
- Carcinogenesis 2, 17, 95, 105-107
- Cardiac atrophy 75-77, 80, 86
cachexia-induced 75, 77, 86
- Cardiomyocytes 75-78, 80-86
in cancer cachexia 80, 82-86
- Casein K2 peptide inhibitor 31
- Castration-resistant prostate cancer (CRPC)
150, 151
- Celecoxib 14, 15
- Cell lines 11, 19, 50, 54, 56, 57, 65, 93, 95, 96,
104, 123, 126, 187
glioblastoma 95
human cancer 65
leukemia 54, 126
- Cells 97, 201, 202
hepatoblastoma 201, 202
isobavachalcone-treated 97
- Cellular lactate consumption 133
- Cervical cancer 32, 126, 152, 157
- Cetuximab 36, 58, 59, 145
- Chalcone 98, 99, 101
analogues, heterocyclic 98, 99
flavone derivatives 101
- Chalcone derivatives 93-99, 102, 103, 105
boronic 95, 102
claimed 96
coumarin 98
hetero aromatic 98
heterocyclic 93
novel boronic 95
novel pyrazolic 95
substituted 102
vinyllogous 98, 103
- Chalcones 93, 98, 100, 102, 104
boronic 98, 104
isoxazolyl aryl 98, 102
novel anticancer 93
substituted 98, 100
- Chalcones derivatives 94, 95, 105, 107
synthetic 107
- Chemoreceptor trigger zone (CTZ) 160
- Chemotherapeutic agents 10, 40, 51, 57, 97,
159-163
- Chemotherapeutic drugs 34, 159
classical 11, 15, 19, 20, 39, 58, 61, 65, 161
- Chemotherapeutics 11, 15, 19, 20, 39, 161
- Chemotherapy 1, 3, 6, 10, 19-21, 23, 34, 35,
49, 51, 57, 58, 66, 67, 125, 143, 144-152,
158, 160, 161, 162
CIK combinations 146
conventional 67, 160, 161
cytokine combinations 151
first-line 149
marrying 143, 144
immune checkpoint inhibitor
combinations 146, 148
mAbs combinations 145, 147
regimens 148, 149
rituximab combinations 148
vaccine combinations 146, 149
- Cholesterol-conjugated miRNA duplex 183
- CHOP chemotherapy 145, 146
- Chromosomal abnormalities 180
- Chronic lymphocytic leukemia (CLL) 145,
147, 151, 180
- Cisplatin 41, 43, 49, 55, 59, 145-149, 151,
161, 202
- CK2 phosphorylation 55
- Colon cancer 8, 35, 162, 193
- Colorectal cancer 1-3, 22, 53, 56, 122, 151,
181, 185
metastatic 151
- Colorectal mucosa 2
- Combination 143-145, 153, 162
chemotherapy-immunotherapy 143, 145
chemotherapy-immunotherapy drug 143
chemotherapyoncolytic immunotherapy 153
diverse interventional 143, 162
implementing multiple
- Combination 49, 57, 150, 151, 162, 202, 203
chemotherapy protocols 57
of gemcitabine 150, 151
therapy 49, 162, 202, 203
invention unveils 202
- Combretastatin 31, 62, 63, 64
- Combretastatin A-1 62, 63

phosphate 62, 63
 Congestive heart failure (CHF) 77, 81, 83
 Cremaphor 48
 Cryptophycin 43, 44
 Curcuminoids 13
 Cyclic dinucleotide 162
 Cyclooxygenases 1, 3, 9
 Cyclophosphamide 55, 59, 145-147, 150, 151
 Cytokine-induced killer (CIK) 145
 Cytotoxic activity 65, 97, 103

D

DC-CIK-chemotherapy combinations 152
 Deregulated miRNA expression 180
 Dihydrochalcones 95, 96, 101
 Dihydronicotinamide 120, 121
 DNA linking agents 59
 Docetaxel 31, 46-48, 50, 51, 55, 60, 61, 96, 145, 146, 150, 151
 Dolastatins 43
 Domain, substrate binding 118
 Double-blind RCT 8, 9
 Double stranded nucleic acid 124, 125
 Doxorubicin 8, 11, 55, 59, 145-147, 159, 162, 202
 Drug development 115, 118
 Drug resistant lung cancer 197
 Durvalumab 146

E

Elongases 1, 3, 11
 Elpamotide 150
 Endogenous angioinhibitors 35, 36
 Enzyme 114, 115, 117, 123, 134
 lactate dehydrogenase 134
 glycolytic 114, 115, 117, 123
 EPA placebo 9
 Epigallocatechin 126
 Epileptic treatment 114, 129, 130
 Epothilones 31, 49, 50, 66
 Epstein-Barr Virus (EBV) 146
 Erlotinib 58, 59
 Esophageal cancer 146

Ethyl esters (EE) 19, 22
 Etoposide 55, 145, 148
 Expression 4, 10, 11, 13, 14, 17, 20, 55, 80, 81, 95-98, 100, 103, 104, 122-124, 160, 184, 188, 189, 191, 195, 196, 199, 201, 204
 elevated 17
 increased 17, 80, 81, 123
 LDH-A 122, 123
 Extracellular matrix 52

F

Familial adenomatous polyposis (FAP) 12, 14
 Fatty acids (FAs) 1, 2, 6-12, 16-18, 20-23, 51, 81, 82, 86
 binding proteins (FABPs) 5, 16-18
 elongase 9
 oxidation 81, 82
 free 9, 22, 23
 utilisation 81, 82
 Fibroblasts 122, 123, 153
 Fish oils 2, 3, 5, 9, 13, 19-23
 Flavonoids 93, 94
 Fludarabine 55, 145, 147, 151
 Fluorouracil 57, 58, 162
 Forster resonance energy transfer (FRET) 114, 133
 Free fatty acid (FFA) 9, 22, 23
 Front-line chemoimmunotherapy 147

G

Galloflavin 126
 GalNAc-conjugated antimiRs 183
 Gastrointestinal cancers 1, 8, 146
 neoplasms 146
 Gefitinib 36, 58, 59
 Gemcitabine 8, 43, 125, 128, 146, 147, 149-152, 161-163
 Genome-wide miRNA expression profiling 180
 Glucose 82, 85, 122, 123
 deprivation 122, 123
 utilisation 82, 85

Glycolysis 81, 85, 114-116, 122, 135
 anaerobic 85, 114, 117
 Glycolytic pathways 114, 118, 136
 Glyoxylate 123
 Growth, hormones impact cancer 35
 GTP-bound β -tubulin 38, 39

H

Hairpin 124, 125
 Halichondrin 31, 42, 43, 66
 Heart failure 75-79, 81-84, 86
 cachexia-induced 76, 78, 83, 84
 congestive 77, 81, 83
 HeLa cancer cell line and compromised cell proliferation 128
 Hematologic malignancies 98, 203
 Hepasphere/quadrassphere microspheres 8
 Hepatitis C virus (HCV) 194
 Hepatoblastoma 198, 201
 Hepatocellular carcinoma 59, 147, 178, 199-201
 cells 201
 Hepatocytes 200, 201
 primary human 201
 Heteroaromatic chalcone derivatives 102
 Histone deacetylases (HDACs) 100
 Homobutein 100
 Homoisoforn 119
 Human 10, 100
 bladder cancer cells 100
 cancer cells 10
 Human cancers 53, 115, 116, 122, 136, 157, 184, 198, 199
 diverse 53
 multiple 157
 Hybrid liposomes (HLs) 19
 Hydrolysable site 38
 Hydroxydaunorubicin 145, 146

I

IDO-mediated catabolism 158
 Indole derivatives inhibitors 134
 Induced STAT3 phosphorylation 82

Inhibitors 50, 51, 114, 162, 185
 miRNA-pathway 185
 mTOR 155, 162
 selective 50, 51, 114
 Inhibitory 125, 126, 129, 131, 136
 activity 125, 126, 129, 131, 136
 effect, synergistic 125
 Insulin-resistance 75, 85
 Interactions, stromal-immune-cancer cell 154
 Interferons 19, 35, 36, 59
 Interphase microtubule mass 39
 Intraductal papillary mucinous neoplasms (IPMNs) 180
 Ipilimumab 148, 149, 162
 combinations 148
 Isoenzyme 131, 133, 135
 Isoflavonoids 93, 94
 Isoliquiritigenin 100
 Isostere of pyruvate 114, 125
 Ixabepilone 49, 50, 61, 62

J

J-Series prostaglandin-ethanolamides 5

K

Ketone bodies 83

L

Lactate 114, 117-124, 127, 129, 131, 133, 134-136
 dehydrogenase (LDH) 114, 117-122, 124, 127, 129, 131, 133-136
 dehydrogenase inhibitor 134, 135
 production 122, 123, 135
 cellular 133
 transporters activity 133
 LDHA 132, 133
 activity 132
 stable inhibition in LDHB 133
 LDH 117, 119, 131, 132, 133-135
 activity 132, 133
 in cancer cells 117

inhibitors 117, 119, 131, 135
 in cancer diagnosis 133, 134
 ratio of H-type 132
 total 132, 133
 Lipid 16, 17, 201
 droplet contents 201
 trafficking 16, 17
 Lipxygenases 1, 3, 4
 Liver cancer 32, 53, 96, 101, 187, 196, 199-202
 advanced primary 202
 cell lines 96, 200
 glypican-3-targeting miRNAs 201
 human 199
 treatment of 196, 199
 LNA-modified antisense inhibitor 183
 Locked nucleic acids (LNAs) 193, 195, 197
 LOX inhibitors 15
 Lung cancer 53, 61, 65, 96, 98, 122, 145, 147, 162, 178, 187, 191, 196, 197
 cell lines 98
 resistant 178, 196
 small cell 61, 65, 122
 treat prostate cancer bone metastasis and drug resistant 196, 197
 Lycofalone 15
 Lymphomas 19, 40, 41, 46, 49, 53, 61, 127, 145, 147, 148
 Lysine 100, 123
 5 acetylation 123
 residue 100

M

Malignant pleural mesothelioma 146, 193
 MAPK 156, 157
 inhibition 157
 signaling 156, 157
 Membrane permeability transition pore (MPTP) 78, 83, 85
 Metabolic 116, 163
 alteration 116
 targeting cancer cells 163
 Metabolism 1, 3, 10, 23, 45, 58, 81, 114, 115, 135, 204

cancer cell 114, 115
 Metastasis 5, 13, 18, 32, 93, 99, 122, 123, 131, 156, 158, 180, 184, 186-188, 193, 194, 196
 omental 18
 prostate cancer bone 196
 Metastatic castrate-resistant prostate cancer 146
 Microtubule 31, 39, 40, 43-46, 48, 50, 56, 66
 associated proteins (MAPs) 39
 catastrophe 39
 depolymerising agent 45
 depolymerizers 31
 destabilization 43
 destabilizers 31, 39, 66
 destabilizing agents 39
 dynamicity 45, 46, 48
 dynamics 31, 39, 40, 44, 66
 formation 56
 polymerizers 31
 stabilizers 31, 39, 46, 66
 stabilizing properties 49, 50
 Mimic combination therapy 178
 Minimal residual disease (MRD) 147
 MiR, mature miRNAs of 201
 MiRagen therapeutics 182, 183
 miRNA 178, 179, 181, 189, 190, 193-195, 203, 204
 antagonists 189, 190, 193
 based drugs, developing 181
 based therapeutics 178, 179, 203, 204
 based therapies 189
 based treatment 204
 binding site 194, 195
 biogenesis 181
 developing 189, 203
 miRNA biogenesis 179, 181, 182, 185, 195, 204
 enhancing 204
 inhibiting 185
 inhibiting oligonucleotides 204
 inhibitor molecules 197
 inhibitors 179, 182, 195
 pathways 181
 miRNA mimic(s) 179, 182, 186, 187, 189-191, 202, 203

believed 191
 double-stranded 190
 sorafenib 202
 therapeutic 186
 therapy 190
 tumor suppressor 186
 miRNA processing machinery 180
 Mitochondrial pyruvate consumption 133
 Mitogen-activated protein kinase (MAPK) 10,
 143, 154, 156, 157, 158
 mRNA degradation 190
 M-type LDH activity 132
 Multiple advanced cancers 143, 152, 154, 157,
 162, 163
 Multiplex RNAi inhibition strategy 184, 185
 Muscle protein degradation 80

N

NADH competition assay 127, 128
 NADPH oxidase 75, 76, 78
 Nasopharyngeal cancer 146
 Natural killer (NK) 13, 152, 153
 Neoplasia 127
 Neoplastic diseases 62-64, 97
 Neuroblastoma 195
 Neutrophils 10
 Next-generation sequencing (NGS) 187
 NFκB signalling in human bladder cancer cells
 100
 N-hydroxy-indole 114
 Nivolumab 148, 149
 Non-Hodgkin lymphoma (NHL) 115, 151
 Non-small cell lung cancer (NSCLC) 21, 41,
 43, 49, 146, 148, 149, 152, 155, 183, 192
 Novel 5, 95, 97, 103, 178, 194, 203
 anti miRNA 194
 chalcone(s) derivatives 95, 97, 103
 miRNA-formulations 203
 therapeutics 5, 178
 Novel anti-cancer 18, 179
 agents 18
 paradigms 179

O

Ofatumumab 145
 Oligomer nucleotide units 197
 Oncogenic 144, 179, 182, 189, 194
 miRNAs 179, 182, 189, 194
 signaling pathways 144
 Oncolytic 147, 152, 153
 adenovirus 147
 immunotherapy 152, 153
 OncomiRNA 194
 O-phenyl chalcone compounds 102
 Ovarian cancer 18, 19, 53, 56, 58, 61, 146,
 156, 184, 193
 microenvironment 156
 Overall response rate (ORR) 149, 151
 Overexpression 122, 123
 of c-Myc 123
 of hLDH-A 122, 123
 Oxadiazole scaffold 53, 54
 Oxalate 123, 124
 Oxaliplatin 20, 57, 58, 59, 146, 151, 152
 Oxamate 119, 120, 125, 126
 Oxidative capacity 82, 83

P

Paclitaxel 48, 55, 57, 96
 docetaxel 48, 55, 96
 stabilizing microtubules 57
 Pancreatic cancer 53, 83, 122, 123, 125, 127,
 146, 147, 152, 157, 162, 191
 advanced 152
 cell growth 123
 cell lines 125
 initiation 123
 Pancreatic ductal adenocarcinoma cells
 (PDACs) 19
 Pembrolizumab 146, 148, 149
 Pentoxifylline 31, 52, 53
 Peroxisome 16, 81, 82
 proliferator-activated receptor (PPAR) 16,
 81, 82
 Phosphorylation, oxidative 82, 115, 116, 118
 Polypeptide, therapeutic 160

Polyunsaturated fatty acids 1, 2, 3, 5, 9
 Prednisone 145, 146, 147, 151
 PremiRNA 198
 Prevention of gastrointestinal cancers 8
 Progression-free survival (PFS) 148, 149, 151
 Proliferating cancer cell 117
 Proliferating cancerous lymphoma cells 34
 Promising direct anti-cancer, demonstrated 155
 Properties 14, 15, 20, 22, 40, 43, 45, 46, 60, 62, 104, 105, 106, 203
 anti-cancer 22
 anti-inflammatory 105, 106
 antimitotic 45, 46, 62
 Prostate cancer 15-18, 35, 43, 49, 53, 56, 96, 149, 150, 162, 178, 180, 187
 advanced 149, 150
 castration-resistant 150
 cell lines 17
 metastatic 35
 Proteasome 99, 104
 Protein kinase(s) 12, 14, 82
 B (PKB) 14
 C (PKC) 82
 Proteins 10, 12, 19, 54, 55, 67, 78, 80, 81, 104-106, 118, 119, 151, 161, 162, 182, 185
 fibronectin domain antibody-IL-2 fusion 161, 162
 miRNA biosynthesis protein TAR RNA-binding 185
 Pyrazolyl-thiazole 129, 130
 Pyruvate 114, 116-122, 125-127, 131, 135, 136
 competitive inhibitor of hLDH-A 125
 conversion of 114, 117, 120, 121
 dehydrogenase complex 114, 131
 molecules of 116
 reduction 121, 122, 135
 structural isostere of 114, 125

Q

Quinolone chalcone compounds 101
 Quinoxaline-chalcone derivatives 97

R

Radiation therapy 35, 66, 115
 Radiotherapy 6, 20, 21, 135
 Reactive oxygen species (ROS) 76-78, 85, 94, 125
 Regulation/inhibition of hLDH-A by small molecules 124
 Regulators, immune-suppressive 153, 154
 Regulus therapeutics 183
 Residues 118-120
 catalytic 120
 Resistance 10, 11, 20, 48, 51, 55, 57, 58, 61, 125, 135
 taxane 48
 Rituximab 35, 145-147, 151
 RNA-induced silencing complex (RISC) 179, 182, 193
 ROS regulation in cardiomyocytes in cancer cachexia 83

S

Sensitize cancer stem 20
 Serum alpha-fetoprotein 199
 Small lymphocytic lymphoma 145
 Small molecules inhibitors of miRNAs 185
 Sorafenib 181, 196, 202
 microRNA combination therapy for liver cancer 196
 miRNA combination therapy for liver cancer 202
 resistance 181
 Stabilizing microtubules 57
 Stimulate apoptosis, non-accustomed cells 11
 Structure activity relationship (SAR) 93, 100, 104-107, 127
 Substrate 120, 127, 128
 binding regions 128
 pyruvate 120, 127
 Superparamagnetic nanoparticles 5
 Synergistic 51, 143, 144, 161, 162
 anti-tumour effects 143, 144, 161, 162
 drug combinations 51

treatment 161, 162
 Synergy 20, 51, 150, 151
 therapeutic 51
 Synthesis of collagen 52, 131

T

Taccalonolides 31, 50, 66
 Tasidotin 43
 Taxane- binding site 46, 49
 Therapeutic 75, 77, 84 86, 160, 178, 179, 187, 188
 agents 160, 188
 approaches, miRNA-based 178
 interventions 75, 187
 strategies 75, 77, 84, 86
 miRNA-based 179
 Thymosin 160
 Tobacco smoking 2
 Topoisomerase inhibitors 59
 Transcription factor-miRNA interplay 189
 Trastuzumab 36, 49, 59, 125, 145, 147
 Tricyclic pyrimidine tyrosine kinase inhibitors 61, 62
 Tryptophan 157, 158
 Tubulin 38, 39, 53, 54, 57, 60, 61, 63, 64, 65, 95
 assembly 57, 64
 inhibitors of 53, 60
 polymerization 54, 57, 61, 63, 64, 65, 95
 subunits 38, 39
 Tumors 5, 6, 10, 18, 34, 36, 50, 51, 52, 55, 56, 58, 60, 67, 95, 96, 114, 116, 117, 122-124, 135, 161, 193
 cancer advances 161
 cells 6, 34, 36, 50-52, 55, 56, 58, 60, 67, 95,

96, 116, 122-124, 135
 chemoresistance 10, 55, 56
 glycolysis 114, 116, 117
 malignant 5
 primary ovarian 18
 regression 193
 Tumorigenesis 1, 122, 123, 157, 159
 Tumor suppressor(s) 144, 154, 164, 179, 180, 182, 184-186, 189, 190, 201, 202
 miRNAs 179, 180, 182, 185, 189, 201, 202
 Tumour-associated immune suppression 144, 154, 164
 Tumour cells 144, 152, 154, 155, 157, 158, 160, 162, 163
 direct killing of 144, 154
 high ARG-expressing 158

U

Ubiquitin-proteasome system (UPS) 80, 81, 85
 Upregulated IL-6 signalling 85

V

Vasoactive intestinal peptide (VIP) 160
 Vasoconstriction 10

W

World health organization (WHO) 115
 WT1-targeted DC vaccination 146

X

Xanthohumol 98



PROF. DR. ATTA-UR-RAHMAN, FRS

Atta-ur-Rahman, Ph.D. in organic chemistry from Cambridge University (1968), has 1080 international publications in several fields of organic chemistry including 751 research publications, 37 international patents, 69 chapters in books and 221 books published largely by major U.S. and European presses. He is the Editor-in-Chief of eight European Chemistry journals. He is Editor of the world's leading encyclopedic series of volumes on natural products "Studies in Natural Product Chemistry" 54 volumes of which have been published under his Editorship by Elsevier during the last two decades.

Prof. Rahman won the UNESCO Science Prize (1999) and was elected as Fellow of the prestigious Royal Society (London) in July 2006. He has been conferred honorary doctorate degrees by many universities including (Sc.D.) by the Cambridge University (UK) (1987). He was elected Honorary Life Fellow of Kings College, Cambridge University, UK, conferred the TWAS (Italy) Prize and the Austrian government has honoured him with its high civil award ("Grosse Goldene Ehrenzeischen am Bande") (2007). He is Foreign Fellow of Chinese and Korean Academy of Sciences, Foreign Fellow of the Chinese Chemical Society and former President of Pakistan Academy of Sciences.

See discussions, stats, and author profiles for this publication at: <https://www.researchgate.net/publication/340925757>

The Transition of Political Imperialism to Economic Imperialism: A Historical Study of the genesis of the native capitalist class in the Textile Industry of Bombay in the 19th cent...

Chapter · July 2017

CITATIONS

0

READS

23

1 author:



Avkash Jadhav

St. Xavier's College, Bombay

12 PUBLICATIONS 0 CITATIONS

SEE PROFILE



**Masters International
R&D Center**

MIRDEC 2017

MIRDEC – 4th
International Academic Conference
Social Science, Multidisciplinary and Globalization
Studies

CONFERENCE PROCEEDINGS
MADRID, SPAIN

Full Paper Series

Editors
Kemal Cebeci
Adam Pawlicz
Slagjana Stojanovska

Holiday-inn Piramides, Madrid, Spain
04-07 JULY 2017

Masters International Research & Development Center

MIRDEC
International Academic Conference

MIRDEC-4th,
International Academic Conference on
Social Science, Multidisciplinary and Globalization
Studies

CONFERENCE PROCEEDINGS

FULL PAPER SERIES

Editors:

Kemal Cebeci
Adam Pawlicz
Slagjana Stojanovska

ISBN: 978-605-82290-0-6

Holiday Inn Madrid – Pirámides, Madrid, Spain
04-07 July, 2017

MIRDEC-4th, International Academic Conference on
Social Science, Multidisciplinary and Globalization Studies,
04-07 July 2017, Madrid, Spain

KEYNOTE SPEAKERS

Professor Marta Mas Machuca

Universitat Internacional de Catalunya,
Barcelona, Spain

Dr. Avkash Jadhav

St. Xavier College, University of Mumbai,
India

Dr. Maria del Populo Pablo Romero

University of Seville, Spain

Keynote Session Chair:

Diana K. Kerr

University of St. Andrews, United Kingdom

**MIRDEC-4th, International Academic Conference on
Social Science, Multidisciplinary and Globalization Studies.
04-07 July 2017, Madrid, Spain**

Special thanks to Keynote Speakers:

Professor Marta Mas Machuca

Universitat Internacional de Catalunya,
Barcelona, Spain

Dr. Avkash Jadhav

St. Xavier College, University of Mumbai, India

Dr. Maria del Populo Pablo Romero

University of Seville, Spain

Special thanks to Session Chairs:

Diana K. Kerr

University of St. Andrews, United Kingdom

Paulo Mourao

University of Minho, Portugal

Rafat Fazeli

*University of Redlands & California State University, Fullerton,
USA*

Pere Ayling

University of Suffolk, United Kingdom

Alexander Christopher Rauchstaedt

UCAM Universidad Católica San Antonio de Murcia, Spain

Gloria Maria Canales Vargas

Universidad Europea Madrid, Spain

We are very pleased to introduce the proceedings(full paper series) of the Masters International Research and Development Center, MIRDEC 4th, International Academic Conference on Social Science, Multidisciplinary and Globalization Studies, 04-07 July 2017, Madrid, Spain. MIRDEC thanks to all our participants for their academic and social contributions.

Mirdec-4th Madrid 2017 Conference Proceedings, Full Paper Series

Masters International Danismanlik Arastirma Yayincilik
Masters International Consultancy Research and Publishing

ISBN: 978-605-82290-0-6

MIRDEC Publishing

Editors:

Kemal Cebeci

Adam Pawlicz

Slagjana Stojanovska

Copyright © 2017 Masters International Danismanlik Arastirma Yayincilik, editors and the authors. All rights reserved. No part of the material protected by this copyright may be reproduced or utilized in any form or by any means, without the prior written permission of the copyright owners, unless the use is a fair dealing for the purpose of private study, research or review. The authors and editors reserve the right that their material can be used for purely educational, scientific and research purposes.

Publisher: Masters International Danismanlik Arastirma Yayincilik
Masters International Consultancy Research and Publishing

ISBN: 978-605-82290-0-6

MIRDEC Publishing

Cinarlicesme sk. No: 21/13 PK: 34303 Kucukcekmece Istanbul Turkey

Publisher certificate no: 35822

Publication date: 30 August 2017

www.mirdec.com

info@mirdec.com

Published at: Form Baskı Teknolojileri Reklam ve Paz. Tic. Ltd. Şti, Şerifali Mah. Şehit Sokak,
No:16/1 Umraniye / Istanbul, Turkey Tel: (+90)(216) 337 37 96, Matbaa Sertifika
No/Certificate no: 31613, www.formbaski.com.tr

MASTERS INTERNATIONAL

Research & Development Center

MIRDEC

MADRID 2017

TABLE OF CONTENTS

AVKASH JADHAV

THE TRANSITION OF POLITICAL IMPERIALISM TO ECONOMIC IMPERIALISM: A HISTORICAL STUDY OF THE GENESIS OF THE NATIVE CAPITALIST CLASS IN THE TEXTILE INDUSTRY OF BOMBAY IN THE 19th CENTURY..... 4

ANTONIO FOCACCI

PROJECT REVERSIBILITY MEASUREMENT TO MANAGE “ADOPTION-WAITING” DECISIONS..... 18

TAMAZ ZUBIASHVILI

GLOBALIZATION AND MIGRATION PROCESSES..... 29

ANDREA RACZ AND ANDREA HOMOKI

RESULTS OF EFFICIENCY AND RESILIENCE MEASUREMENTS IN THE HUNGARIAN CHILD PROTECTION SYSTEM..... 36

AYKUT BEDUK AND NERIMAN CELIK

A RESEARCH ON ASSESSING THE IMPACTS OF HUMAN RESOURCES APPLICATION ON WORK-LIFE BALANCE..... 51

SHERIN MOHAMED FAHMY MOHAMED

“REVOLUTION OF THE DISENFRANCHISED” AND THE DIMINISHING SUPPORT OF THE EGYPTIAN PUBLIC OPINION ON MUSLIM BROTHERHOOD 60

CHI LI CHIU, TANG TSENG CHUNG, DENG HSU TONG, AND CHEN SHEN HOU

EARLY WARNING INDICATORS OF FINANCIAL CRISIS INCIDENCE: EVIDENCE FROM TAIWAN 80

MANJUSHAA BATTLE

GLOBALIZATION AND AGEING POPULATION IN RURAL INDIA: ISSUES AND CHALLENGES..... 85

NARJES BANAYE SHAHANI AND MOHAMMAD HOSEIN NASER BAKHT

METHODS OF ADAPTION FROM IRANIAN WOMEN’S LITERARY WORKS IN DARIUSH MEHRJUYI’S MOVIES 97

NABIL CHERIET , REDOUANE AHNYNE AND GHIZLANE SAAD

CO-CREATION IN ORGANISATIONS 109

CATIA MAGALHAES , EMILIA MARTINS, ROSINA FERNANDES AND FRANCISCO MENDES

USE OF TECHNOLOGY-BASED INTERVENTION FOR PROGRAM DELIVERY AND DISSEMINATION OF FAMILY EVIDENCE-BASED PROGRAMS (EBP)..... 116

MASTERS INTERNATIONAL

Research & Development Center

MIRDEC
International Academic Conference

MIRDEC-4th
International Academic Conference on
Social Science, Multidisciplinary and
Globalization Studies
04-07 July 2017, Madrid, Spain

CONFERENCE PROCEEDINGS

FULL PAPERS

ISBN: 978-605-82290-0-6

Holiday Inn Madrid – Pirámides, Madrid, Spain
04-07 July, 2017

AVKASH JADHAV¹

THE TRANSITION OF POLITICAL IMPERIALISM TO ECONOMIC IMPERIALISM: A HISTORICAL STUDY OF THE GENESIS OF THE NATIVE CAPITALIST CLASS IN THE TEXTILE INDUSTRY OF BOMBAY IN THE 19th CENTURY

Abstract

The advent of the British in India as the East India Company is documented with the arrival of Vasco De Gama when he landed at Calicut in 1498. In the span of four centuries the country witnessed various stages of imperialism, from requesting concessions for trading to acquiring their spheres of influence over certain territories, then declaring certain provinces as their protectorate to establishing it as their presidency. In all we can easily distinguish the changing interests in the way they slowly and gradually unfolded their demands in India. What started as merely exploration of the world soon became a tool of introducing trading designs which ultimately culminated into establishing political hegemony in India. The administrative control of the country became their primary goal until the revolt of 1857, which brought with it the beginning of a new chapter of governance in the colonial history of India. The direct control of the British crown was responsible for changing the priorities of controlling the affairs in the country. We strongly witness the influx of various ideas and patterns of England being experimented, under the ‘mirage of development’ and creating new opportunities for growth. This paper will discuss the introduction of a new native class of Indian capitalist, who equally contributed towards Britain’s economic imperialism in India. This new class was became the native entrepreneurs and the capitalists of the country. The introduction of cotton textile mills in Bombay gave them enough scope to follow the same exploitative imperialist designs of their European masters. It is equally interesting to note that the mill owners of Bombay established their association called ‘Bombay Mill Owners Association’ in 1875, ie within two decades of the opening of the first mill in Bombay in 1854. Whereas the workers working in these mills almost took six decades to realize the importance of forming union or association to safeguard their rights. This paper will primarily discuss that how the native Indian capitalist class slowly replaced the colonial masters in exploiting co Indians under the ‘mirage of development’ which was never inclusive, but exclusive unlike the Europeans.

Keywords: Bombay, mills, working class.

JEL Codes: N85

Acknowledgement: The author would like to express special gratitude to the **Indian Council of Historical Research (ICHR)**, New Delhi India for providing the travel grant to participate and present my research paper as the Keynote speaker in the IVth MIRDEC International Conference(4th -7th July,2017) at Madrid, Spain. I sincerely thank the Member Secretary and the Council members for approving my foreign Travel Grant (FTG).

India has always attracted the attention of the world since ancient times, through the writings of Herodotus to the era of geographical discoveries in medieval age. The European world was always intrigued with the unexplainable wealth of the Far East, which was equally adorned by its rich heritage and culture. The expansion of the West to the East under the pretext of reviving the sea or trade routes opened up the new eco- political developments, there by introducing a new pattern under the guise as the foreign policy which was defined as imperialism. The term ‘imperialism’ came into common usage in England in the 1890s as a development of the older term “empire” by the advocates of a major effort to extend the British Empire.

¹ Associate Professor, Department of History, St. Xavier’s College, Mumbai. India.

The term was rapidly taken into other languages to describe the contest between rival European states to secure colonies and spheres of influence in Africa and Asia, a contest that dominated international politics from the mid-1880s to 1914, and caused this period to be named the “age of imperialism”. The first systematic critique of imperialism was made by the English bourgeois social-reformist and economist John Atkinson Hobson (1858-1940) in his 1902 book ‘Imperialism: A Study’². Imperialism emerged as the development and direct continuation of the fundamental characteristics of capitalism in general. But capitalism only became capitalist imperialism at a definite and very high stage of its development, when certain of its fundamental characteristics began to change into their opposites, when the features of the epoch of transition from capitalism to a higher social and economic system had taken shape and revealed themselves in all spheres. If it were necessary to give the briefest possible definition of imperialism ‘we should have to say that imperialism is the monopoly stage of capitalism’³.

Karl Kautsky, the principal Marxian theoretician defines it as follows: ‘Imperialism is a product of highly developed industrial capitalism. It consists in the striving of every industrial capitalist nation to bring under its control or to annex all large areas of agrarian territory, irrespective of what nations inhabit it’⁴.

The characteristic features of imperialism is precisely that it strives to annex not only agrarian territories, but even most highly industrialised regions (German appetite for Belgium; French appetite for Lorraine), because (1) the fact that the world is already partitioned obliges those contemplating a redivision to reach out for every kind of territory, and (2) an essential feature of imperialism is the rivalry between several great powers in the striving for hegemony, i.e. for the conquest of territory, not so much directly for themselves as to weaken the adversary and undermine his hegemony.

English writer Hobson puts it as, ‘the new imperialism differs from the older, first, in substituting for the ambition of a single growing empire the theory and the practice of competing empires, each motivated by similar lusts of political aggrandisement and commercial gain; secondly, in the dominance of financial or investing over mercantile interests’⁵.

The description of ‘British imperialism’ in Schulze-Gaevernitz’s book *British Imperialism and English Free Trade at the Beginning of the 20th century* reveals the same parasitical traits. The ‘national income of Great Britain approximately doubled from 1865 to 1898, while the income from abroad increased nine fold in the same period’⁶. The US President Abraham Lincoln believed ‘When the white man governs himself, that is self-government; but when he governs himself and also governs others, it is no longer self-government; it is despotism’⁷. In its economic essence imperialism is monopoly capitalism. This in itself determines its place in history, ‘for monopoly that grows out of the soil of free competition, and precisely out of free competition, is the transition from the capitalist system to a higher socio-economic order’⁸.

We need to understand the different aspects of monopoly; firstly, monopoly arose out of the concentration of production at a very high stage. This refers to the monopolist capitalist associations, cartels, syndicates and trusts. We have seen the important part these play in present-day economic life. Secondly, monopolies have stimulated the seizure of the most important sources of raw materials, especially for the basic and most highly cartelised industries in capitalist society: the coal and the iron

² Lenin.V.I, Imperialism the Highest stage of Capitalism, Resistance Marxist Library, Resistance Books , Australia, 1999, p. 7.

³ Ibid. p. 91.

⁴ Schulze-Gaevernitz, British Imperialism and English Free Trade at the Beginning of the 20th Century, Leipzig, 1906, p. 318.

⁵ Siegmund Schilder, Trends of Development of World Economy, Berlin, pp. 1912. 160-161.

⁶ Schulze Op.cit. p. 104

⁷ Patouillet O J , L’impérialisme Américain, Dijon, 1904, p. 272.

⁸ Lenin. Op.cit. p. 104.

industries. Thirdly, monopoly has sprung from the banks. 'The banks have developed from humble middleman enterprises into the monopolists of finance capital'⁹. The British imperialism was more pragmatic than that of other colonial powers. Its motivation was economic, not evangelical. The main changes which the British made in Indian society were at the top, but the lower strata were thoroughly exploited. They replaced the wasteful warlord aristocracy by a bureaucratic-military establishment, carefully designed by utilitarian technocrats, which was very efficient in maintaining law and order. The greater efficiency of government permitted a substantial reduction in the fiscal burden, and a bigger share of the national product was available for landlords, capitalists and the new professional classes. Some of this upper class income was siphoned off to the UK, but the bulk was spent in India. However, the pattern of consumption changed as the new upper class no longer kept harems and palaces, nor did they wear fine muslins and damascened swords. We witness the emergence of the new towns and urban amenities with segregated suburbs and housing for them were created. Their habits were copied by the new professional elite of lawyers, doctors, teachers, journalists and businessmen. Within this group, old caste barriers were eased and social mobility increased.

The striking thing about the British raj is that it was operated by few people. There were only 31,000 British in India in 1805 (of which 22,000 were in the army and 2,000 in civil government)¹⁰. The number increased substantially after the revolt of 1857, but thereafter remained steady. In 1911, there were 164,000 British (106,000 employed, of which 66,000 were in the army and police and 4,000 in civil government)¹¹.

In the Mughal period or medieval India, the zamindars would usually keep a tenth of the land revenue to themselves, but by the end of the British rule their income from rents was a multiple of the tax they paid to the state. In Bihar, for instance, it was five-sixths of the total sum levied was rent by 1850 and only one-sixth revenue¹². Under the British, transfers became much more frequent, particularly into the hands of moneylenders. The moneylenders frequently presented as squeezing out poor peasants and tenantry and thus promoted the concentration of wealth, but the evidence of what happened to zamindar estates suggests that 'village moneylenders may also have helped to break up concentrations of wealth'¹³. Nevertheless, there were some economic consequences of the new legal situation as because of the emergence of clear titles under the British rule, it was now possible for the Indian farmers to mortgage their land. The status of moneylenders also improved due to the change from the Muslim law to the British law. Though the moneylenders were there during the Mughal period, but 'their importance grew substantially under the British rule, and over time a considerable amount of land changed hands through foreclosures'¹⁴.

It was close to 1870's, India built up her own textile manufacturing industry which displaced the British imports. The British imports entered India duty free, and when a small tariff was required for revenue purposes Lancashire pressure led to the imposition of a corresponding excise duty on Indian products to prevent them gaining a competitive advantage. This undoubtedly handicapped industrial development. If India had been politically independent, her tax structure would probably have been different. In the '1880s, Indian customs revenues were only 2.2 per cent of the trade turnover, i.e. the lowest ratio in any country. In Brazil, by contrast, import duties at that period were 21 per cent of trade turnover'¹⁵. Indian firms in industry, insurance and banking were given a boost from 1905 onwards by the swadeshi movement, which was a nationalist boycott of British goods in favour of Indian enterprise. During the

⁹ Lenin. Op.cit. p. 120.

¹⁰ D.A.B. Bhattacharya (ed.), Report on the Population Estimates of India (1820-30), Census of India 1961, Government of India, Registrar General, Delhi, 1963, pp. 4-5.

¹¹ Census of India 1911, Vol. I, India, Part II, Tables, Calcutta, 1913, pp. 374-6.

¹² D. Warriner, Land Reform in Principle and Practice, Oxford University Press, 1969, p. 158.

¹³ W.C. Neale, Economic Change in Rural India, Yale University Press, 1962, p. 63.

¹⁴ M.L. Darling, The Punjab Peasant in Prosperity and Debt, Oxford University Press, London, 1947, p. 178.

¹⁵ M.G. Mulhall, The Dictionary of Statistics, Routledge, London, 1899, pp. 172 and 258.

First World War, lack of British imports strengthened the hold of Indian firms on the home market for textiles and steel. After the First World War, under the nationalist pressure, the government started to favour Indian enterprise in its purchase of stores and it agreed to create a tariff commission in 1921 which started raising tariffs for protective reasons. By 1925, the average tariff level was 14 per cent compared with 5 per cent pre-war¹⁶. The procedure for fixing tariffs was lengthy and tariff protection was granted more readily to foreign-owned than to Indian firms, but in the 1930s protection was sharply increased¹⁷.

From the beginning of the British conquest in 1757 to its independence in 1947, it seems unlikely that per capita income could have increased by more than a third and it probably did not increase at all. In the UK it there was a tenfold increase in the per capita income over these two centuries. The most noticeable change in the economy was the rise in population from about 170 million to 420 million from 1757 to 1947 in India. However, there were some significant changes in social structure and in the pattern of output. The social pyramid was truncated because the British lopped off most of the top three layers of the Mughal hierarchy, i.e. the Mughal court, the Mughal aristocracy and quasi-autonomous princes (a quarter of these survived), and the local chieftainry (zamindars who survived in about 40 per cent of India). In place of these people the British installed a modern bureaucracy which took a smaller share of national income. The newcomers had a more modest life-style than the Mughals, but siphoned a large part of their savings out of the country and provided almost no market for India's luxury handicrafts. The modern factory sector which they created produced only 7.5 per cent of national income at the end of British rule and thus did little more than replace the old luxury handicrafts and part of the village textile production.

The British reduced the tax squeeze on agriculture and turned warlords into landlords, but the new order had little dynamism. A good deal of the old fuzziness about property rights remained, and landlords were still largely parasitic. The bigger zamindars copied the Mughal lifestyle by maintaining hordes of retainers and huge mansions, the smaller landowner's ambition was to stop working and enhance his ritual purity by establishing a seedy gentility, very little incentive was provided for investment and almost nothing was done to promote technical changes in agriculture. At the bottom of society the position of sharecropping tenantry and landless labourers remained wretched.

In urban areas a new westernized 'middle class' of Indians emerged and became the major challenge to the British raj. The British were a very thin layer at the top of society but they took about 5 per cent of national income. Their allies, the native princes and zamindars, took about 3 per cent, eight per cent is a sizeable proportion for the ruling class but, under the Mughal regime, the equivalent group collected 15 per cent of national income in taxes and spent most of it on their own consumption. Immediately under this group were two new indigenous classes - capitalists and professionals - who became the dominant elite in independent India. They were relatively larger in number and probably had a bigger share of national income than their counterparts. Within the village society the social structure was probably similar to that in Mughal India, with the two top economic groups corresponding to the old dominant castes, the next group to peasant castes, and the bottom group consisting largely of untouchables.

The main difference from the Mughal economy is that village proprietors and tenants-in-chief were no longer heavily squeezed by taxation and their share of national income had probably increased. Thus the main gainers from the British regime (apart from the British) were the so called 'middle' class of Indian capitalists and professionals, and the village hierarchy. Most of these were high caste Hindus though the Parsis and Sikhs did fairly well. The main losers were the Muslims who had formed the major part of the Mughal aristocracy, officer corps, lawyers, and artisans in the luxury handicrafts.

¹⁶ W.A. Lewis, *Economic Survey, 1919-39*, Allen and Unwin, London, 1949, p. 48.

¹⁷ M. Kidron, I, Oxford University Press, London, 1965, p. 13.

The development of capitalism in India was therefore a very tortuous process. The Indian working-class movement too consequently also developed at a much later stage. The development of capitalism, so to say, a change-over from the old feudal economic order to the modern capitalist economy was a long drawn-out process starting from the mid 16th century to 18th century, the characteristics form of capitalist production was to manufacture for pure profit. In the era before manufacture also, the workers depended on selling their labour-power, but they still had the real chance of eventually becoming independent. But the manufacture era involved extensive division of labour between many workers concentrated under one roof. The social division of labour and specialization of functions led to the downgrading and subordination of the individual who became a completely isolated component, cut off from the process of production as a whole and subjected to rigorous discipline. All that the worker required was a highly specialized dexterity losing his general skill as a craftsman and his ability for independent work. The products of seasoned craftsmanship in the era before manufacture transformed itself into the products of 'collective labour' of a few skilled and many unskilled workers in the era of manufacture. However, it was characterized not only by the invention and introduction of machinery, but also by the appearance of new basic classes, 'the bourgeoisie who owned the factories and the means of production and exploited the workers, and the proletariat, i.e. the wage-laboures who did not possess the means of production'¹⁸.

The industrial revolution had therefore two aspects-the technical, i.e. the invention and the introduction of machinery and the passing away of the old feudal order and the formation of new basic social classes-the bourgeoisie and the proletariat in the newly emerging capitalist economic order. But in this continuously expanding revolution with the machine as the starting-point, the human organ was superseded by mechanical organization, independent of the limitations of human power. This totally transformed the whole production process. At the beginning of this process, in manufacture era the organization of social labour was purely subjective in the sense that it required a combination of different operations, but the new modern industry had in machinery a purely objective productive organism which converted the labour into a mere appendage of an already existing material condition of production¹⁹.

It is interesting to note that when Marx analysed the Indian society, industrial revolution had already taken place in England and capitalism was expanding from England to the other European countries. This traditional system of Indian industry and agriculture was, however, laid waste by the imperialist plunder age. The establishment of the railways and certain connected industries in furtherance of colonial interest completed the process. In the first volume of 'Capital' Marx presents a vivid description of this old economic and social system.

'These small and extremely ancient Indian communities, some of which have continued down to this day, are based on possession in common of the land, on the blending of agriculture and handicrafts, and on the unalterable division of labour, which serves, whenever a new community is started, as a plan and scheme ready cut and dried. Occupying areas of from 100 up to several thousand acres, each forms a compact whole producing all it requires. The chief part of the products is destined for direct use by the community itself and does not take the form of a commodity. Hence, production here is independent of that division of labour brought about in Indian society as a whole, by means of the exchange of commodities. It is the surplus alone that becomes a commodity and a portion of even that, not until it has reached the hands of the State, into whose hands from time immemorial a certain quantity of these products has found its way in the shape of rent in kind. The constitution of these communities varies in different parts of India. In those of the simplest form, the land is tilled in common and the produce divided among the members. At the same time spinning and weaving are carried on in each family as subsidiary industries. Side by side with the masses thus occupied with one and the same work, we find the 'Chief inhabitant', who is judge, police and tax-gatherer in one; the book-keeper, who keeps the

¹⁸ Sukomal Sen Working class of India. History of Emergence and Movement 1830-1970, published by K.P. Bagchi & Company, Calcutta, 1970. pp. 1-2.

¹⁹ Ibid. p. 3.

amounts of the village and registers everything relating thereto; another official who prosecutes criminals, protects strangers travelling through and escorts them to the next village; the boundary man, who guards the boundaries against neighbouring communities; the water-overseer, who distributes the water from the common tanks for irrigation; the Brahmin, who conducts the religious services; the schoolmaster who on the sand teaches the children reading and writing; the calendar-Brahmin, or astrologer, who makes known the lucky or unlucky days for seed-time and harvest, and for every other kind of agricultural work; a smith and a carpenter, who make and repair all the agricultural implements; the potter, who makes all the pottery of the village; the barber, the washer man, who washes clothes, the silversmith, here and there the poet, who in some communities replaces the silversmith, in others the school-master. This dozen of individuals is maintained at the expense of the whole community. If the population increases, a new community is founded, on the pattern of the old one, on unoccupied land. The whole mechanism discloses a systematic division of labour; but a division like that in manufacturer is impossible, since the smith and the carpenter find an unchanging market, and at the most there occur, according to the sizes of the villages, two or three of each, instead of one. The law that regulates the division of labour in the community acts with irresistible authority of law of nature, at the same time that each individual artificer, the smith, the carpenter, and so on conducts in his workshop all the operations of his handicraft in the traditional way, but independently and without recognizing any authority over him. The simplicity of the organization for production in these self-sufficient communities that constantly reproduces themselves in the same form, and when accidentally destroyed, spring up again on the spot and with the same name-this simplicity supplies the key to the secret of the unchangeableness of Asiatic societies, an unchangeableness in such striking contrast with the constant dissolution and refounding of Asiatic contrast states, and the never-ceasing changes of dynasty. The structure of the economical elements of society remains untouched by the storm-clouds of the political sky'²⁰.

The colonial rule and the exploitation by the British imperialists completely ruined the system of production of these traditional and self-sufficient societies. As the British army advanced and occupied different territories of India as a sequel to the victory in the battle of Plassey in 1757, the old economic system and social divisions of labour obtaining in those territories were also shattered simultaneously. Along with this occupation, the surplus products of the occupied zones also fell into the hands of the colonialists who then started direct plundering of and exporting the wealth of India to England. Imposing a high rate of taxation on internal trade of India and simultaneously engaging itself in money-lending business, the East India Company extorted a huge sum of money from the Indian people.

Referring to this direct plunder Karl Marx observed 'during the whole course of the 18th century the treasures transported from India to England were gained much less by comparatively insignificant commerce, than by the direct exploitation of that country, and by the colossal fortunes there extorted and transmitted to England'²¹. The sweat and blood of the Indian people, reduced to money became one of the principal sources of the primitive accumulation of capital in Britain. According to obviously minimized statistics, the British colonialists derived from India during the period of 55 years between 1757 and 1812 a direct income exceeding 100,000,000. Conspiratorial military victory and plundering of the wealth of the conquered areas continued for several decades. Subsequently the colonialists realized that the exploitation should be regulated and legalized in some way to ensure permanent revenues and consolidations of British rule, Data of Committee of Correspondence, submitted to the Board of Directors of the East India Company, February 9, 1813²².

²⁰ Karl Marx, Capital, Vol 1, Foreign Languages Publishing House, Moscow, 1954, pp. 357-58.

²¹ Karl Marx, The East India Company-Its History And Results, contained in On Colonialism by K. Marx and F. Engels, Foreign Languages Publishing House, Moscow, p. 51.

²² Hansard's Parliamentary Debates, Vol. 25, p. 28, quoted from Capitalism in India-Basic Trends in its Development, Peoples' Publishing House, New Delhi, by A. I. Levkovsky, p. 10.

In 1793 during the viceroyalty of Lord Cornwallis, the British colonialists passed the Act of Permanent Settlement, fixing a constant rate of taxation to be exacted from the zamindars of Eastern India. The 1793 Act and subsequent Acts conferred the formal rights of land-ownership to none but the zamindars, the hereditary rights of village community members and petty feudal lords were not recognized. This system at once dispossessed the people of Bengal Presidency of their hereditary claims to the soil in favour of the native tax-gatherers called zamindars. Despite securing a kind of right to land-ownership, the feudal zamindars were compelled to turn over ten-eleventh of the rent to the British colonial state, failing which, the state sold his land to anyone who could pay the sum. So in a bid to satisfy the colonial rulers and also for self-aggrandizement, the zamindars intensified exploitation of the peasantry and not infrequently raised it to inhuman heights. While their own instalments payable to the government were fixed, the zamindars enjoyed the freedom of increasing the rate of taxation on the peasantry. With the disappearance of the ancient Indian feudal nobility through this process, a new stratum of landlords originated from the moneylenders, tradesmen and colonial officials appeared on the scene. A series of parasitic middlemen also sprouted out between the cultivator and the zamindars each one of whom sought to extort his own pound of flesh thereby sharpening the exploitation of the peasantry beyond all proportions.

In the south of India the colonialists introduced the ryotwari system to exploit the peasantry. The peasant became a permanent tenant of a plot of land, a toiler of the soil entangled with obligations and in fact chained to the land as a serf. Thus the zamindari and the ryotwari systems were the two principal ways in which the modified feudal methods of exploiting the peasants were preserved. These systems while fully sub served the interests of the colonialists degraded the peasantry to the position of semi-slaves. Simultaneously with the exporting of the plundered wealth of India to England in furtherance of her Industrial Revolution, the English capitalists felt the need of marketing in India the industrial products of England. This in other words meant a free trade with India. The exclusive monopoly of trade with India so long enjoyed by the East India Company since the battle of Plassey in 1757 did not protect the interest of free trade of the British capitalist class as a whole. Prompted by an energetic search for new markets, the English bourgeoisie unleashed a large-scale campaign to abolish the privileges of the East India Company.

In 1813 the East India Company's monopoly in the trade with India was abolished opening the door of free trade with India. This in fact indicated a new phase in the economic exploitation of India. The East India Company had so long been earning profits mainly by exporting to and selling Indian silk, muslin and other luxury goods in England, but after 1813 Indian market was laid open to the British industrial commodities resulting in a rapid increase of British exports to India. 'From 1,600,000 in 1814 it grew to 5,800,000 in 1828, or one-eighth of all British export. The total tonnage of British ships engaged in trade with India in 1828 reached nearly 110,000 tons. In 1814 Britain shipped 213,000 yards of plain and 800,000 yards of coloured cotton textiles to India; in 1826 the totals were 16 and 26 millions yards respectively'²³. The process of development of capitalism over the ruins of feudalism as it was in the case of Europe was not to be found in India. Although the Imperialist rulers devastated the old Indian economy, they did not supplant it by unleashing the forces of modern capitalist economy. So, the growth of capitalist economy in India followed a different path with accompaniment of strange contradictions, impediments and untold sufferings for the Indian people. Despite this, British rule in India produced two kinds of results, one destructive and the other regenerating. To consolidate colonial exploitation, it on the one hand annihilated the old Asiatic society and on the other was constrained to take some steps, the objective consequences of which rendered the growth of capitalist economy irresistible, although through a very halting and painful course.

²³ A. I. Levkovsky, Capitalism in India-Basic Trends in its Development, Peoples' Publishing House, New Delhi, p. 19.

Introduction of railways in India was the event of foremost importance in this process. For transportation of the goods imported from England from the ports of India to the interior of the country, for carrying the raw materials from the countryside to the ports, for quick movement of the army to suppress the Indian people through military might-in a word to consolidate colonial rule and exploitation, introduction of railways in India turned out to be an indispensable task for the British imperialists.

In 1853 Viceroy, Lord Dalhousie, wrote his 'famous minute' pointing out the great social, political and commercial advantages to be gained from connecting the three Presidency cities-and these with the north-west frontier-by rail'. It is pertinent to quote here Marx's famous saying regarding the future results of British rule in India as he keenly observed this new aspect. With an amazing capacity of foresight he wrote: "I know that the English millocracy intend to endow India with railways with the exclusive view of extracting at diminished expenses the cotton and other raw materials for their manufacturers. But when you have once introduced machinery into the locomotion of a country, which possesses iron and coals, you are unable to withhold it from its fabrication. You cannot maintain a net of railways over an immense country without introducing all those industrial processes necessary to meet the immediate and current wants of railway locomotion and out of which there must grow the application of machinery to those branches of industry not immediately connected with railways. The railway system will therefore, become, in India, truly the forerunner of modern industry'²⁴.

The exploitation of the Indian working class was expressed chiefly in the fact that both British and Indian capitalists secured absolute surplus value. The working hours of maximum length, from dawn to dusk and often even longer was the most striking indication of the brutal manner in which labour was exploited. Even the official report of the Indian Factory Labour Commission which was appointed in 1908 to enquire into various recommendations made by the Freer Smith Committee in respect of certain amendments in the existing Factory Act, could not hide this inhuman picture. According to this report, in Ahmedabad the average working period in a day was 12 hours, and at some factories using electric power it was no less than 14 hours. In 'Bombay also the average was 12 hours, but in 60 out of 85 cotton mills where electricity was used, the labourers had to work not less than 13 to 15 hours. In Broach, Gujarat division the working period lasted 14 ½ to 13 ½ hours, in Delhi 13 ½ to 14 ½ hours in Agra it ranged from 13 hours 45 minutes to 15 hours 15 minutes, in Amritsar and Lahore from 13 to 13 hours 40 minutes. But the British capitalists owning the jute mills of Calcutta set the record making the weavers of these mills worked for 15 hours and also from 15 ½ to 16 hours in some cases²⁵. The employers did not show any sense of proportion or any human consideration in exploiting female and child labour. Children even of such tender age as between 5 years and 7 years were employed most cruelly everywhere. Investigations conducted by the Indian Factory Labour Commission of 1908 revealed that half of the time, 30 to 40 per cent of those employed in the factories were tender-aged children.

The Report of the Royal Commission on Labour in India, 1931 also testified, 'When the Factory Commission of 1908 made the investigations, many textiles mills were working from 13 to 15 hours a day with a single set of workers, and before that this practice had been fairly general'²⁶. This was further confirmed by Mr. N. A. Moss, the Chief Inspector of Factories, Bombay, according to whom, 'Strikes have been many two should be put down every year for each factory, but all of them have been short-lived and in the end it is always the operatives who have given in, in some cases with fines and in some

²⁴ Karl Marx, The Future Results of the British Rule in India, contained in On Colonialism by K. Marx and F. Engels, Foreign Languages Publishing House, Moscow, p. 87.

²⁵ Sen.op.cit. pp. 37-38.

²⁶ Ibid. p. 40

cases with loss of arrears of wages. The reasons leading the men to strike are mostly temporary, reduction of wages without any notice whatsoever²⁷.

Class-consciousness of the proletariat, as a general rule, rises in proportion to the advance of economic struggles against capitalist exploitation. But in a colony, some specific features of exploitation as distinct from that in a metropolitan country impede the development of economic struggles of the workers and in consequences retard the growth of their class political consciousness. The general nature of capitalist exploitation in colonial India was thus rendered further oppressive by these specific features.

A colonial regime means the open dictatorship of imperialism in its crudest forms, implying thereby that in a colony the oppression of the working class grows immeasurably, assuming the most ugly and monstrous character. In the context of colonial India it is not only national oppression to which, in addition to social oppression, the working class was subjected as a section of the entire Indian people, the capitalist exploitation in itself was so altered that its burden on the working class increased tremendously. Successful economic struggle is admittedly one of the main factors determining the conditions for the sale of labour-power and the degree of capitalist exploitation. Further, 'this struggle puts a limit on the otherwise boundless quest for profit to which capital is impelled by competition'²⁸. When the capitalists were confronted with growing resistance from the workers, a tendency on the part of the employers and the government officials towards presenting a little exaggerated view of the situation cannot, however, be ruled out-in this respect, the study made by the Factory Labour Commission of the existing labour condition seems to be more balanced.

According to the Commission, 'The history of movement in Bombay and of similar movements in other industrial centres shows clearly that while the operatives fully understand the machinery of local strikes and have repeatedly forced employers to comply with their demands in isolated cases, they are as yet unable to combine over any large area with the object of securing a common end by concerted action'²⁹. The history of all the countries shows that the working class exclusively by its own effort, is able to develop only trade-union consciousness, i.e. 'the conviction that it is necessary to combine in unions, fight the employers, and strive to compel the government to pass necessary legislation, etc'³⁰. Japan's bid to capture market as a formidable rival to western capital further worsened the condition of India's industry. This cut-throat competition of international capital combined with extreme impoverishment of the Indian people resulted in a serious shrinkage of the country's market. Confronted with this unfavourable situation, the Indian textile mill owners conveniently directed attacks on the working class in a bid to strengthen their own position. The mill owners attempted 'to reduce wages of the workers. It is a particular misfortune of the colonial working class that they have ultimately to fall victim to the intense rivalry between the imperialists and native capitalists'³¹. Economic development is as much concerned with human and institutional aspects as its more emphasized aspects of capital-formation and better exploitation of the resources of a given country. Hence in a developing economy (or a country aspiring for economic development) 'it is highly essential to minutely examine all the existing and potential forms of human institutions so as to discover their impacts upon the economic growth of the community'³².

Although both industrial and agricultural workers are affected by this process of capital accumulation, usually the industrial workers first try to protect themselves from the impact of the strains of capital

²⁷ British Parliamentary Paper, XXXVI, Vol. II, Part V, 1892, p. 120, British Parliamentary Paper, XXXVI, Vol. II, Part V, 1892, p. 107, British Parliamentary Paper, XXXVI, Vol. II, Part V, 1892, p. 107.

²⁸ Sen.op.cit. pp. 87-88.

²⁹ Report of the Factory Labour Committee, 1906, 1907.

³⁰ V. I. Lenin, Collected Works, Moscow, 1964, Vol. 5, p. 375.

³¹ Sen.op.cit. p. 205.

³² Subrathesh Ghosh, Trade Unionism In The Underdeveloped Countries, published by Book and Private Ltd, Calcutta, 1960. p 3.

accumulation, since in the urban society they have to come more directly in contact with the rich industrialists and other people belonging to the upper income brackets and hence, are more exposed to what is now known as the demonstration effect. The high standard of living and the conspicuous consumption by the rich in the urban society make them more conscious about their own miseries and intensify their discontent. And in spite of higher money wages 'the loss of the accustomed way of life which the new recruits to the industries in the developing economy used to enjoy in their rural society together with the great insecurity of the industrial life make their condition more intolerable an compared to those of the agricultural workers'³³.

Machinery-inventions made large-scale production an easy possibility. But this large-scale production, of course, necessitated large investments in machinery plantations, which brought into existence a class of men, who could make such investments, the class of capitalists. Mechanical production and output far superseded that of the home industries and village or guild-industries of the Medieval Ages. The ruin of these industries threw the old independent guild-labour-hands out of employment and thus brought them to the feet of the capitalists, who could propose their own terms to the labourers. The 'labourers could do nothing but offer themselves to these new masters on their conditions'³⁴. The possibility to carry on unlimited production through machinery in a very short time, gave to the capitalists means of making vast profits, which intensified the feeling of acquisitiveness, vanity, rivalry and love of power. This made them disregard the condition of the labourers that they employed and whom in course of time they began to consider as another piece of machinery.

The old lords, serfs and slaves were abolished; but new kinds of lords and slaves came into existence, without those obnoxious titles, under the name of capitalists and wage-earners. The whole wealth thus obtained by starving the labourers at home and ruining labourers abroad, through competition, went to satisfy the lust of the capitalists. The labourers were men as much as the capital-owners were. Political parties and state mechanisms with their pretention of democratic representations were dominated by their purses. Thus becoming 'masters of the political wheel, which alone is competent to effect reform in society, they could suppress the cry of lessening the miseries of the working-class'³⁵.

The capitalists that is the possessors of the means and implements of labour, namely lands, factories, ready money and raw material; contractors that is the heads and initiators of labour, commercial men, who represent or ought to represent intellect and the working men, who represent manual labour. The capitalists have become 'the masters of the new slaves, who are not given the rights of human beings, even'³⁶. Capital cares nothing for the length of life of labour power. All that concerns it is, simply and solely the maximum of labour power that can be rendered fluent in a working day. It attains this end by 'shortening the extent of the labourer's life as a greedy farmer snatches increased produce from the soil by robbing it of its fertility'³⁷. Large populated areas in industrial towns well exhibit to what level of life wage-earners was being reduced. The industrial society began to breed the class of capitalists, which was the source of so much evil in Europe. When the cost of living rose extraordinarily high in times of war, the Indian capitalists showed as much implacability towards the demands of starving Indian labour as the European capitalists did. White capitalists can at least be excused on the ground that it is in their very nature and breeding to behave so towards the Indians, but the Indian capitalists equally followed their path. Most of the trade in foreign and inland was centered in the hands of the Shethias of the Gujarathi community, the Marwaris, the Parsis and the Bohras. These capitalist communities were naturally opposed to the attempts of native Indian workers emancipation. And they in their turn

³³ Ghosh. op.cit.pp. 21-22

³⁴ Bani Deshpande, Roza Deshpande, Umakant Mokashi, (ed), Selected Writings : S. A. Dange, Lok Vangmaya Griha publication vol-1 Bombay, 1974. p. 63.

³⁵ Ibid. pp. 63-65.

³⁶ Ibid. p. 66.

³⁷ Ibid. p. 66.

‘exhibited all the greediness, idleness and cruelty, luxurious and demoralized life consequent upon capitalism in every form and in every country’³⁸.

Thus the Indian capitalists were committing three sins. ‘They supported the foreign despotism, they demoralize and ruin the peasantry and the wage-earning classes of the society, by doing this they supported and fed the capitalists of Europe and thus helped the cause of misery of the workers of that continent also’³⁹.

The instruments like the mills in Bombay were not merely for producing cloth; they also had special, social significance. They were instruments of exploiting wage-labourers. The capitalist obtained their profits because they owned the means of production which ultimately meant capital. Thus the distribution of the means of production determined the distribution of the products. The owner of the means of producing cloth had all of it while to the share of the worker falls little of it. It was ‘these class relations which determined in the first place the outline of society, its economic structure’⁴⁰.

In the modern imperialist epoch it requires very little arguments to prove that the State conforms to the economic structure. In capitalists society, ‘the capitalists control the means of production, naturally they control the State also’⁴¹. The British imperialism in India deprived the feudal order of its political power, but retained its socio-economic character, making it serve the needs of the imperialist country. To ‘serve the needs of British industry means to serve as its suppliers of raw materials and markets’⁴².

The ‘vast wealth taken from India had given a stable basis of liquid capital to British industries, but it disturbed state of affairs in India, and unregulated rapacious Company control, wherein the traders were directly administrators both of the political rule and commercial development’⁴³. The handicraft and manufactures were destroyed first by extra-economic force and violent destruction and a vast number of artisans were thrown out of the land. The character of agriculture was thoroughly changed. The growing of crops was subjected to the needs of the exchange market and the peasants economy was brought within the orbit of capital it market. The low level of productive forces, ‘the poor national income was burdened with an expensive bureaucracy and disproportionate militarism, the resulting discrepancy was filled up by high taxation and public debt’⁴⁴.

A highly industrialized society requires a great deal of accounting to be done. The needs of modern production and distribution have to carry out the inventory of the world in all matters. Without the ‘highly organized system of accounting, modern large scale production and international exchange of goods would not be possible’⁴⁵. The Indian worker had to work in such inhuman conditions that for all time there was a permanent fund of grievances justifying a strike. The material conditions of the working-class, on the admission of responsible commissions, were forcing them into class struggle. While in capitalist Europe they secured the 8-hour day and in Soviet Russia they were working even a 7-hour day, but in India during the same time they had to work for minimum 10 hours a day and in the native states it was extended to 14 hours also. He had ‘no right to fall sick and become old and if he does, he must starve and die, there was no insurance for him’⁴⁶.

³⁸ Ibid. pp. 71-72.

³⁹ Ibid. p. 73.

⁴⁰ Ibid. p. 34.

⁴¹ Ibid. p. 36.

⁴² Ibid. p. 319.

⁴³ Ibid. p. 325

⁴⁴ Ibid. p. 328

⁴⁵ Ibid. p. 461

⁴⁶ Ibid. p. 478.

The workers assembling in modern industrial sectors were mainly expropriated peasant class from villages, to which they were linked with family ties or through their holdings of increasingly smaller pieces of land heavily encumbered with debt. The poorest and the most downtrodden and the menials were the first to leave the villages, as they found their roots threatened and had to face a precarious survival commensurate with the growing misery in the rural India. The emerging working class had to face crude exploitative control of the employers which had its roots somewhere in the character of a despotic social set up supported from past as well as the very logical need of modern capitalism. The exploitation of the working class reached an extreme step due to the dual socio-economic pressure generated from the well entrenched feudal social set up and "Commoditism" of the modern capitalism⁴⁷. The replacement of the man with machine made his survival merely mechanical and the native Indian capitalists conveniently replaced the British rule and role even after the independence of India.

References List

A R Desai, Labour Movement in India, Documents: 1918-1920 Indian Council of Historical Research, Popular Prakashan, New Delhi, Vol. No.I,II,III. 1989.

Bani Deshpande, Roza Deshpande, Umakant Mokashi, (ed), Selected Writings :S.A.Dange, Lok Vangmaya Griha publication Vol- I,II & III, Bombay, 1974.

British Parliamentary Paper, XXXVI, Vol. II, Part V, 1892,p. 120, British Parliamentary Paper, XXXVI, Vol. II, Part V, 1892.

Census of India 1911, Vol. I, India, Part II, Tables, Calcutta, 1913.

D.A.B. Bhattacharya (ed.), Report on the Population Estimates of India (1820-30), Census of India 1961, Government of India, Registrar General, Delhi, 1963.

D. Warriner, Land Reform in Principle and Practice, Oxford University Press, 1969.

Dipak Malik, Indian Trade Unionism in Development Perspective, Commonwealth publication, New Delhi, 1989,

Economic and Political Weekly: 1970-2011.

Gopal Ghosh, Indian Trade Union Movement, T U publication, Calcutta, First Part, 1963.

Hansard's Parliamentary Debates, Vol. 25,p. 28,quoted from Capitalism in India-Basic Trends in its Development, Peoples' Publishing House, New Delhi.

Indian Journal of Economics, published by the Department of Economics and Commerce, University of Allahabad.1920-1940.

Indian Annual Register, 1920-1940.

Karl Marx, Capital, Vol 1, Foreign Languages Publishing House, Moscow, 1954.

Lenin.V.I, Imperialism the Highest stage of Capitalism, Resistance Marxist Library, Resistance Books, Australia, 1999.

⁴⁷ Dipak Malik, Indian Trade Unionism in Development Perspective, publisher Commonwealth, New Delhi.1989. p 41.

- M.L. Darling, The Punjab Peasant in Prosperity and Debt, Oxford University Press, London, 1947.
- M.G. Mulhall, The Dictionary of Statistics, Routledge, London, 1899.
- M. Kidron, I, Oxford University Press, London, 1965.
- Manorama Savur and Kamala Ganesh (ed), Labour Movement in India 1937-1939 , Pragati publications, Indian Council of Historical Research, New Delhi Vol.17. 2005
- Morris David Morris, The Emergence of an Industrial Labour Force In India, A Study of the Bombay Cotton Mills, 1854-1947. Oxford University Press (OUP), Bombay , 1965.
- N M Joshi, Trade Union Movement in India. Private Papers, Nehru Memorial Library, New Delhi.
- Patouillet O J , L'impérialisme Américain, Dijon, 1904.
- P D Kulkarni, "Textile Trade Unionism in Bombay", The Indian Journal of Social Work (ed), The faculty of Tata Institute of Social Sciences, Bombay, No.3, vol.VII December,1946-47.
- Radhakamal Mukherjee ,The Indian Working Class, Hind Kitabs ltd, Bombay second edition, 1948.
- Report of the Bombay Enquiry Committee, Labour Gazette, Vol. XIX, September 1920- 39.
- Report of the Factory Labour Committee, 1906, 1907.
- Report of the Indian Factory Labour Commission, September, 1890, under the orders of His Excellency, the Governor General in Council, with Proceedings and Appendices, Calcutta, Office of the Superintendent of Government Printing , 1890.Vol. II.
- Reports of The Bombay Millowners Association, Report of The Bombay Millowners Association, 1880-1920.
- Report from the Commissioner of Police, Bombay, dated the 12 April 1924 to the Secretary to the Government of Bombay, Home Dept. File No. 55, Govt. of India.
- Richard Newman, Workers and Unions in Bombay 1918-1929. A study of Organization in the Cotton Mills, Australian National University, Canberra, 1981.
- Revri Chamanlal, The Indian Trade Union Movement. An outline History-1880-1947, Orient Longman, New Delhi-1972.
- Schulze-Gaevernitz, British Imperialism and English Free Trade at the Beginning of the 20th Century, Leipzig, 1906
- Shapurji Saklatwala. Shapurji, A Few Thoughts on Party Work, Private Papers, Nehru Memorial Library.
- Siegmund Schilder, Trends of Development of World Economy, Berlin, 1912.
- Subrathesh Ghosh, Trade Unionism In The Underdeveloped Countries. published by Book and Private Ltd, Calcutta, 1960.

Sukomal Sen, Working class of India History of Emergence and Movement 1830-1970, published by K.P. Bagchi & Company, Calcutta, 1970

The Indian Journal of Social Work (ed), The faculty of Tata Institute of Social Sciences, Bombay, 1930-45.

V.B Karnik, Indian Trade Unions A Survey, published by Manaktalas, Bombay, October 1988.

V. I. Lenin, Collected Works, Moscow, 1964.

W.C. Neale, Economic Change in Rural India, Yale University Press, 1962.

W.A. Lewis, Economic Survey, 1919-39, Allen and Unwin, London, 1949.

Masters International Research & Development Center

MIRDEC
International Academic Conference

MIRDEC-4th,
International Academic Conference on
Social Science, Multidisciplinary and Globalization
Studies

CONFERENCE PROCEEDINGS

FULL PAPER SERIES

Editors

Kemal Cebeci
Adam Pawlicz
Slagjana Stojanovska

ISBN: 978-605-82290-0-6

Holiday Inn Madrid – Pirámides, Madrid, Spain
04-07 July, 2017

We are very pleased to introduce the proceedings(full paper series) of the Masters International Research and Development Center, MIRDEC 4th, International Academic Conference on Social Science, Multidisciplinary and Globalization Studies, 04-07 July 2017, Madrid, Spain. MIRDEC thanks to all our participants for their academic and social contributions.

Mirdec-4th Madrid 2017 Conference Proceedings, Full Paper Series

Masters International Danismanlik Arastirma Yayincilik
Masters International Consultancy Research and Publishing

ISBN: 978-605-82290-0-6

MIRDEC Publishing

Editors:

Kemal Cebeci

Adam Pawlicz

Slagjana Stojanovska

Copyright © 2017 Masters International Danismanlik Arastirma Yayincilik, editors and the authors. All rights reserved. No part of the material protected by this copyright may be reproduced or utilized in any form or by any means, without the prior written permission of the copyright owners, unless the use is a fair dealing for the purpose of private study, research or review. The authors and editors reserve the right that their material can be used for purely educational, scientific and research purposes.

Publisher: Masters International Danismanlik Arastirma Yayincilik

Masters International Consultancy Research and Publishing

ISBN: 978-605-82290-0-6

MIRDEC Publishing

Cinarlicesme sk. No: 21/13 PK: 34303 Kucukcekmece Istanbul Turkey

Publisher certificate no: 35822

Publication date: 30 August 2017

www.mirdec.com

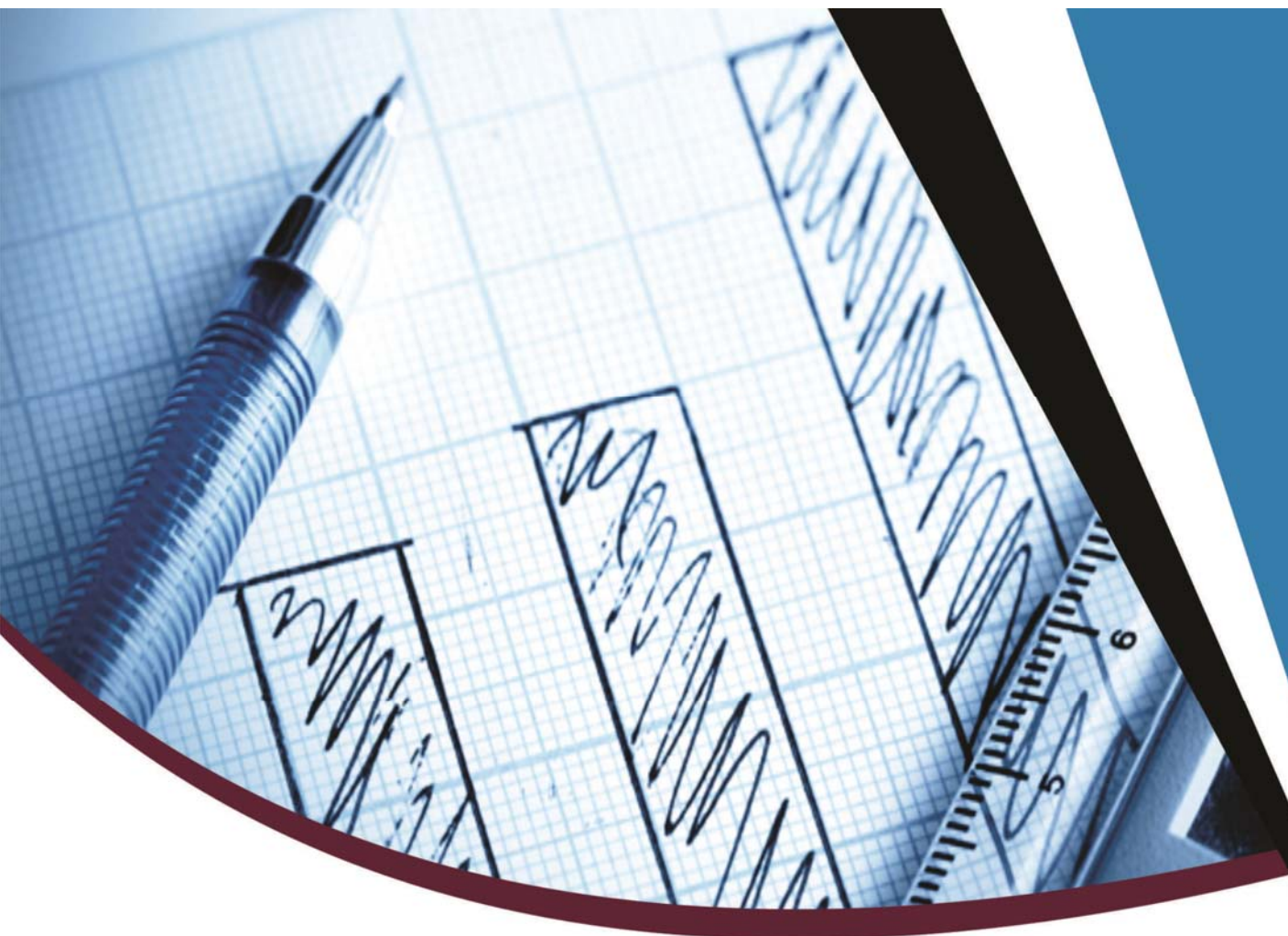
info@mirdec.com

Published at: Form Baskı Teknolojileri Reklam ve Paz. Tic. Ltd. Şti, Şerifali Mah. Şehit Sokak,
No:16/1 Umraniye / Istanbul, Turkey, Tel: (+90)(216) 337 37 96, Matbaa Sertifika
No/Certificate no: 31613, www.formbaski.com.tr

**MASTERS INTERNATIONAL
Research & Development Center
MIRDEC MADRID 2017**

Masters International
Research & Development Center
Madrid 2017





Commerce - II

- Megha Somani
- Suvaiba S. Pirani
- Jyoti M. Bhatia
- Shital Mehta

Himalaya Publishing House

ISO 9001:2008 CERTIFIED

COMMERCE - II

(As per the Revised Syllabus 2016-17 of Mumbai University for
B.Com., Semester - II)

Dr. Megha Somani

*M.Com., MMM, Ph.D., UGC-SET
Assistant Professor, Dept. of Commerce,
M.M.K. College of Commerce & Economics,
TPS III, 32nd Road, Bandra (West).*

Ms. Suvaiba Sajjad Pirani

*M.Com., NET
Assistant Professor, Dept. of Commerce,
St. Xavier's College,
5, Mahapalika Marg, Metro, Mumbai - 400001.*

Ms. Jyoti M. Bhatia

*M.Com., ACS, ACMA, LL.B., NET
Assistant Professor, Dept. of Accountancy,
St. Andrew's College of Arts, Science & Commerce,
Bandra (West).*

Dr. Shital Mehta

*M.Com., NET (JRF), Ph.D.
Assistant Professor, Dept. of Commerce,
Smt. Kamala Mehta College of Commerce,
Seven Bungalows, Versova, Andheri (West).*



Himalaya Publishing House

ISO 9001:2008 CERTIFIED

© **Authors**

No part of this publication may be reproduced, stored in a retrieval system, or transmitted in any form or by any means, electronic, mechanical, photocopying, recording and/or otherwise without the prior written permission of the publisher.

First Edition : 2017

Published by	: Mrs. Meena Pandey for Himalaya Publishing House Pvt. Ltd. , “Ramdoot”, Dr. Bhalerao Marg, Girgaon, Mumbai - 400 004. Phone: 022-23860170, 23863863; Fax: 022-23877178 E-mail: himpub@vsnl.com; Website: www.himpub.com
Branch Offices	:
New Delhi	: “Pooja Apartments”, 4-B, Murari Lal Street, Ansari Road, Darya Ganj, New Delhi - 110 002. Phone: 011-23270392, 23278631; Fax: 011-23256286
Nagpur	: Kundanlal Chandak Industrial Estate, Ghat Road, Nagpur - 440 018. Phone: 0712-2738731, 3296733; Telefax: 0712-2721216
Bengaluru	: Plot No. 91-33, 2nd Main Road Seshadripuram, Behind Nataraja Theatre, Bengaluru - 560020. Phone: 08041138821; Mobile: 9379847017, 9379847005
Hyderabad	: No. 3-4-184, Lingampally, Besides Raghavendra Swamy Matham, Kachiguda, Hyderabad - 500 027. Phone: 040-27560041, 27550139
Chennai	: New No. 48/2, Old No. 28/2, Ground Floor, Sarangapani Street, T. Nagar, Chennai - 600 012. Mobile: 9380460419
Pune	: First Floor, “Laksha” Apartment, No. 527, Mehunpura, Shaniwarpeth (Near Prabhat Theatre), Pune - 411 030. Phone: 020-24496323, 24496333; Mobile: 09370579333
Lucknow	: House No. 731, Shekhupura Colony, Near B.D. Convent School, Aliganj, Lucknow - 226 022. Phone: 0522-4012353; Mobile: 09307501549
Ahmedabad	: 114, “SHAIL”, 1st Floor, Opp. Madhu Sudan House, C.G. Road, Navrang Pura, Ahmedabad - 380 009. Phone: 079-26560126; Mobile: 09377088847
Ernakulam	: 39/176 (New No. 60/251), 1st Floor, Karikkamuri Road, Ernakulam, Kochi - 682011. Phone: 0484-2378012, 2378016; Mobile: 09387122121
Bhubaneswar	: 5 Station Square, Bhubaneswar - 751 001 (Odisha). Phone: 0674-2532129; Mobile: 09338746007
Kolkata	: 108/4, Beliaghata Main Road, Near ID Hospital, Opp. SBI Bank, Kolkata - 700 010. Phone: 033-32449649; Mobile: 07439040301
DTP by	: Sonali
Printed at	: Rose Fine Art, Mumbai. On behalf of HPH.

PREFACE

It gives us immense pleasure to present the first revised edition of *Commerce - II* at the F.Y.B.Com. level, Semester II, as per University of Mumbai, revised syllabus w.e.f. June 2016.

In this edition, the subject matter in the entire chapter is added, updated and modified in a compact manner. The book contains objectives and question bank after each and every module. There are three sets of model question paper at the end of the book. We hope that this book will meet all the requirements of students in their regular study and semester end examination.

Our special thanks to the publisher Mr. Pandey and his team have been very helpful and supportive.

We welcome constructive suggestions for improving the contents of the book.



Authors

SYLLABUS

Sr. No.	Models	No. of Lectures
1.	Concept of Services	12
2.	Retailing	12
3.	Recent Trends in Service Sector	10
4.	E-Commerce	11
	Total	45

Sr. No.	Models/Units
1	<p>Concept of Services Introduction: Meaning, Characteristics, Scope and Classification of Services, Importance of Service Sector in India. Marketing Mix Services: Consumer Expectation, Service Mix, Product, Place, Price, Promotion, Process of Services Delivery, Physical Evidence and People. Service Strategies: Market Research and Service Development Cycle, Managing Demand and Capacity, Opportunities and Challenges in Service Sector.</p>
2	<p>Retailing Introduction: Concept of Organised and Unorganised Retailing, Trends in Retailing, Growth of Organised Retailing in India, Survival Strategies for Unorganised Retailers. Retail Format: Store Format, Non-store Format, Store Planning – Design and Layout. Retail Scenario: Retail Scenario in India and Global Context, Prospects and Challenges in India, Mall Management, Retail Franchising, FDI in Retailing, Careers in Retailing.</p>
3	<p>Recent Trends in Service Sector ITES Sector: Concept and Scope of BPO, KPO, LPO and ERP. Banking and Insurance Sector: ATM, Debit and Credit Cards, Internet Banking, Opening of Insurance Sector for Private Players, FDI and its Impact on Banking and Insurance Sector in India. Logistics: Networking, Importance and Challenges.</p>
4	<p>E-Commerce Introduction: Meaning, Features, Functions and Scope of E-Commerce, Importance and Limitations of E-Commerce Types of E-Commerce: Basic Ideas and Major Activities of B2C, B2B and C2C. Present Status of E-Commerce in India: Transition to E-Commerce in India, E-Transition Challenges for Indian Corporates, Online Marketing Research.</p>

PAPER PATTERN

Maximum Marks: 100

Questions to be Set: 06

Duration: 03 Hours

Sr. No.	Questions	Marks
Q.1	Objective Questions (A) Sub-questions to be Asked (12) and to be Answered (any 10) (B) Sub-questions to be Asked (12) and to be Answered (any 10) (*Multiple Choice/True or False/Match the Columns/Fill in the Blanks)	20
Q.2	Full Length Practical Question OR	15
Q.2	Full Length Practical Question	15
Q.3	Full Length Practical Question OR	15
Q.3	Full Length Practical Question	15
Q.4	Full Length Practical Question OR	15
Q.4	Full Length Practical Question	15
Q.5	Full Length Practical Question OR	15
Q.5	Full Length Practical Question	15
Q.6	(A) Theory Questions (B) Theory Questions OR	10 10
Q.6	Short Notes To be Asked (06) To be Answered (04)	20

CONTENTS

1	Concept of Services	1 – 22
2	Retailing	23 – 46
3	Recent Trends in Service Sector	47 – 70
4	E-Commerce	71 – 90
5	Module Question	91– 96



C
H
A
P
T
E
R

1

Concept of Services

1.1 Introduction

Service sector represents different services used by the consumers and different sectors of the society. All the services like banking, insurance, telecommunication, hotel, construction, retail trade, transport, catering, tourism and so all collectively come under the service sector. It helps to increase the GDP of the country, which in turn helps to increase the economy of the country.

Service sector is the fast growing sector in India. It helps to generate employment, increase the nation's income and also helps to raise the standard of the living of the consumers. The service is of two types: one who provides the service and the other who uses the service. Service cannot be stored and the same service cannot be used again and again. Any person who uses the service has to actually pay for the same. Service are intangible, they cannot be seen or touched but they can be felt. Service most of the time is followed by the goods.

The service sector has flourished after the liberalisation, privatisation and globalisation in India.

Definitions of Service

1. Philip Kotler defines Service as “A service is any activity or benefit that one party can offer to another that is essentially intangible and does not result in the ownership of anything.”
2. The American Marketing Association defines Services as, “activities, benefits and satisfactions which are offered for sale or are provided in connection with the sale of goods.”

Characteristics/Features of Services

The characteristics or distinctive features of service are given below:

1. **Intangibility:** Services are intangible in nature. They cannot be seen or touched but can be felt. The services provided by the banks, insurance, hotels, trade, transport and so on are only felt by the consumer. A student paying fees for tuition is actually paying for getting knowledge, which is not seen or touched, but it is felt. So, unlike products, services are only felt, not seen or touched. It is the feeling of satisfaction or may be dissatisfaction.
2. **Inseparability:** Services cannot be separated from the service providers. It comes hand in hand with the service provider. The production and consumption of service happens at the same time. For example, a travel agent explaining to the tourist about the travel place. As the service cannot be separated, direct sale is the only channel of distribution and a maid has

to be physically present to cook the food; a teacher has to be present to explain things to the students. The rights of ownership of service are not transferred to the service users.

3. **Perishability:** The service cannot be stored. It is perishable in nature. One cannot store the service and use it another time. For instance, the railway or bus ticket will last only for that particular journey and not for the next one. Similarly, a movie ticket will be used for that particular show and not for the next show.
4. **Non-transferable Ownership:** Services are of non-transferable ownership. A person pays to get the service, but not to own the service. For instance, a person makes the payment to book a room in a hotel or books a table for dinner, but he does not own the hotel or restaurant. This shows that the ownership lies with the service provider.
5. **Inconsistency:** Services are inconsistent or heterogeneous in nature. A repeated service will never be same. It also depends upon the customer's wants and desires. Services are customised in nature. For instance, a musician performing in Navratri for nine nights may definitely be inconsistent, as it is very difficult to perform same all the time. A teacher in a school or college may explain differently the same topic every year. Services cannot be homogeneous.
6. **Cannot be Produced in Anticipation:** Service cannot be produced in anticipation or expectations of demand. On the contrary, products are made for future demand like pen, television sets, books and so on. Services cannot be produced first and then used later. For instance, a maid service cannot be taken on one day and used the same for the remaining six days of the week. It has to be produced or taken every day.
7. **Non-returnable:** Services once taken cannot be returned back to the service providers. If it were a product, it would be easily returned or exchanged, but it is not possible in case of service. A defective mobile or laptop can be exchanged or returned back, but poor service of banks, employees or maids cannot be given back to them, nor can they be exchanged. Only after using the entire service, one understands its quality.
8. **Needs Communication:** To provide service, one needs to communicate with the service users or customers. The needs have to be clearly understood by the service provider. For instance, in a restaurant, the customers order for spicy food or less spicy food, it has to be communicated properly. Even for the services provided by the lawyer, both the victim and lawyer need to interact about their views, so that the service is properly provided.
9. **Proper Recruitment:** It is very important to have the right person for the right job. Recruiting the right staff and also providing training to them helps in delivering service in the correct manner. The consumers or customers compare the service and make their ideas for the next time.
10. **Goods Follow Services:** Services at times are followed by the goods. No product can be sold without service. For instance, a television set cannot be sold without a salesman giving his service. In this case, television is a product which is tangible and the salesman's service is an intangible in nature. So, service may come alone (separately) like in case of a singer or teacher or it may be followed by a product.

Scope of Services

Services have many sub-sectors, they are connected with each other. The following are the various areas/scope of services:

1. **Transportation:** It brings place utility. It helps in moving the goods from one place to another. Transportation includes roadways, railways, waterways and airways. It helps in bridging the gap between service provider and service user. Transportation helps to distribute the goods and services. It promotes industrial, agricultural and economic development.
2. **Warehousing:** Warehousing creates time utility. It is the storehouse or godown. It helps in storing the goods on a large scale. Warehousing also protects the goods from the sun, wind and rain. It helps in providing regular supply of goods to the consumer.
3. **Banking:** Banks provide loans and advances and also help in depositing money. Banks provide financial services to the individual as well as the corporate. It provides the ATM services, debit cards, credit cards, net banking, locker facility and so on. These services are helpful in increasing the companies' capital which in turn raises the GDP of the country.
4. **Insurance:** Insurance service is taken by the individuals as well as by the corporates. It helps in reducing risk in the personal as well as in the corporate life. Insurance is taken for protection of various risks and uncertainties like health, theft, natural calamity, fire and so on.
5. **Education:** Education is also a part of the service sector. It also adds to the economy of India. There are many universities and courses, nationally and internationally. Indian universities also have tie-ups with other foreign institutions. It gives a very wide variety and exposure in service sector.
6. **Information Technology (IT) Services:** IT is the leading service in our country now. It is important for the economic growth of India. Wipro, Oracle Financial, Tata Consultancy and Infosys are a few well-known IT companies. IT generates a lot of employment in India and also adds to the economic growth of our country.
7. **Tourism:** This sector has given a tremendous income to our country. It is the faster growing sector in India. There has been a steady rise in the total number of foreign tourists arriving in India. While 33.04 lakh foreign tourists arrived in 2014, it rose to about 36.36 lakh in the first five months of 2016 – a 13.48% rise. It brings a lot of money to our country. The Government of India has allowed 100% FDI in this sector.
8. **Health:** Healthcare is one of the most important service sectors. There are many private and public hospitals in the society. Not only hospitals but there are many dispensaries, clinics and nursing homes to provide quality medical service to the patients. Health services also add to the GDP of our country.
9. **Retailing:** Retail sector is the growing sector in India. In India, unorganised retailing is very common like kirana shops, paan beedi shops, etc. India has a lot of scope in the organised sector like malls, departmental stores, multi-brand outlets, etc. The Government of India allowed 100% FDI in single brands and 51% FDI in multiple brands. It helps to promote the economic development of India.

10. **Hotel Industry:** Hotel industry is also an important service sector. As India has good holiday destinations, many tourists visit India. Internally also, many people travel from one part of the country to the other. This gives rise to the hotel industry, which also plays an important role in the economic development of India. Taj Group of Hotels, J.W. Marriott and ITC Hotels are a few reputed names in the hotel industries in India, with the best amenities and arrangements.
11. **Other Services:** There are many other services that are important in the day-to-day activities of the people. They also boost the economic and social development of India. They are:
 - (a) Media
 - (b) Communication
 - (c) Repairs and maintenance
 - (d) Recreation services
 - (e) Defence services
 - (f) Courier services.

Classification of Services

The services can be classified or segregated into various types. Following are the types of services.

1. **Degree of Tangibility:** Services are mostly intangible. Few services are followed by tangible goods or products. For instance, if a banker tries to explain the benefits of a new scheme, then he is purely selling services. But if a banker tries to explain the benefits of the new credit card and tries to sell it, then in this case, services are followed with goods.
2. **Degree of Relation with Customer:** Services are classified on the basis of customers' demands. It may be formal or informal in nature. For instance, if a teacher teaches in a class, then it is formal, but if a teacher gives tuition to weak student at home, then it is informal in nature.
3. **Degree of Customisation:** When the services are classified on the need of the customers, it is said to be customised service. For instance, if a readymade garment shop sells its products, it is less customised, but if it is stitched by a tailor as per the customer's needs, then it is giving a high degree of customisation.
4. **Degree of Skill:** Services are also divided on the basis of the employee's skill. If a service is divided on the basis of expertise, it may be professional or non-professional. Professional services may be like those of doctors, teachers and bankers. Non-professional services may be like the services provided by maids or domestic workers.
5. **Degree of Labour-intensiveness:** Services can also be divided on the basis of labour-intensiveness. Few services require a lot of labourers, like people working in a handicrafts or repairs unit. But few services are low labour-intensive, like ATMs or automatic packing of goods and other mechanised services.
6. **Degree of Discretion:** Services are biased in nature and differs from customer to customer. There may be discretion done by the service provider when he gives his service to the

customers. A doctor may be very polite and kind with his patients, but may be very harsh and strict with his staff at the dispensary.

7. **Degree of Business Goals:** Services may be also depend upon the business goals or objectives of a business objective is to earn profit, then the services provided may be very costly and if the business objective is to increase market share, the services may be slightly cheaper.
8. **Depends on Place and Time:** Services also depend on the place and time of service delivery. The service may be given at the place and time of service provider, like a beauty saloon, or it may be given at the place and time of the services user, *e.g.*, a beautician visiting home as per the customer needs.
9. **Degree of Government Rules:** Some services are highly governed by the government, like railways, defence, and so on. While some services have very little or no government control, like private cars or private schools and colleges.
10. **Facilities or Equipments Used:** Services may also be classified on the basis of facilities or equipments provided. For instance, a fitness centre may require more facilities and equipments than a house painter.

Importance of the Service Sector in India

Service sector is the fastest growing sector in India. It has contributed to the growing employment, increase in GDP and also raised the standard of living of the people in India. It also helps to generate foreign exchange in the country. The following are the details of growing importance of the service sector in India.

1. **Share of Services in GDP:** The contribution of service sector to GDP has tremendously grown. After 1991, the concept of liberalisation, privatisation and globalisation increased. It gave rise to the service sector. In 1950-51, GDP was 25%, it increased to 42.5% in 1990-91. After that in 2015-16, it was 64% and it will reach approximately to 90% by 2020.
2. **Revenue to the Government:** The sector provides revenue to the government in the form of service tax, corporate tax and individual tax. Services as a percentage of Gross Domestic Product (GDP) has increased from 50% in 2000-01 to nearby 60% in 2013-14. The effective service tax rate now is 12.36%
3. **Generates Employment:** It plays a very important role in generating employment. India has the second fastest growing service sector after China. The share of service sector is more in the urban areas, as compared to the rural areas in India. The service sector was contributing about 28% of the total employment in India in 2012.
4. **Service Sector Supports Other Sectors:** The other sectors like primary and secondary sector get the boost from service sector. The different services like banking, transportation, communication, warehousing, insurance services and so on are all very important for the primary as well as secondary sectors.
5. **It Helps in Social Development:** Service sector helps in the development of the society. It strengthens the social development. The services like education, health, insurance, media, etc. are very important for the social development of India.

6. **It Helps in Regional Development:** Service sector is the backbone of regional development. It helps in strengthening the region's infrastructure like transport, Information Technology (IT), warehouse facility, etc. It makes the region developed and reduces the regional imbalance. Service sector plays a very important role in backward areas like Bihar, Uttar Pradesh, Assam, Meghalaya, etc.
7. **It Helps to Improve Efficiency:** It plays an important role to improve the efficiency of the people in an organisation. It helps to improve through good education, training and development, research and development, etc. in various fields. The improvement in the person helps to improve the quality of the goods and services and reduce wastage of resources.
8. **It Builds Reputation:** Service sector development helps to build reputation of the country. It helps to create a goodwill in the global market for India due to the improvement in goods and efficiency in the service provided.
9. **Various Sector-wise Growth in Service Sector:** Service sector is the fastest growing industry in India. It includes various sectors like banking, insurance, telecommunication, IT, transportation, warehousing, hotels, travel and tourism, finance, real estate and so on. There is a rapid growth in the services. The fast growing sectors in the recent years are IT, travel and tourism, finance, banks and insurance sector.
10. **Increase in standard of living:** The service sector has a tremendous growth in the GDP of India. It has contributed to GDP about 64% in 2015-16 and employment of about 20% of the total workplace in 2012. This has led to improved standard of living for the citizens of our country, as the income of the employees has increased. The purchasing power of the people has also increased the country.

1.2 Marketing Mix Services

Customer Expectations

Customer expectations are beliefs about service delivery that function as standards or reference point against which performance is judged. Customers compare their perceptions of service delivery with these reference points when evaluating service quality and therefore knowing what customers expects is critical in gaining competitive advantage.

Levels of Customer Expectations

Customers hold different types of expectations about service, the highest type of these are desired service and adequate service.

1. **Desired Service:** This is the highest level of customer expectation. **Desired service is the level of service the customer hopes to receive.** It is a combination of what customers believe "can be" and "should be". It signals the level of customer hopes and wishes and belief that they may be fulfilled. Thus, failure to meet these expectations may result to customers cutting down on purchase.

For instance, in case of banking services, one expects prompt service, better complaint handling, high interest on savings, online availability of services, etc.

2. **Adequate Service:** Customers generally accept that the service would not always be performed according to their expectations and this is known as adequate service. **Adequate service is the level of service that customers will accept.** Though customers' hopes and wishes may still be high, they have a certain level of understanding in cases where receiving desired service does not seem possible at all.

For example, customers are used to the self-service approach used in supermarket and therefore have certain levels of understanding or tolerance towards food retailers' service delivery.

The Zone of Tolerance

Services are heterogeneous in that performance may vary across providers, across employees from the same provider, and even with the same service employee. The extent to which customers recognise and are willing to accept this variation is called the zone of tolerance. The zone of tolerance is defined as the degree to which customers recognise and are willing to accept service performance variations. Customers assess service performance on the basis of two boundaries: what they desire and what they consider acceptable.

- If service drops below adequate service level, customers get frustrated and this may cause dissatisfaction with the service provided by the company.
- If service is above the zone of tolerance, where service performed by the business exceeds the desired level, customers will have favourable responses to the business.

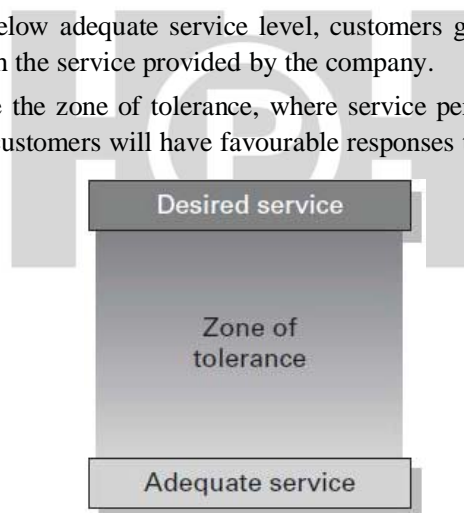


Fig. 1.1: The zone of tolerance

It is to be noted that:

- Different customers possess different zones of tolerance
- Zones of tolerance vary for service dimensions/attributes/factors

Factors Influencing Customer Expectations of Service

Because expectations play such a critical role in customer evaluation of services, marketers need and want to understand the factors that shape them.

Factors Influencing Desired Service Expectation: One of the largest influences on desired service level are personal needs and personal service philosophy.

1. **Personal Needs:** Personal needs are, those states essential to the physical or psychological well-being of the customer. For instance, a cinema-goer who regularly goes to see films straight from work, and is therefore thirsty and hungry, hopes and desires that the food and drink counters at the cinema will have short queues and attentive staff, whereas a cinema-goer who regularly has dinner elsewhere has a low or zero level of desired service from the food and drink counters.
2. **Personal Service Philosophy:** It is the customer's underlying generic attitude about the meaning of service and the proper conduct of service providers. For instance, if a person have ever been employed as a member of waiting staff in a restaurant, he is likely to have standards for restaurant service that were shaped by his training and experience in that role. He might, for example, believe that waiters should not keep customers waiting longer than 15 minutes to take their orders.
3. **Other Miscellaneous Factors:**
 - Generally, better the **image of the service organisation**, higher is the customer service expectation.
 - Customers expect high service level from the high charged/**priced** services, and *vice versa*.
 - Information about service.

Factors influencing adequate service expectations:

1. **Temporary Service Intensifiers:** It consists of short-term, individual factors that make a customer more aware of the need for service. Personal emergency situations in which service is urgently needed such as an accident raise the level of adequate service expectation.
2. **Perceived Service Alternatives:** These are other providers from whom the customer can obtain service. If customers have multiple service providers to choose from, or if they can provide the service for themselves (such as lawn care or personal grooming), their levels of adequate service are higher than those of customers who believe it is not possible to get better service elsewhere.

For instance, an airline customer who lives in a provincial town with a small airport, for example, has a reduced set of options in airline travel. This customer will be more tolerant of the service performance of the carriers in the town because few alternatives exist. The customer's perception that service alternatives exist raises the level of adequate service and narrows the zone of tolerance.
3. **Customer's Self-perceived Service Role:** It is the customer perceptions of the degree to which customers exert an influence on the level of service they receive. In other words, customers' expectations are partly shaped by how well they believe they are performing their own roles in service delivery. For instance, a customer may give special instructions to the air hostess regarding specific services required, which raises his expectation level.
4. **Situational Factors:** It is defined as service performance conditions that customers view as beyond the control of the service provider. For example, during monsoon, delay in railway service. Customers who recognise that situational factors are not the fault of the service company may accept lower levels of adequate service given the context.

- 5. Predicted Service:** It is the level of service that customers believe they are likely to get. This type of service expectation can be viewed as predictions made by customers about what is likely to happen during transaction. Higher the level of predicted service, higher the expectation of the customers.

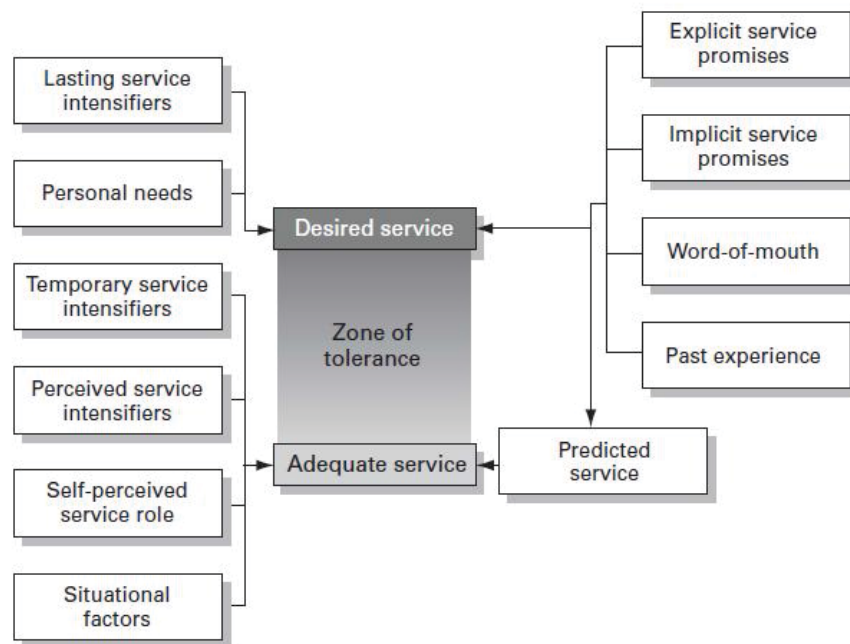


Fig. 1.2: Factors that influence desired and predicted service

Marketing Mix for Services

The service marketing mix is also known as an extended marketing mix. The product marketing mix consists of the 4Ps which are Product, Pricing, Promotions and Placement. The extended service marketing mix places 4 further Ps which include People, Process, Physical Evidence and Productivity and Quality. All of these factors are necessary for optimum service delivery.

- 1. Product:** The product in service marketing mix is intangible in nature. At the same time, service products are heterogeneous, perishable and cannot be owned. The service product, thus, has to be designed with care. In the service industry, the production and consumption of the product are simultaneous and the product is intangible. The nature of this 'product' allows for on-the-spot customisation. Firms must try to differentiate its service product from other competitors so as to get competitive advantage in the market. For instance, ICICI Bank offers account opening at the doorstep of the customers. Some of the elements of a firm's product mix are:

- core service offered
- quality of service
- service warranties and after-sale service
- product line and related services, etc.

2. **Place:** Place in case of services determine where the service product is going to be located. As mentioned, the service is produced and consumed in the same place. It cannot be owned and taken away from the location. This is why the place at which this transaction occurs is of vital importance. The location of the service provision is carefully analysed to allow ease of access and the desire to make the effort to reach it. For instance, the best place to open up a petrol pump is on the highway or in the city. A place where there is minimum traffic is a wrong location to start a petrol pump. Elements of place mix are:
 - channel selection
 - transportation
 - area coverage
 - location of store.
3. **Price:** Since a service cannot be measured by what material goes into its creation nor is the actual tangible cost of production measurable, it can be challenging to put a price tag on it. There are some tangibles of course, such as the labour costs and overheads. Firm can add mark up to the cost to determine price. But additionally, the ambiance, the experience and the brand name are also factors in the final price offering.
4. **Promotion:** Promotions have become a critical factor in the service marketing mix. Services are easy to be duplicated and hence it is generally the brand which sets a service apart from its competitors. To prevent a service from becoming interchangeable with its competitors, it becomes vital to create a desirable brand image and name in the market. Thus, firm should adopt proper promotion mix. This will not only create awareness but will also attract both new and repeat customers. Elements of promotion mix are:
 - advertising
 - sales promotion (discounts, free gifts, etc.)
 - salesmanship
 - publicity.
5. **People:** This is a vitally important element of the service marketing mix. When a service is being delivered, the person delivering it is not different from the service itself. Customers make judgments about service provision and delivery based on the people representing organisation. This is because people are one of the few elements of the service that customers can see and interact with. The staff requires appropriate interpersonal skills, aptitude, and service knowledge in order to deliver a quality service. For instance, when dining at a restaurant, if a rude waiter is encountered, the entire experience will be labeled as bad service. Elements of people mix are:
 - recruitment
 - selection
 - training and development
 - performance appraisal
 - compensation
 - process.

6. Process: This is the way in which a service is delivered to the end customer. This element of the marketing mix looks at the systems used to deliver the service. All services need to be underpinned by clearly defined and efficient processes. This will avoid confusion and promote a consistent service. In other words, processes mean that everybody knows what to do and how to do it. For instance, service process, in case of Domino's pizza, states home delivery in 30 minutes. In case of banking service, process for depositing money can be:

- taking token number
- waiting for one's turn, till the token is called
- interaction between service person and customer, whereby service persons verify details of the customers from record and counts the money
- entering transaction details in the bank record by staff
- issuing counterfoil to customer after stamping

Elements of process mix are:

- service procedure and process
- quality control
- follow-up of customers

7. Physical Evidence: Services are intangible in nature. However, to create a better customer experience, tangible elements are also delivered with the service. Often, physical evidence is used as a differentiator in service marketing. The level of comfort and attractiveness of a service location may make a lot of difference to the user experience. A calm and soothing environment, with thoughtful comfortable measures, may provide a sense of security to a new customer which will make them return. Customers will make judgments about the organisation based on the physical evidence. For example, imagine a private hospital and a government hospital. A private hospital will have plush offices and well-dressed staff. Same cannot be said for a government hospital. Thus, physical evidence acts as a differentiator.

Key elements in physical evidence mix are:

- appearance/ambience of the store
- internal store decor (furniture, sanitation, lighting, ventilation, etc.)
- appearance of the staff

Apart from these 7Ps, there is one more additional 'P' of service marketing.

8. Productivity and Quality: Productivity, which is sometimes known as performance, examines how well a company's services compete in the marketplace. This may include how consistent the service is and how well its features translate into benefits as it is being delivered.

Productivity and quality must work hand in hand. Improving productivity is key to reducing costs. Improving and maintaining quality is essential for building customer satisfaction and loyalty. Ideally, strategies should be sought to improve both productivity and quality simultaneously.

1.3 Service Strategies

Market Research

Marketing research is a systematic method of finding solution to the marketing problems in the areas of product, pricing, promotion and distribution.

According to American Marketing Association, “Marketing research is systematic gathering, recording and analysing of data about problems relating to marketing of goods and services.”

1. **Formulating the Problem:** Formulating a problem is the first step in the research process. In many ways, research starts with a problem that the management is facing. This problem needs to be understood, the cause diagnosed, and solutions developed. For instance, if the problem is ‘declining sales’, the firm should try to identify its cause from customers or sales staff. It may be due to poor quality, improper pricing, faulty promotion, poor after-sales service, etc. It provides information needed to solve the problem.
2. **Determining Data Needs:** The marketer should decide data needs and sources. Depending on the nature of the problem, primary or secondary data should be selected. Again, firm should decide on internal or external sources of data.
3. **Preparation of the Research Design:** Research design is an overall plan of the research investigation. It lays down structure within which research would be conducted. Research design involves the following elements:
 - (i) Areas of study
 - (ii) Sources of data
 - (iii) Techniques and tools for collecting and analysing data
 - (iv) Time available for research
 - (v) Cost factor relating to study.

The preparation of such design facilitates research to be efficient with minimum time, effort and money.

4. **Collection of Data:** This is the most important step in the research process. In this step, researcher collects information to solve a research problem. Data can be primary or secondary.
 - Primary data is first hand information. Primary data can be collected through experiments, surveys, observation, interview, or questionnaires.
 - Secondary data is in the form of various published sources such as journals, magazines, reports, etc.
 - The researcher should select appropriate method of data collection keeping in mind objective and scope of inquiry, finance and time available and nature of study.
5. **Organisation/Processing of Data:** The collected data is available in a raw form and needs to be processed. This processing involves classification, coding, editing and tabulation. This process is known as organising. Such organisation of data makes it ready for analysis.

6. **Data Analysis and Interpretation:** Data analysis involves application of different statistical tools such as percentages, coefficients on organised data. This enables a researcher to establish relation between the problem and the information.

Analysis refers to conclusions arrived at from research findings. It involves generalisation of research findings.

7. **The Research Report:** The research process concludes with the research report. This report will include all of your information, including an accurate description of your research process, the results, conclusions, and recommended courses of action. The report should provide all the information the decision-maker needs to understand the project.
8. **Implementation of Findings:** Researchers should submit research report to the management for implementation. Findings and recommendations of research report should be implemented to solve research problems.

New Service Development Cycle

Developing a new service includes the complete journey from generating the initial idea to bringing the service product to the market.

By setting out the steps involved, and sticking to them, your product development will become a more focused and flexible approach that can be adapted for all different types of products and services.

1. **Idea Generation:** The development of a new service will start with the generation of idea about new service. Ideas can come from many different directions. Ideas can be generated through:
 - Undertaking market research
 - Listening to suggestions from target audience – including feedback on your current products' strengths and weaknesses
 - Encouraging suggestions from employees and partners
 - Looking at competitors' successes and failures.
2. **Idea Screening:** This step is crucial to ensure that unsuitable ideas, for whatever reason, are rejected as soon as possible. Ideas need to be considered objectively, ideally by a group or committee. Specific screening criteria like return on investment, affordability and market potential of the idea needs to be considered. Proper answers to above criteria help in avoiding product failure.
3. **Concept Development and Testing:** In case of new service development, concept testing means formulating the basic product definition and then presenting the same to consumers with descriptions to get their reactions. Along with clear definition of the concept, description of the service representing its specific features and characteristics are produced to the customers and employees to determine their response to the service. For instance, a bank may intend to offer insurance policy to their customers. Bank can explain the types and benefits of policies orally to their customers to find out their reactions. These reactions will help to understand the following things:
 - Do they understand the concept?
 - Do they want or need it?

This stage gives you a chance to develop the concept further, consider their feedback, and also start thinking about what your marketing message will be.

4. **Business Analysis:** Once the concept has been tested and finalised, a firm should assess whether the new product/service will be profitable. This should include a detailed marketing strategy, highlighting the target market, product positioning and the marketing mix that will be used.

This analysis needs to include: whether there is a demand for the product, a full appraisal of the costs, competition and identification of a break-even point.

5. **Service Development:** If the new service is approved in analysis stage, it will be passed to the technical and marketing development stage. This means the firm investigates exact design and specifications. It develops value-added service attributes that brings customer satisfaction. At this stage, concept is refined after considering inputs from customers, employees and other stakeholders and service blueprint is developed. Blueprint describes service in terms of people, process and physical evidence.
6. **Test Marketing:** At this stage, before launching the service on large scale, it is launched in a limited market area to a small group of customers for a limited period at a special price. This stage aims to obtain customer feedback.
7. **Commercialisation:** If the test marketing results are favourable, final decisions needs to be made to move the product to its launch into the market. At this stage, the service goes live and introduced to the marketplace. Pricing and marketing plans need to be finalised and the sales teams and distributors are briefed, so that the service and company is ready for the launch. Proper promotion mix should be adopted to create product awareness.
8. **Post-production Evaluation:** At this stage, the information gathered during the commercialisation stage is reviewed and changes are made in the delivery process, staffing, marketing mix variables, etc. on the basis of the market response to the offerings.

Managing Demand and Capacity

Service, being intangible and perishable, cannot be produced in anticipation of demand and cannot be stocked. These features of service creates problem of managing demand and capacity in case of demand fluctuations. Demand fluctuations, in relation to supply capacity, results in three possible outcomes:

1. Excess demand (lower capacity)
2. Balance between demand and capacity
3. Excess capacity (lower demand)

There are two general approaches for accomplishing demand and capacity.

- A. To smooth the demand fluctuations themselves by **shifting demand** to match existing supply.
- B. To **adjust capacity** to match fluctuations in demand.

A. Shifting Demand to Match Capacity

By shifting demand and capacity, an organisation seeks to shift customers away from periods in which demand exceeds capacity. Perhaps by convincing them to use the service during periods of slow demand.

During periods of slow demand, the organisation seeks to attract more and/or different customers to utilise its productive capacity. Firm can use a variety of approaches listed below to increase demand to match capacity.

1. **Vary the Service Offering:** One approach is to change the nature of the service offering, depending on the season of the year, day of the week, or time of day. For instance, airlines can change the configuration of their plane seating to match the demand from different market segments. In some planes, there may be no first-class section at all. On routes with a large demand for first-class seating, a significant proportion of seats may be placed in first class. Movie theaters are sometimes rented during weekdays by business groups. It is an example of varying the service offering during a period of low demand.
2. **Communicate with Customers:** Another approach for shifting demand and capacity is to communicate with the customers. It helps them know the times of peak demand so that they can choose to use the service at alternative times and avoid crowding or delays. For example, signs in banks and post offices which let customers know their busiest hours and busiest days of the week can serve as a warning. This allows customers to shift their demand to another time if possible. In addition to signage communicating peak demand times to customers, advertising and other forms of promotion can emphasise different service benefits during peak and slow periods.
3. **Modify Timing and Location of Service Delivery:** Some firms adjust their hours and days of service delivery to match customer demand. For instance, banks can operate for extended hours, specially till evening or may operate on weekends to cater to working customers. Theaters also accommodate customer schedules by offering matinees on weekends and holidays when people are free during the day for entertainment.
4. **Differentiate on Price:** A firm can offer services at discounted prices during slow demand of the service. This strategy relies on basic economics of supply and demand, *i.e.*, demand rises when price falls. Any hotel, airline and restaurant can offer discounts during off-season. But the goal is always to ensure the highest level of capacity utilisation without sacrificing profits.

B. Adjust Capacity to Meet Demand

A second strategic approach to matching demand and capacity focuses on adjusting or flexing capacity. The idea here is to adjust, stretch and align capacity to match customer demand. During periods of peak demand, the organisation seeks to stretch or expand its capacity as much as possible. During periods of slow demand, it tries to shrink the capacity so as not to waste resources.

(i) Stretch Existing Capacity

The existing capacity of service resources can often be expanded temporarily to match demand. In such cases, no new resources are added. Rather people, facilities, and equipment are asked to work harder and longer to meet demand.

1. **Stretch Time:** It may be possible to extend the hours of service temporarily to accommodate demand. For instance, retailers are open longer hours during the Diwali shopping season. And accountants have extended appointment hours (evenings and Saturdays) before tax deadlines.
2. **Stretch Labour:** In many service organisations, employees are asked to work longer and harder during periods of peak demand. For example, service personnel in banks, tourist attractions, restaurants and telecommunication companies are asked to serve more customers per hour during busy times.
3. **Stretch Facilities:** Theatres, restaurants and classrooms can sometimes be expanded temporarily by the addition of tables, chairs, or other equipment needed by customers. Again, extra coach can be added in a train during peak season.
4. **Stretch Equipment:** Computers, telephone lines and maintenance equipment can often be stretched beyond what would be considered the maximum capacity for short periods to accommodate peak demand.

(ii) Align Capacity with Demand Fluctuations

By adjusting service resources creatively, organisations can match capacity with customer demand patterns. Specific actions might include the following:

1. **Use Part-time Employees:** In this case, the organisation's labour resource is being aligned with demand. Retailers hire part-time employees during the holiday rush, tax accountants engage temporary help during tax season, tourist resorts bring in extra workers during peak season and so on.
2. **Outsourcing:** Firms that find they have a temporary peak in demand for a service that they cannot perform themselves may choose to outsource the entire service. For example, firm can outsource some of its services like after-sales service (repairs and maintenance), especially when it is difficult to hire and train new staff for the same.
3. **Rent or Share Facilities or Equipment:** For some organisations, it is best to rent additional equipment or facilities during periods of peak demand. For example, express mail delivery services rent or lease trucks during the peak holiday delivery season. It would not make sense to buy trucks that would sit idle during the rest of the year.
4. **Schedule Downtime during Periods of Low Demand:** If people, equipment, and facilities are being used at maximum capacity during peak periods, then it is imperative to schedule repair, maintenance and renovations during off-peak periods. This ensures that the resources are in top condition when they are most needed. With regard to employees, this means that vacations and training are also scheduled during slow demand periods.

Opportunities in the Service Sector

Service sector is the fastest growing sector in India. It contributes to the GDP of India and provides a huge employee generation. Service sector has grown after 1991, where liberalisation, privatisation and globalisation (LPG) was introduced in the New Industrial policy, 1991.

Following are the reasons for the opportunities in the service sector.

1. **Various Sector-wise Growth in Service Sector:** Service sector is the fastest growing sector in India. After China, India ranks second in the contribution of service sector. The service sector includes IT, banking, insurance, telecommunication, transportation, warehousing, hotel, travel and tourism, finance, real estate and so on.
2. **Growing Income:** As the service sector grows, it generates a lot of employment. As the income of the people increase, their spending capacity also increases. Thus, the disposable income increases. It also increases the standard of the people in India.
3. **Globalisation:** Globalisation is migration of people from rural to urban areas. It is seen that cities have far better services and lifestyles. They have many facilities in health, education and telecommunication services. This attracts the people to shift to cities. This gives rise to the service sector of urban areas.
4. **Foreign Direct Investment (FDI):** FDI is one of the main reasons in the growth of service sector in India. It brings inflows of capital, skills, technology and also professionalism. Government of India has allowed FDI upto 100% in many of the sectors of services. For instance, single brand retailing, telecom, tourism and so on have 100% FDI inflow in India.
5. **Professionalism:** As FDI has increased, this has brought a lot of change in the working style of the service sector. After 1991, the Government has adopted the policy of liberalisation, privatisation and globalisation. This has led to change in the working system. This gave rise to professionalism in the service sector.
6. **Increase in Population:** India is the second largest country after China in terms of population. The population is growing at an alarming rate and soon India will be in the first position in the world in terms of population. As the population increases, so do services used or provided. This will give rise to services like education, health, telecommunication, tourism, travel, banking, insurance, hotels and so on.
7. **Socio-economic Changes:** Nowadays, there is a lot of influence of the western culture among Indians. The society has undergone tremendous change in their living and earning style. The influence is seen on every service sector like banking, insurance, retailing, education, health, telecom, food or restaurants and so on. This is giving service more opportunity to expand.
8. **Foreign Trade:** The rise in the import and export services in India has brought opportunities for the entrepreneurs. India's total export contributes nearly 30% of the service sector. This will definitely increase in the near future and give a boost to the service sector.

Challenges in the Service Sector

1. **Challenges of Intangibility:** As the services are intangible, it becomes very difficult for the service user to decide whether to take the service or not. It cannot be judged or examined before taking the service. Hence, the service provider should provide learning to their employees, create trust among the customers by providing quality service and they should also try and maintain their loyal customers so that positive verbal publicity is spread.

2. **Challenges of Inseparability:** Services are not separated from the service provider. Both the service provider and user have to be present while getting the service. It may give rise to the problem of mobility of the service provider. For instance, a service provided by maids or servants is inseparable. The maid may not be in the position to go anywhere and everywhere to give her service, due to time and location constraints. This problem may be solved by outsourcing people, training, use of automated machines wherever possible.
3. **Challenges of Perishability:** Services cannot be stored nor can they be mass produced by the service provider. In this case, the service provider should give offers or discounted rates at non-peak hours. The service provider may use different pricing policies and various other complimentary services at non-peak hours.
4. **Challenges of Inconsistency:** Services are not consistent. They definitely differ and it is an uncontrollable feature. It may also affect the quality of the service. In order to solve the inconsistency problem, the service provider should give training to their employees, motivate the employees by giving them monetary or non-monetary benefits. The customers may also be allowed to give their suggestion and comments.
5. **Challenges in Trained Manpower:** The service sector requires skilled and qualified employees. The employees have to be uniformly trained in their field like banking, telecom, tourism, medical, insurance and so on. The training facilities are inadequate and less. The tools or techniques used for training are also insufficient. Thus, there are no sufficiently trained manpower.
6. **Challenges in Customer Retention:** In this competitive world, customers have a lot of variety in products and services. They tend to shift or switch to a new service very quickly. It becomes very difficult for the service provider to retain the customers. To retain the customer, the service provider must satisfy the customer needs, also take the customers' feedback and consider their suggestions. Innovative services should be introduced for retaining consumers.
7. **Gap between Education and Application in Career:** It is seen that the education system in India does not sync with the application in service sector careers. Theoretical knowledge and the curriculum does not help in the practical application. The youngsters, who have finished their education, find it difficult to apply the same knowledge practically in different service sectors, especially banking, insurance, health, telecommunication and so on.
8. **Employee Turnover:** It is very difficult to retain the employees in the same job, as there are growing opportunities in the service sector. The areas like IT, medical, banking, media, etc. require frequent change for the employees' experience and advancement in career. This results in lot of employee turnover. To retain employees, they should be given retention bonus, and motivated with monetary or non-monetary benefits. Exit interviews should be conducted to understand the problems and to get a solution to retain the employees.

Exercise

Match the Following

I.

A	B
1. Service product	(a) Channel of distribution
2. Price	(b) Physical presence
3. Place	(c) Core benefit
4. Physical evidence	(d) Exchange value
5. Process of delivery	(e) Delivery system/sequence

Ans: 1. (c), 2. (d), 3. (a), 4. (b), 5. (e).

II.

A	B
1. Concept testing	(a) Fastest growing service in India
2. Medical tourism	(b) Leads to customer dissatisfaction
3. Poor service quality	(c) Product acceptability
4. Services	(d) Rejection of ideas
5. Idea screening	(e) Perishable/intangible

Ans: 1. (c), 2. (a), 3. (b), 4. (e), 5. (d).

State Whether the Following are True/False

1. Service provider needs to conduct marketing research.
2. Idea generation is the starting point in new service development.
3. Services are durable in nature.
4. Services can be produced in anticipation of demand.
5. Services are produced and consumed at the same time.
6. Service sector plays an important role in the economic development of the nation.
7. Marketing mix variables are same for goods and services.
8. Services are tangible in nature.
9. When service performance is above the zone of tolerance, customers are dissatisfied.
10. Service sector faces challenge of customer retention.

Ans: True: 1, 2, 5, 6, 10.

False: 3, 4, 7, 8, 9.

Fill in the Blanks with Suitable Word/Words

1. Purchasing of service does not result in _____.
(leadership, liability, ownership)

2. Services are _____.
(tangible, intangible, durable)
3. _____ is an important feature of service.
(Perishability, Tangibility, Durability)
4. Demand for services exceeds the maximum capacity during _____ period.
(peak, non-peak, off-season)
5. Currently, service sector contributes about _____ of GDP.
(30%, 50%, 60%)
6. Pre-purchase evaluation of service is _____.
(simple, possible, not possible)
7. Extent to which customer are willing to accept the variation in service performance is called _____.
(desired service, zone of tolerance, adequate service)
8. _____ is the highest level of service, which customer hopes to receive.
(Desired service, Maximum service, Adequate service)
9. If service performance is below adequate level, customer is likely to be _____.
(satisfied, dissatisfied, delighted)
10. _____ is a systematic method of finding solutions to the marketing problem.
(Marketing research, Decision-making, Planning)
11. _____ is overall plan for research investigation.
(Research design, Sample design, Data collection)
12. _____ can be offered to customers to shift demand during non-peak periods.
(incentives, comfort, disincentives)

Ans: 1. ownership, 2. intangible, 3. perishability, 4. peak, 5. 60%, 6. not possible, 7. zone of tolerance, 8. Desired service, 9. dissatisfied, 10. Marketing research, 11. Research design, 12. comfort.

Define

1. Services
2. Consumer Expectations
3. Marketing Mix
4. Marketing Research
5. Market Testing

Write Shorts Notes on the Following

1. Service Sector
2. Classification of Service
3. Desired Service Expectations

4. Product
5. Price
6. Place
7. Promotion
8. Productivity
9. Physical Evidence
10. Process
11. People
12. Marketing Research
13. Managing Demand and Capacity
14. Challenges in Service Sector

Answer the Following

1. Define service. Explain its characteristics.
2. Discuss the scope of service.
3. Bring out the classification of services.
4. Explain the importance of service sector in Indian country.
5. Discuss different levels of consumer expectations.
6. What are the elements of service mix?
7. What are the steps in marketing research?
8. Briefly discuss new service development cycle.
9. Bring out various strategies of managing demand and capacity in service.
10. What are the opportunities in service sector?
11. What are the challenges in service sector?

Case Study 1 – Palace on Wheels

E-Tailors

Palace on Wheels takes tourists on a journey through Rajasthan, the land of sand dunes, and regal palaces and Agra, the land of the Taj Mahal. Rated as one among the ten best luxurious rail journeys in the world, it aims to provide the ultimate royal experience. The Palace on Wheels was started in 1982 as a heritage holiday train by joining the coaches of the original royal saloons owned by princely states of Gujarat, Rajputana, the Nizam of Hyderabad and the Viceroy of British India. Later, these coaches were replaced by modern air-conditioned coaches, but the royal ambience was maintained. Presently, the train has 14 saloons. Each coach has four twin-bedded chambers decorated in colourful Rajasthani art and the panels and ceilings are covered with miniature traditional motifs that reflect courtly life. The saloons are equipped with world-class facilities such as channel music, intercom, attached toilets, running hot and cold water, shower stalls and wall-to-wall carpeting. Each saloon has personal attendants called Khidmatgars at the beck and call of the guests. The train also has two

restaurants named “The Maharajah” and the “The Maharani” with a princely ambience where guests have a choice of Continental, Chinese, Indian and Rajasthani cuisines, prepared by the chefs in the adjacent kitchens. In addition, the train also has well-stocked bar and a library. The train travels mostly in the night and stops during the day to allow the guests to visit the palaces and the forts. The Palace on Wheel experience has become one of the most sought-after luxuries for international tourists, and has long passenger lists that require guests to book months in advance in order to get their share of the royal experience.

Questions:

1. What characteristics of services of Palace on Wheel make it attractive?
2. How services are modified over a period of time?

Case Study 2 – Service Differentiation

Indian Post, with more than 0.15 million post offices, majority of which are in rural areas, provides a useful distribution channel of non-postal services. Although the advent of courier services has reduced the business of postal services in urban areas, it continues to be patronised by rural population of savings deposits, postal insurance, village telephone, etc., in addition to the regular postal services. Sensing an opportunity to increase the revenue and compensate for the reduction in person-to-person mailing, India Post has decided to increase the number of life insurance schemes in rural area along with the introduction of new schemes. The Indian postal network, the largest in the world, plans to tap rural families with its array of products. India Post is leveraging the personal contact and first-hand understanding of the local people by the postal staff to market the new service products in the rural areas. India Post is targeting about 50% of the revenue being generated from non-postal services for the effective implementation of the marketing efforts. India Post has decentralised the marketing and sales function to the divisional level.

Questions:

1. What are the reasons for decline in business of postal services?
2. What differentiation in services are offered to customers to maintain competitiveness?





www.himpub.com



ISBN: 978-93-5262-418-8

PCF 421

₹ 75/-

साहित्य सौरभ



हिन्दुस्तानी प्रचार सभा प्रकाशन

हिन्दुस्तानी प्रचार सभा द्वारा संचालित
'सरल हिन्दी' परीक्षा
की पाठ्यपुस्तक

साहित्य-सौरभ

प्रधान संपादक
डॉ. रीता कुमार

सह संपादक
डॉ. सुमन जैन
डॉ. संतोष कौल 'काक'
डॉ. भगवतीप्रसाद उपाध्याय



हिन्दुस्तानी प्रचार सभा प्रकाशन

ISBN NO.: 978-93-82287-12-4

प्रकाशन

: श्री फ़िरोज़ पैच
ट्रस्टी व मानद सचिव
हिन्दुस्तानी प्रचार सभा
महात्मा गाँधी मेमोरियल बिल्डिंग
7, नेताजी सुभाष रोड, मुंबई - 400 002

पुस्तक सलाहकार समिति

: श्री सतीश शाह
ट्रस्टी व अध्यक्ष, हिन्दुस्तानी प्रचार सभा
: श्री फ़्रांसिस मैथ्यू
ट्रस्टी, हिन्दुस्तानी प्रचार सभा
: श्री अरविन्द डेगवेकर
ट्रस्टी व कोषाध्यक्ष, हिन्दुस्तानी प्रचार सभा

प्रथम संस्करण

: सन् 2017

मूल्य

: रुपये 65/-

आवरण

: राकेश कुमार त्रिपाठी

सर्वाधिकार

: हिन्दुस्तानी प्रचार सभा

मुद्रक

: इंडिगो प्रिंटर्स, बीआरएम - 10
गुप्ता मिल्स, देवीदयाल कंपाऊंड
ब्रिटानिया कंपनी के पास, रे रोड (पश्चिम)
मुंबई - 400 010



महात्मा गाँधी मेमोरियल बिल्डिंग, हिन्दुस्तानी प्रचार सभा



हिन्दुस्तानी प्रचार सभा प्रकाशन