

Innovation Ecosystem

Entrepreneurship Cell

Overview:

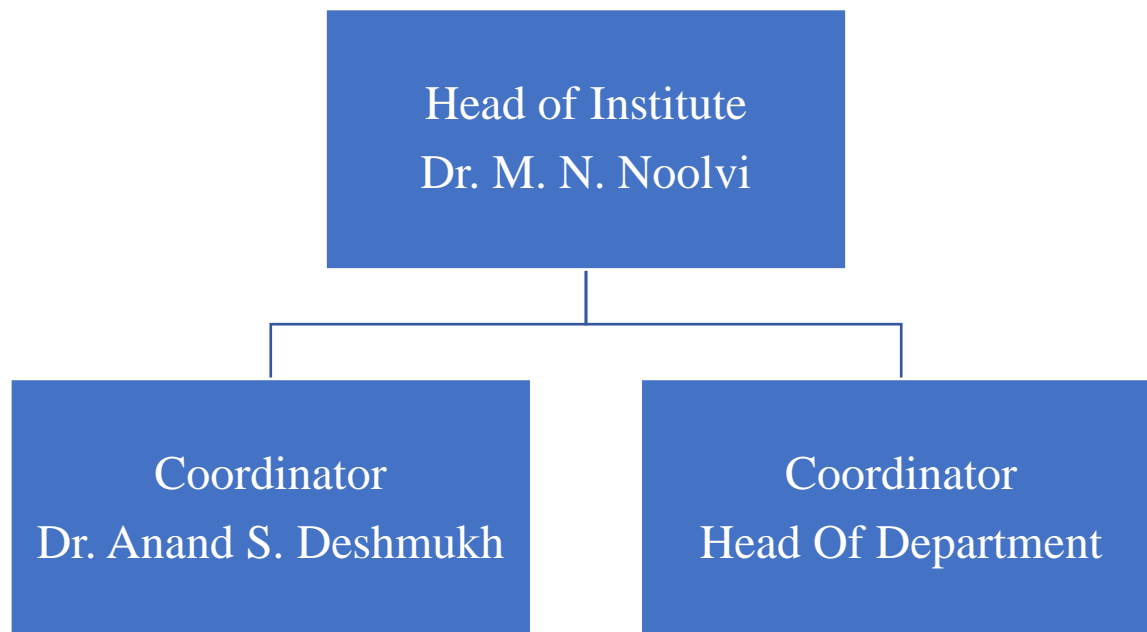
The Entrepreneurship Cell popularly known as EDC, at SDPC, Kim Campus is aimed at fostering entrepreneurial skills among the students on campus.

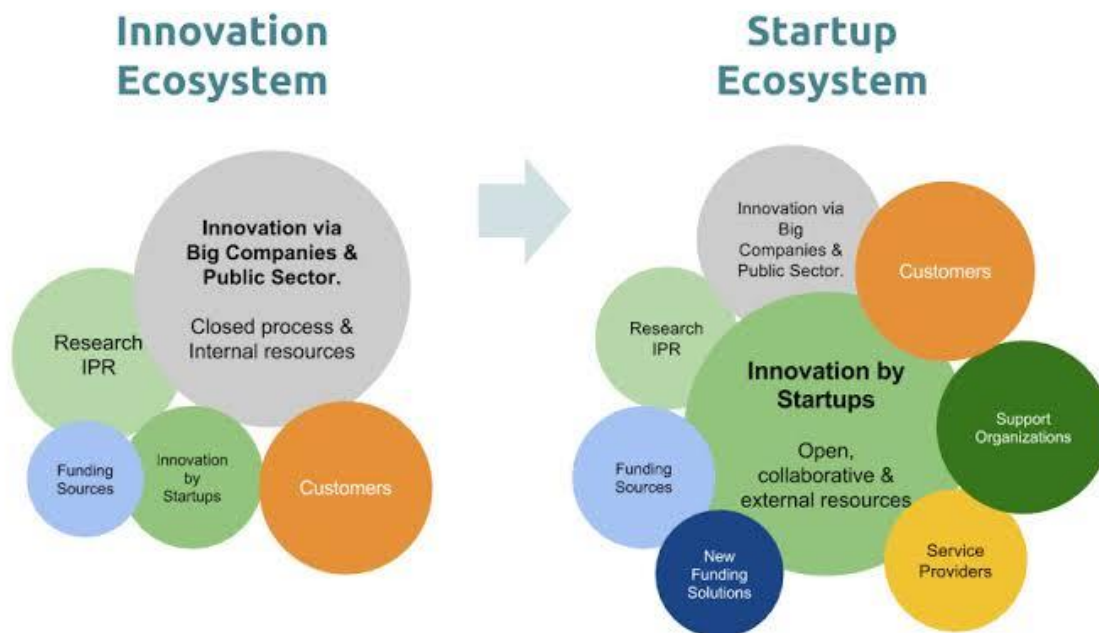
In the process of identifying, nurturing, training and establishing start-ups, EDC is involved in a wide range of activities.

Vision:

EDC is a consortium dedicated to find the entrepreneur in the students of the SDPC, Kim. EDC connects students with solid entrepreneurial skills, in the campus, to form a close-knit network, working together to spread the word of entrepreneurship. In EDC, the belief is simple; Entrepreneurship is not just a “you” and “me” game. It’s an “us” game which takes off successfully only when people come together. EDC promotes openness of knowledge, network, contacts and opportunities. Since its inception on 1st May 2012, it has successfully conducted several competitions of challenging formats, workshops, guest lectures and it has also provided mentors to several “now-successful” start-ups on campus.

Organizational Structure:





Institution has created an eco-system for innovations including incubation centre (Shree Dhanvantary Pharmaceutical Analysis & Research Centre (SDPARC)) and other initiatives for creation and transfer of knowledge. The college has Research Committee to monitor and address issues of research activities under the chairmanship of M. N. Noolvi, Professor and Principal, senior professor Dr. Pallavi KJ & Dr. Suresh Jain and associate professors Dr. Sushil Raut, Dr. Uttam A. More. The incubation facility at the institute is beneficial to both nurture new companies in the region and support the training (skill development) and gainful employment of college students. Institute constituted research advisory committee in 2012 and revised committee members in 2014. Research Advisory Committee provides advisory regarding pertaining of new research project and grants. It also advises the faculty to undertake new and innovative research project and assist them in writing for new research proposal and grants. Various projects and funds were received through the Research Advisory Committee.

Functions of the committee:

The main function of Research Committee is to encourage faculty members to write research proposals and submit to various funding agencies or industries. It motivates faculty and students to publish research work in high impact peer-reviewed journals. Also motivates to present their research work in national and international conferences. To sign MOU for collaborative research with universities and industries it also helps to provide seed fund to transform an idea into a product.



Outcomes:

- Faculty members received research grant from GUJCOST (24.45 lakhs) and BRNS (67.37 lakhs)
- One of the faculty (Dr. Anand Deshmukh) won the second prize (3.00 lakhs) for his project on safe and economic biodegradable hydrogels for water conservation in agriculture using green technology from vibrant Gujarat Startup Summit 2016.
- Faculty members received industrial projects from Good Health Pvt. Ltd. GreenKem Organics Pvt Ltd., CPA Pharma LLP, Nanotherapeutics etc.
- With the help of incubation facility “diet-o-fit” product developed for CPA Pharma LLP and many more product/processes developed.
- Faculty members filed 12 Indian patents for their research work.
- Faculty members Published their research work in high impact Peer-reviewed journals.
- Faculty members are awarded PhD (04) enrolled for PhD (08).
- Signed MOU with several international universities (Texas State US, Lamar University-Texas, Jinan University-China, University of KwaZulu-Natal- South Africa, University of Ulsan-South Korea), and pharmaceutical industries.
- Signed MoU with Jinan University China for the research project with amount of RMB 9000000 (9 Million)
- SDPARC, Linkages and MoUs All Students of final year B. Pharm and M. Pharm undergo industrial training for one month successfully.
- Several research awards have been received by faculty from various professional and research organization.
- Alumni of the institute have started their marketing firms, Pharmacies/Drug stores, Wholesale and other allied pharmacy related business.
- With these R & D activities institute received SIRO-DSIR approval.

Awards Received by faculty members:

- Received 2 Awards by **Dr. M. N. Noolvi** for his excellent teaching and scientific skill Dr. APJ Abdul Kalam Awards for **Teaching Excellence** & Dr. APJ Abdul Kalam Awards for **Scientific Excellence** on 21 October, 2017 from MARINA LABS.
- Received 1 Award by **Dr. Uttam A. More** for his research work Dr. APJ Abdul Kalam Awards for **Young Scientists** on 21st October, 2017 from MARINA LABS.



Dr. M. N. Noolvi



Dr. Uttam A. More



- Received 3 Lakhs to **Dr. Anand S. Deshmukh** for his idea on “The wet soil An economic approach for water conservation” from Vibrant Gujarat Start Up Summit 2016, with platform of ICREATE, (International Centre for Entrepreneurship and Technology), 21-22 October, 2016.



Dr. Anand S. Deshmukh

- Received 1 Award by Dr. M. N. Noolvi for his teaching skills & research work “Best Academician of the Year” on 11th August, 2018 from ICCI.



- India-South Africa Joint proposals, DST, between our college and Kwazulu-Natal university, South Africa, Proposal Title: “Investigations of Peptide Based CNT Guided Delivery Device for Cancer Therapy”



- Received Best Researcher and Academician Award to **Dr. Suresh Kumar Jain** from VDGGOOD.



- Prof. M. N. Noolvi invited as guest speaker at china, Asia Pacific Pharmaceutical science summit forum.



**Invited Speaker
Dr. M. N. Noolvi at China
9th-10th January 2019**



 暨南大學藥學院
Jinan University College of Pharmacy
601 Huangpu Avenue West, Guangzhou, Guangdong, CN

Issue Date: 07/10/2019

To: Embassy of People's Republic of China, Delhi, India

LETTER OF INVITATION

Subject: Application for F Visa with Double entry

Dear Sir,
We have invited Prof. Malleshappa Nosbi to people's Republic of China as a Visiting Professor, to visit the College of Pharmacy, Jinan University, China from 2019/11/24 to 2019/11/30

Applicant's Details are as follows:
Name: Nosbi Malleshappa
Date of Birth: 01/06/1973
Gender: Male
Passport No: T7098316
Issue Date: 06/09/2019
Date of Expiry: 05/09/2029
Place of Issue: Surat
Nationality: Indian

Furthermore, we confirm that Prof. Malleshappa Nosbi expenses incurred in China will be borne by College of Pharmacy, Jinan University.

We will provide you with

- A round-trip economic class international air ticket issued by our university-designated travel agency.
- Any other related local transportation cost.
- Hotel accommodation during your visit.

Thank you in advance for your assistance.

Yours Sincerely,
Yu Cai,
Professor,
College of Pharmacy,
Jinan University.

Invitation Letter (Jinan University China)

R & D Facility

R & D FACILITY



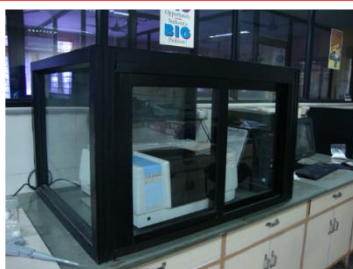
HPLC



GC



AAS



FT-IR



Dissolution Test

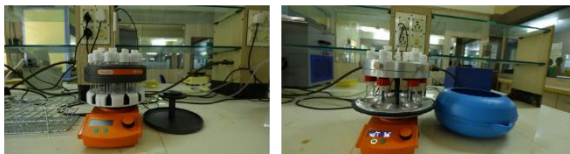


Distillation Unit

R & D FACILITIES



Heidolph Rota Evaporator



Parallel Synthesizer



Scale-up Reactor

Research Lab-II



R & D FACILITIES



DDPRL



Training Room



Research Lab-I



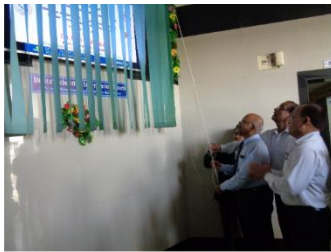
GC



HPCL



Cold Centrifuge



Digital Scientific Information Board



GUJCOST Sponsored Lab



- Interlink with international & national companies
Tie-up with these companies our R & D center commercialized various formulations & API synthesis process/methods, tech transfer details.

Linkage with Industry








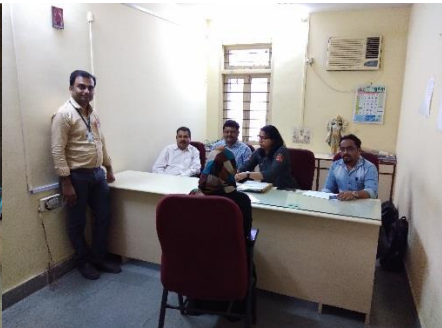
Innovation Made:

1.	<p>Company Name: CPA Pharma LLP – Surat India Project leader: Dr. M. N.Noolvi Assistant: Dr. Uttam A. More, Mr. Stephen Avvaru</p> <p>R & D work: Process Chemistry, API Synthesis “Pantoprazole” the synthetic route for pantoprazole was changed using synthon approach and same process was commercialized to CPA Pharma LLP – Surat India</p>	 <p>CPA PHARMA LLP Live your life happy</p>
2.	<p>Company Name: CPA Pharma LLP – Surat India Project leader: Dr. M. N. Noolvi & Dr. Anand Deshmukh Assistant: Ms. Payal Jain</p> <p>R & D work: Product Development, “DIET-O-FIT” DIET-O-FIT is a perfect blend of balanced nutrition, packed with high quality soy protein isolate, carbohydrates, fibers, vitamins, antioxidants, low calorie sugars, poly-unsaturated fatty acids and selected plant extract serving low in calorie essential nutrients. It Provides an ideal way to maintain a balanced calorie intake by avoiding unnecessary Fat consumption. The added plant extract additionally aid in managing basal metabolic rate (BMR) http://www.cpapharma.com/product/diet-o-fit/</p> 	 <p>CPA PHARMA LLP Live your life happy</p>
3.	<p>Company Name: Good Health Pvt Ltd, Sector 15A, Noida Project leader: Dr. Basavaraj Assistant: Mr. Washim Hussain</p> <p>R & D work: Product Development, “Unique Parenteral formulation”</p>	 <p>GOOD HEALTH</p>
4.	<p>Company Name: Oniosome Healthcare Pvt. Ltd. Mohali. Project leader: Dr. M. N.Noolvi Assistant: Dr. Uttam A. More</p> <p>R & D work: Since our research lab is having in silico molecular modeling software with the help of that small project was carried out entitled “Interaction between Phospholipid and Bleomycin A2”</p>	 <p>oniosome www.oniosome.com</p>



5.	<p>Company Name: Krishna Bio-Organic Agro Project leader: Dr. Anand Deshmukh Assistant: Ms. Payal Jain R & D work: Formulation, “Bio-waste Based Novel, Safe, Green and Economic Hydrogels”</p> <p>Availability of water for agricultural is a major challenge in rural lands of many developing countries like India, South African and Asian continents. Here availability of potable water for drinking and agriculture greatly depends upon rain water with a scarce supply by groundwater sources. Recent reports of WHO and UNICEF suggests that 84% of the people globally, who don't have access to improved water, live in rural areas, where they live principally through subsistence agriculture. The measures of government in such countries can only be effective if possible water harvesting plans are laid at the grass root levels starting from villages and small towns, where agriculture is the main source of income and survival. Further, as a solution to situation, use of Acrylic hydrogels is widely acknowledged. But again these Acrylic hydrogels are well known for their toxic effect on the ecosystem. And for many developing countries with poor economies, these acrylic hydrogels are the only available choices for managing agricultural needs with poor water resources as in arid areas.</p> <p>Keeping this in mind, we propose here a feasible and practical means of developing ecofriendly, biodegradable, economic, yet effective hydrogel system using the easily available bio-wastes (such as coconut coir, sugarcane bagasse or fibrous cereal bran) in conjunction with natural polysaccharides and bio-safe cross-linkers in accordance to the concepts of green chemistry.</p> <p>These hydrogels help in conserving water, preventing toxicity to farming land, better crop yield, enhancing soil quality and prevention of leaching of nutrients. In addition, this technology of hydrogel synthesis is simple, yet easy to scale up and does not require expensive equipment/instrumental set up, which is a desirable feature in view of the economic viability of rural areas of the developing countries.</p> <p>This project helps in reducing the use of hazardous synthetic polymer-based hydrogels and their leach outs in land and water. The project is also planned to be extended for delivery of plant nutrients and natural biocid.</p>






<p>6.</p>	<p>Company Name: Greenkem Organics, Ahmedabad Project leader: Dr. M. N. Noolvi Assistant: Dr. Uttam A. More, Mr. Stephen Avvaru, Mr. Ankit Vaghasiya</p> <p>R & D work: Process Chemistry, API Synthesis “Albendazole”</p> 
<p>7.</p>	<p>Company Name: Onpharno Project leader: Dr. M. N. Noolvi & Dr. Anand Deshmukh Assistant: Mr. Stephen Avvaru, Mr. Ankit Vaghasiya R & D work: Product Development, “Dermatological product development”</p> 
<p>8.</p>	<p>Company Name: Zydus Cadila Project leader: Dr. M. N. Noolvi & Dr. Suresh Jain Assistant: Mr. Sanket Gandhi R & D work: Training, “Students Training & placement”</p> 
<p>9.</p>	<p>Company Name: Sun Pharma Project leader: Dr. M. N. Noolvi & Dr. Suresh Jain Assistant: Mr. Sanket Gandhi R & D work: Training, “Faculty Training”</p>   



<p>10.</p>	<p>Company Name: Triokaa Project leader: Dr. M. N. Noolvi & Dr. Suresh Jain Assistant: Mr. Rohan Barse R & D work: Training, “Training to faculties on research proposal writing”</p>  
<p>11.</p>	<p>Company Name: Onpharno Project leader: Dr. M. N. Noolvi Assistant: Mr. Rohan Barse R & D work: “New Cosmetic formulation development for dermal application”</p>  

Sponsoring agencies for R & D Projects

Name of Principal Investigator	Project Titles	Amount sanctioned (Lakhs)	Sponsoring Agency
 Dr. M. N. Noolvi	Design and synthesis of radiolabeled heterocycles as aromatase inhibitors: a novel approach for breast cancer diagnosis and therapy	25.89 (On going)	<i>Board of Research in Nuclear Sciences (BRNS), Trombay, Mumbai</i>
	Development of small molecule of protein tyrosine kinase inhibitors: a hope for cancer therapy	5.00 (Completed)	Gujarat Council on Science & Technology (GUJCOST)
 Dr. Anand S. Deshmukh	Electron beam irradiation induced graft polymerization of polysialic acid onto polycaprolactone for effective and biocompatible encapsulation of therapeutic agents	16.93 (Completed)	<i>Board of Research in Nuclear Sciences (BRNS), Trombay, Mumbai</i>
	Bio-waste based novel, safe, green and economic hydrogels	Sanctioned	The Federation of Indian Chambers of Commerce and Industry (FICCI), New Delhi
 Dr. Uttam A. More	Design and synthesis of radiolabeled heterocyclics as potential diagnostic imaging agents for neurodegenerative disorders	24.55 (On going)	<i>Board of Research in Nuclear Sciences (BRNS), Trombay, Mumbai</i>



	<p>Development and evaluation of biodegradable polymeric system based long acting depot formulation of (LHRH) agonist goserelin acetate.</p>	<p>5.00 (Completed)</p>	<p>Gujarat Council on Science & Technology (GUJCOST)</p>
	<p>To investigate the antidepressant and anxiolytic effects of <i>Mimosa Pudica</i> extracts and <i>Curcuma longa</i> extracts on chronic administration of some synthetic drugs in rats</p>	<p>4.20 (Completed)</p>	<p>Gujarat Council on Science & Technology (GUJCOST)</p>
	<p>Development and isolation of bioactive component from endophytic fungi of <i>Musa Paradisiaca</i> as antidiabetic agent</p>	<p>4.00 (Completed)</p>	<p>Gujarat Council on Science & Technology (GUJCOST)</p>
	<p>Design and development of polyherbal formulation for skin repigmentation therapies in vitiligo</p>	<p>2.50 (Completed)</p>	<p>Gujarat Council on Science & Technology (GUJCOST)</p>




Dr. Bhavesh V. Akbari

Mr. Samaresh Pal Roy

Dr. Hasumati A. Raj

Dr. Gautam Sonara



	<p>Synthesis of chitosan derivatives for antimicrobial, antifungal and antitumor activity</p>	<p>2.50 (Completed)</p>	<p>Gujarat Council on Science & Technology (GUJCOST)</p>
	<p>Evaluation of endophytes for their antimicrobial activity from indigenous medicinal plants in south Gujarat</p>	<p>2.25 (On going)</p>	<p>Gujarat Council on Science & Technology (GUJCOST)</p>
	<p>Development of new pro drugs for HIF-1 inhibitions: A new hope for cancer therapy.</p>	<p>1.50 (On going)</p>	<p>Gujarat Council on Science & Technology (GUJCOST)</p>

Dr. Manish Goyani

Dr. Nirav K. Gheewala

Dr. Meghna Patel



- Constituted research advisory committee in 2012 and revised in 2014 under chairmanship of Prof. Tejraj M. Aminabhavi.

Advisory Committee, SDPC, Kim, Surat

 <p>Prof. Tejraj M. Aminabhavi PhD (The University of Texas, Austin, USA) Polymer Science and Engineering Discipline Editor, Chemical Engineering Journal (Elsevier, New York) and Emeritus Professor, College of Pharmacy, Dharwad, India. Chairman</p>	 <p>Dr. Mallikarjuna N. Nadagouda PhD (Gulbarga University, Gulbarga, India) Physical Scientist, National Risk Management Laboratory US Environmental Protection Agency (US EPA). Member</p>
 <p>Prof. Walter E. Rudzinski PhD (The University of Arizona, Tucson, USA) Professor, Texas State University, USA. Member</p>	 <p>Dr. Sudipta Chakraborty PhD (Mumbai University, Mumbai, India) Scientific Officer-F (Group Leader, Radiochemical Group) Bhabha Atomic Research Centre, Mumbai, India. Member</p>
 <p>Prof. Kap Seung Yang PhD (North Carolina State University, Raleigh, North Carolina, USA) Professor, Department of Polymer & Fiber System Engineering and Director of Alan G. MacDiarmid Energy Research Institute at Chonnam National University. Member</p>	 <p>Dr. Padmakar V. Kulkarni PhD (Rensselaer Polytechnic Institute, Troy, New York, USA) Adjunct Professor in Radiology, department of UT Southwestern Medical Center, Dallas. Member</p>
 <p>Prof. Han Mo Jeong PhD (Seoul National University, Gwanak, South Korea) Professor, Department of Chemistry, University of Ulsan, Ulsan, South Korea. Member</p>	 <p>Dr. M. Surianarayanan PhD (Kanagawa Industrial Technology Research Institute, Kanaga Wa-Ken, Japan) Call for Industrial Safety and Risk Analysis, Chemical Engineering Department, CLRI (CSIR), Chennai, India Member</p>
 <p>Prof. Hyung IL Lee PhD (Carnegie Mellon University, Pittsburgh, USA) Professor, Department of Chemistry, University of Ulsan, Ulsan, South Korea. Member</p>	 <p>Prof. Kuei-Hsien Chen PhD (Harvard University, USA) Research Fellow & Deputy Director, Institute of Atomic & Molecular Sciences, Academia Sinica, National Taiwan University, Taipei, Taiwan. Member</p>

- Institute has signed **5 MOU's with international universities** and has good outer world connections.
 - MOU with Texas State University USA
 - MOU with University of Ulsan South Korea
 - MOU with University of KN South Africa
 - MOU with Lamar University, Beaumont, Texas, USA
 - MOU with Jinan University, China



MOU WITH TEXAS STATE UNIVERSITY USA



MEMORANDUM OF UNDERSTANDING

BETWEEN

Shree Dhanvantary Pharmacy College
Kim-Surat-39 4110, Gujarat, India

AND

Department of Chemistry and Biochemistry
Texas State University, San Marcos, Texas, USA, 78666

Department of Chemistry and Biochemistry, Texas State University, San Marcos TX, USA and Shree Dhanvantary Pharmacy College, Kim, Surat, Gujarat, India seek to implement opportunities for joint academic co-ordination, develop research collaborations and promote mutual understanding between the two institutions for academic teaching and research. To these ends, both the institutions agree to foster and promote academic co-operation in education and research activities.

Contact Person

Taraj Aminabhavi
Research Director
Shree Dhanvantary Pharmacy College
Kim-Surat, India

Walter Rudolph
Department of Chemistry and Biochemistry
Texas State University
San Marcos, Texas, USA

Signing Authorities

Shree Dhanvantary Pharmacy College
Kim-Surat, India

Texas State University
San Marcos, Texas, USA

Date: 2/27/2013
Dr. N.B. Jirani
President

Date: 2/27/2013
Dr. Bruce M. French
President

PRESIDENT
SHREE DHANVANTARY PHARMACY COLLEGE
Managed By
SHREE BAHUKAR EDUCATION TRUST
KIM, SURAT, GUJARAT, INDIA



MOU WITH UNIVERSITY OF ULSAN SOUTH KOREA



MEMORANDUM OF UNDERSTANDING

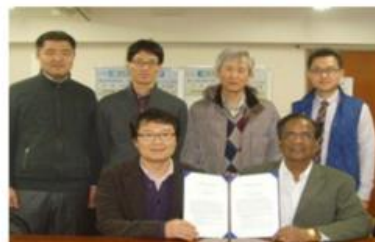
BETWEEN

Shree Dhanvantary Pharmacy College
Kim, Surat-39 4110, Gujarat, India

AND

Department of Chemistry, University of Ulsan, South Korea

Department of Chemistry, University of Ulsan, located in Ulsan, SOUTH KOREA and Shree Dhanvantary Pharmacy College, Kim, Surat, Gujarat, India seek to implement opportunities for joint scholarship to facilitate academic co-operation; to develop research collaboration and to promote mutual understanding as well as applying joint research funds between the two countries. To these ends, the institutions agree to foster and promote academic co-operation in education and research between the two institutions. This memorandum of understanding (MOU) provides a general framework for the development of such collaboration including joint research projects between the two countries and holding of international seminars. A subsequent implementation agreement will specify institutional activities, which will be agreed upon and developed by Shree Dhanvantary Pharmacy College, Kim-Surat, India and Department of Chemistry, University of Ulsan, at a future date. This will include deployment of students for Ph.D training or post-doctoral research.



Signing Authorities, Signature, Date

1) Indian: Taraj Aminabhavi, Ph.D.
Director of Research,
Shree Dhanvantary Pharmacy College
Kim-Surat, India

Dec. 18, 2013

2) Korean: Youngil Lee, Professor,
Department of Chemistry (Department Chair)
University of Ulsan
Ulsan 680-749
Korea (South)
TEL: +82 (0)52-259-2341
FAX: +82 (0)52-259-2348

Dec. 18, 2013

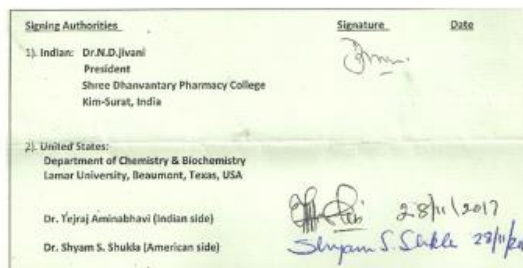




MOU WITH UNIVERSITY OF KN SOUTH AFRICA



MOU WITH LAMAR UNIVERSITY, BEAUMONT, TEXAS, USA





Funding Agency



<https://brns.res.in/index.php>



<https://gujcost.gujarat.gov.in/>



<https://www.greenkem.org/>



<https://goodhealth.com>



<https://vgstartup.com/>



<http://ficci.in/>



<https://gujcost.gujarat.gov.in/>



Institute h-index and citation from scopus

Shree Dhanvantary Pharmacy College

KIM [East], Near Railway Station, Dhanvantary College Road, Taluka : Olpad, Surat GJ, India
Affiliation ID: 60116042

Other name formats: [Shree Dhanvantary Pharmacy College](#) [Shree Dhavantary Pharmacy College](#) [Shree Dhanvanvantary Pharmacy College](#)
[Dhanvantary College Of Pharmacy](#) [Shri Dhanvantary Pharmacy College](#) [Shree Dhanvantry Pharmacy College](#) [View all](#)

[Follow this affiliation](#)

[Give feedback](#) [Set feed](#)

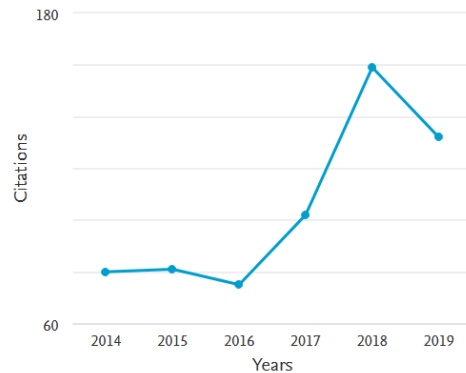
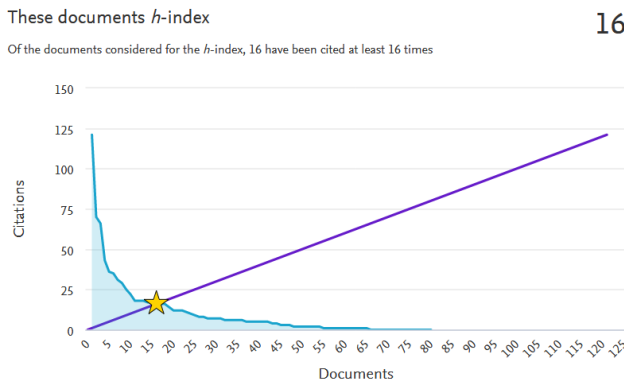
Documents, affiliation only
80

Authors
58

This is an overview of citations for the documents you've selected. Document *h*-index : 16 [View *h*-graph](#)

80 cited documents [+ Save to list](#)

Date range: 2014 to 2019 Exclude self citations of all authors Exclude citations from books [Update](#)

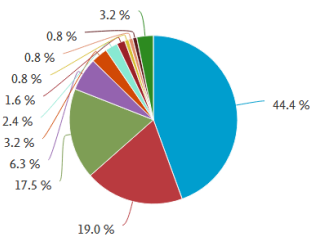


Documents by subject area Collaborating affiliations Documents by source

Sort by: [Document count \(high-low\)](#)

Subject Area	Count	Subject Area	Count
Pharmacology, Toxicology and Pharmaceu...	56	Agricultural and Biological Sciences	1
Chemistry	24	Computer Science	1
Biochemistry, Genetics and Molecular Bio...	22	Dentistry	1
Chemical Engineering	8	Economics, Econometrics and Finance	1
Medicine	4	Energy	1
Engineering	3	Environmental Science	1
Materials Science	2	Physics and Astronomy	1

Shree Dhanvantary Pharmacy College



- Pharmacology, Toxicology and Pharmaceutics
- Chemistry
- Biochemistry, Genetics and Molecular Biology
- Chemical Engineering
- Medicine
- Engineering
- Materials Science
- Agricultural and Biological Sciences
- Computer Science
- Dentistry
- Other

Publication

- [1] H.M. Patel, V. Bhardwaj, P. Sharma, M.N. Noolvi, S. Lohan, S. Bansal, A. Sharma, Quinoxaline-PABA bipartite hybrid derivatization approach: Design and search for antimicrobial agents, *Journal of Molecular Structure* 1184 (2019) 562-568.
- [2] Q. Ma, Y. Ouyang, F. Meng, M.N. Noolvi, S.P. Avvaru, U.A. More, T.M. Aminabhavi, M. Du, H. Liu, Y. Zhuang, M. Pang, T. Cai, Y. Cai, A review of pharmacological and clinical studies on the application of Shenling Baizhu San in treatment of Ulcerative colitis, *Journal of Ethnopharmacology* 244 (2019).
- [3] V. Chauhan, V. Patel, B. Akbari, R. Barse, Development and evaluation of biodegradable polymeric based long acting in situ forming microparticles of lhrh agonist goserelin acetate, *Indian Journal of Pharmaceutical Education and Research* 53(2) (2019) 216-224.
- [4] H. Patel, K. Dhangar, Y. Sonawane, S. Surana, R. Karpoomath, N. Thapliyal, M. Shaikh, M. Noolvi, R. Jagtap, In search of selective 11 β -HSD type 1 inhibitors without nephrotoxicity: An approach to resolve the metabolic syndrome by virtual based screening, *Arabian Journal of Chemistry* 11(2) (2018) 221-232.
- [5] S.P. Avvaru, M.N. Noolvi, T.M. Aminbhavi, S. Chkraborty, A. Dash, S.S. Shukla, Aromatase inhibitors evolution as potential class of drugs in the treatment of postmenopausal breast cancer women, *Mini-Reviews in Medicinal Chemistry* 18(7) (2018) 609-621.
- [6] P. Tanvi, V. Jain, Pharmacognostic standardization of *Oldenlandia corymbosa*, *International Journal of Pharmaceutical Research* 9(1) (2017) 22-25.
- [7] M.A. Shah, H. Patel, H. Raj, Methods for the estimation of ellagic acid and curcumin in antidiabetic herbal formulations - A review, *Eurasian Journal of Analytical Chemistry* 12(4) (2017) 295-311.
- [8] H.M. Patel, R. Pawara, A. Ansari, M. Noolvi, S. Surana, Design and synthesis of quinazolinones as EGFR inhibitors to overcome EGFR resistance obstacle, *Bioorganic and Medicinal Chemistry* 25(10) (2017) 2713-2723.
- [9] H.M. Patel, M.N. Noolvi, N.S. Sethi, A.K. Gadad, S.S. Cameotra, Synthesis and antitubercular evaluation of imidazo[2,1-b][1,3,4]thiadiazole derivatives, *Arabian Journal of Chemistry* 10 (2017) S996-S1002.
- [10] R. Kashyap, Role of secondary metabolites and medicinal plants tested for healing of gastric ulcer – A review, *International Journal of Pharmaceutical Research* 9(1) (2017) 5-15.
- [11] S.M. Kamble, H.M. Patel, S.N. Goyal, M.N. Noolvi, U.B. Mahajan, S. Ojha, C.R. Patil, In silico evidence for binding of pentacyclic triterpenoids to keap1-nrf2 protein-protein binding site, *Combinatorial Chemistry and High Throughput Screening* 20(3) (2017) 215-234.
- [12] S.D. Joshi, U.A. More, S.R. Dixit, S.V. Balmi, B.G. Kulkarni, G. Ullagaddi, C. Lherbet, T.M. Aminabhavi, Chemical synthesis and in silico molecular modeling of novel pyrrolyl benzohydrazide derivatives: Their biological evaluation against enoyl ACP reductase (InhA) and *Mycobacterium tuberculosis*, *Bioorganic Chemistry* 75 (2017) 181-200.

- [13] A.S. Deshmukh, P.N. Chauhan, M.N. Noolvi, K. Chaturvedi, K. Ganguly, S.S. Shukla, M.N. Nadagouda, T.M. Aminabhavi, Polymeric micelles: Basic research to clinical practice, *International Journal of Pharmaceutics* 532(1) (2017) 249-268.
- [14] T.M. Aminabhavi, S.P. Dharupaneedi, U.A. More, The role of nanotechnology and chitosan-based biomaterials for tissue engineering and therapeutic delivery, *Chitosan Based Biomaterials* 2017, pp. 1-29.
- [15] V. Abbot, P. Sharma, S. Dhiman, M.N. Noolvi, H.M. Patel, V. Bhardwaj, Small hybrid heteroaromatics: Resourceful biological tools in cancer research, *RSC Advances* 7(45) (2017) 28313-28349.
- [16] V. Thakor, M. Poddar, S. Dey, S.N. Manjula, S.V. Madhunapantula, R. Pawara, H.M. Patel, M.N. Noolvi, Exploring the anti-breast cancer potential of flavonoid analogs, *RSC Advances* 6(82) (2016) 79166-79179.
- [17] H.M. Patel, M.N. Noolvi, A.A. Shirkhedkar, A.D. Kulkarni, C.V. Pardeshi, S.J. Surana, Anti-convulsant potential of quinazolinones, *RSC Advances* 6(50) (2016) 44435-44455.
- [18] A. Patel, D. Shah, T.R. Desai, M.N. Noolvi, Mucoadhesive buccal films based on chitosan and carboxymethylated feronia limonia fruit pulp mucilage interpolymer complex for delivery of opioid analgesics, *Asian Journal of Pharmaceutics* 10(2) (2016) 137-143.
- [19] M.N. Noolvi, H.M. Patel, S. Kamboj, S.S. Cameotra, Synthesis and antimicrobial evaluation of novel 1,3,4-thiadiazole derivatives of 2-(4-formyl-2-methoxyphenoxy) acetic acid, *Arabian Journal of Chemistry* 9 (2016) S1283-S1289.
- [20] P. Nazema, A. Patel, Role of triptans in the management of acute migraine: A review, *Asian Journal of Pharmaceutics* 10(4) (2016) S437-S443.
- [21] V.C. Jain, B.V. Akbari, G. Sonara, K. Viradiya, H. Bhenjaliya, Design and development of polyherbal cream for skin repigmentation therapies in vitiligo, *International Journal of Pharmaceutical Research* 8(4) (2016) 76-79.
- [22] V. Bhardwaj, M.N. Noolvi, S. Jalhan, H.M. Patel, Synthesis, and antimicrobial evaluation of new pyridine imidazo [2,1b]-1,3,4-thiadiazole derivatives, *Journal of Saudi Chemical Society* 20 (2016) S406-S410.
- [23] P. Adil, D. Shah, T.R. Desai, M.N. Noolvi, Development and evaluation of chitosan and aloe vera gel mucilage interpolymer complex-based mucoadhesive buccal films of tramadol hydrochloride, *Asian Journal of Pharmaceutics* 10(1) (2016) S43-S49.
- [24] S.U. Upadhyay, V.C. Jain, U.M. Upadhyay, HPTLC fingerprinting of different extracts and estimation of phytoconstituents of *Ajuga bracteosa*, *International Journal of Pharmaceutical Research* 7(2) (2015) 119-128.
- [25] S.P. Roy, T. Kannadasan, R. Gupta, Screening of hepatoprotective activity of *Madhuca longifolia* bark on D-Galactosamine induced hepatotoxicity in rats, *Biomedical Research (India)* 26(2) (2015) 365-369.
- [26] M. Poddar, T.M. Aminabhavi, M. Patel, N. Singh, M.N. Noolvi, HIF inhibitors: New hope for cancer therapy, *Letters in Drug Design and Discovery* 12(9) (2015) 736-753.
- [27] H.M. Patel, B. Sing, V. Bhardwaj, M. Palkar, M.S. Shaikh, R. Rane, W.S. Alwan, A.K. Gadad, M.N. Noolvi, R. Karpoomath, Design, synthesis and evaluation of small molecule imidazo[2,1-b][1,3,4]thiadiazoles as inhibitors of transforming growth factor- β type-I receptor kinase (ALK5), *European Journal of Medicinal Chemistry* 93 (2015) 599-613.

- [28] H.M. Patel, P. Bari, R. Karpoornath, M. Noolvi, N. Thapliyal, S. Surana, P. Jain, Design and synthesis of VEGFR-2 tyrosine kinase inhibitors as potential anticancer agents by virtual based screening, *RSC Advances* 5(70) (2015) 56724-56771.
- [29] H. Patel, Y. Sonawane, R. Jagtap, K. Dhangar, N. Thapliyal, S. Surana, M. Noolvi, M.S. Shaikh, R.A. Rane, R. Karpoornath, Structural insight of glitazone for hepato-toxicity: Resolving mystery by PASS, *Bioorganic and Medicinal Chemistry Letters* 25(9) (2015) 1938-1946.
- [30] J. Ma, A.L. Porter, T.M. Aminabhavi, D. Zhu, Nano-enabled drug delivery systems for brain cancer and Alzheimer's disease: Research patterns and opportunities, *Nanomedicine: Nanotechnology, Biology, and Medicine* 11(7) (2015) 1763-1771.
- [31] T.M. Aminabhavi, M.N. Nadagouda, U.A. More, S.D. Joshi, V.H. Kulkarni, M.N. Noolvi, P.V. Kulkarni, Controlled release of therapeutics using interpenetrating polymeric networks, *Expert Opinion on Drug Delivery* 12(4) (2015) 669-688.
- [32] C.M. Setty, A.S. Deshmukh, A.M. Badiger, Hydrolyzed polyacrylamide grafted carboxymethylxyloglucan based microbeads for pH responsive drug delivery, *International Journal of Biological Macromolecules* 67 (2014) 28-36.
- [33] C.M. Setty, A.S. Deshmukh, A.M. Badiger, Hydrolyzed polyacrylamide grafted maize starch based microbeads: Application in pH responsive drug delivery, *International Journal of Biological Macromolecules* 70 (2014) 1-9.
- [34] C.M. Setty, A.S. Deshmukh, A.M. Badiger, Influence of process parameters on the performance of hydrolyzed polyacrylamide grafted maize starch based microbeads, *International Journal of Pharmaceutical Research* 6(2) (2014) 85-93.
- [35] M.P. Patel, M.R. Patel, R. Hasumati, M.N. Noolvi, N. Shah, Simultaneous estimation of amlodipine besylate and indapamide by dual wavelength spectrophotometric method for combined pharmaceutical dosage form, *Indian Drugs* 51(4) (2014) 50-54.
- [36] H.M. Patel, M.N. Noolvi, P. Sharma, V. Jaiswal, S. Bansal, S. Lohan, S.S. Kumar, V. Abbot, S. Dhiman, V. Bhardwaj, Quantitative structure-activity relationship (QSAR) studies as strategic approach in drug discovery, *Medicinal Chemistry Research* 23(12) (2014) 4991-5007.
- [37] R. Gupta, T. Kannadasan, S.P. Roy, Review on potential plant based drugs for hepatoprotective activity, *Journal of Chemical and Pharmaceutical Sciences* 7(2) (2014) 109-112.
- [38] R. Gupta, T. Kannadasan, S.P. Roy, Screening of hepatoprotective activity of *Mimusops elangi* fruit on d-galactosamine induced hepatotoxicity in rats, *Journal of Chemical and Pharmaceutical Sciences* 7(3) (2014) 229-232.
- [39] S.K. Das, P.V. Pansuriya, S.T. Shukla, K.J. Gohil, S.P. Roy, A. Choudhury, V.N. Sutariya, Preclinical evaluation of *Vallisneria spiralis* (roth) kuntze stem for its antiulcer and antioxidant activity in wistar albino rats, *Oriental Pharmacy and Experimental Medicine* 14(1) (2014) 7-13.
- [40] S.P. Roy, R. Gupta, D. Bhadra, T. Kannadasan, A review on some indigenous medicinal plants with hepatoprotective activity, *Journal of Chemical and Pharmaceutical Sciences* 6(2) (2013) 85-92.
- [41] H.M. Patel, M.N. Noolvi, A. Goyal, B.S. Thippeswamy, 2,5,6-Trisubstituted imidazo[2,1-b][1,3,4]thiadiazoles: Search for antihyperlipidemic agents, *European Journal of Medicinal Chemistry* 65 (2013) 119-133.
- [42] P.A. Kasad, Photolytic-Thermal Degradation study and method development of Rivaroxaban by RP-HPLC, *International Journal of PharmTech Research* 5(3) (2013) 1254-1263.

- [43] V. Kanani Vineeta, K.S. Muralikrishna, Development and validation of UV spectrophotometric method for estimation of Dapoxetine HCL in bulk and dosage Form, *International Journal of Drug Development and Research* 5(1) (2013) 161-167.
- [44] S.D. Joshi, K. Manish, A. Badiger, Synthesis and evaluation of antibacterial and antitubercular activities of some novel imidazo[2,1-b][1,3,4]thiadiazole derivatives, *Medicinal Chemistry Research* 22(2) (2013) 869-878.
- [45] V. Bhardwaj, P. Sharma, M.N. Noolvi, H.M. Patel, S. Chauhan, M.S. Chauhan, K. Sharma, Thermo-physical examination: Synthesized 2-furano-4(3H)-quinazolinone and open quinazolinone (diamide) anticancer analogs with sodium dodecyl sulfate, *Thermochimica Acta* 573 (2013) 65-72.
- [46] V. Bhardwaj, P. Sharma, M.N. Noolvi, H.M. Patel, S. Bansal, S. Lohan, G. Badola, Certain 2-furano-4(3H)-quinazolinone analogs: Synthesis, characterization and pharmacological evaluation, *Letters in Drug Design and Discovery* 10(4) (2013) 360-368.
- [47] A. Bhanot, R. Sharma, S. Singh, M.N. Noolvi, S. Singh, In vitro anti cancer activity of ethanol extract fractions of aerva lanata L, *Pakistan Journal of Biological Sciences* 16(22) (2013) 1612-1617.
- [48] H.J. Vekaria, M. Yogesh, K. Patel, Development and validation of HPTLC method for simultaneous estimation of cinitapride and omeprazole in combined dosage form, *Der Pharmacia Lettre* 4(5) (2012) 1467-1474.
- [49] H. Vekaria, K.S. Muralikrishna, M. Sorathiya, Development and validation of HPTLC method for simultaneous estimation of montelukast sodium and fexofenadine hydrochloride in combined dosage form, *Der Pharmacia Lettre* 4(3) (2012) 755-762.
- [50] V. Shah, H. Raj, Development and validation of derivative spectroscopic method for simultaneous estimation of cefixime trihydrate and azithromycin dihydrate in combined dosage form, *International Journal of Pharma and Bio Sciences* 3(3) (2012) P14-P25.
- [51] P.R. Samaresh, R. Gupta, T. Kannadasan, Hepatoprotective activity of Ethanolic extract of Madhuca longifoilia leaves on D-galactosamine induced liver damage in rats, *Journal of Chemical and Pharmaceutical Sciences* 5(4) (2012) 205-209.
- [52] D. Prajapati, H. Raj, Simultaneous estimation of mefenamic acid and dicyclomine hydrochloride by RP-HPLC method, *International Journal of Pharma and Bio Sciences* 3(3) (2012) P611-P625.
- [53] M.R. Patel, G.F. Patel, J.R. Patel, Development and validation of Spectrophotometric methods for simultaneous estimation of Indapamide and Amlodipine besylate in combined dosage form, *Research Journal of Pharmacy and Technology* 5(9) (2012) 1213-1217.
- [54] M.N. Noolvi, H.M. Patel, M. Kaur, Benzothiazoles: Search for anticancer agents, *European Journal of Medicinal Chemistry* 54 (2012) 447-462.
- [55] M.N. Noolvi, H.M. Patel, S. Kamboj, A. Kaur, V. Mann, 2,6-Disubstituted imidazo[2,1-b][1,3,4]thiadiazoles: Search for anticancer agents, *European Journal of Medicinal Chemistry* 56 (2012) 56-69.
- [56] M.N. Noolvi, H.M. Patel, S. Kamboj, A QSAR analysis of 2-phenoxy-N-substituted acetamide analogues as hypoxia-inducible factor-1(HIF-1) inhibitors: A rational approach to anticancer drug design, *Medicinal Chemistry* 8(4) (2012) 599-614.

- [57] L. Nagesh, S. Sivasamy, K.S. Muralikrishna, K.G. Bhat, Antibacterial potential of gall extract of *Quercus infectoria* against *Enterococcus faecalis*-an in vitro study, *Pharmacognosy Journal* 4(30) (2012) 47-50.
- [58] Y.G. Makani, H.A. Raj, Development and validation of first order derivative spectrophotometric method for simultaneous estimation of omeprazole and cinitapride in pharmaceutical dosage form, *International Journal of Pharma and Bio Sciences* 3(3) (2012) P70-P80.
- [59] A.S. Deshmukh, C.M. Setty, A.M. Badiger, K.S. Muralikrishna, Gum ghatti: A promising polysaccharide for pharmaceutical applications, *Carbohydrate Polymers* 87(2) (2012) 980-986.
- [60] I.M. Bagban, S.P. Roy, A. Chaudhary, S.K. Das, K.J. Gohil, K.K. Bhandari, Hepatoprotective activity of the methanolic extract of *Fagonia indica* Burm in carbon tetra chloride induced hepatotoxicity in albino rats, *Asian Pacific Journal of Tropical Biomedicine* 2(3 SUPPL.) (2012) S1457-S1460.
- [61] R. Shukla, S.J. Surana, A.U. Tatiya, S.K. Das, Investigation of hepatoprotective effects of piperine and silymarin on D-galactosamine induced hepatotoxicity in rats, *Research Journal of Pharmaceutical, Biological and Chemical Sciences* 2(3) (2011) 975-982.
- [62] R.J. Mandade, S.A. Sreenivas, D.M. Sakarkar, A. Choudhury, Pharmacological effects of aqueous-ethanolic extract of *Hibiscus rosasinensis* on volume and acidity of stimulated gastric secretion, *Asian Pacific Journal of Tropical Medicine* 4(11) (2011) 883-888.
- [63] A.B. Desai, V.G. Kagathara, H. Joshi, A.T. Rangani, H. Mungra, Evaluation of Antiamnesic effect of Solasodine in Mice, *International Journal of PharmTech Research* 3(2) (2011) 732-740.
- [64] B.V. Akbari, B.P. Valaki, V.H. Maradiya, A.K. Akbari, G. Vidyasagar, Development and evaluation of Orodispersible tablets of Rosuvastatin Calcium-HP- β -CD inclusion complex by using different superdisintegrants, *International Journal of Pharmacy and Technology* 3(1) (2011) 1842-1859.
- [65] M.G. Saralaya, P. Patel, M. Patel, S.P. Roy, A.N. Patel, Antidiarrheal activity of methanolic extract of *Moringa oleifera* lam roots in experimental animal models, *International Journal of Pharmaceutical Research* 2(2) (2010) 35-39.
- [66] S.P. Roy, D. Shirode, T. Patel, C.S. Shastry, N. Gheewala, G. Sonara, S. Ramachandra Setty, S.V. Rajendra, Antioxidant and hepatoprotective activity of *Madhuca longifolia* (koenig) bark against CCl_4 - induced hepatic injury in rats: In vitro and In vivo studies, *Research Journal of Pharmaceutical, Biological and Chemical Sciences* 1(1) (2010) 1-10.
- [67] G.F. Patel, N.R. Vekariya, R.B. Dholakiya, Estimation of Aspirin and Atorvastatin Calcium in combine dosage form using derivative spectrophotometric method, *International Journal of Pharmaceutical Research* 2(1) (2010) 62-66.
- [68] R.R. Parmar, H.A. Bhuvra, A.A. Joshi, R.B. Jadhav, Optimization of extraction method for recovery of antioxidant phenolics from loranthaceae mistletoes, *Asian Journal of Chemistry* 22(5) (2010) 3851-3855.
- [69] S.D. Chavan, N.L. Shetty, M. Kanuri, Comparative evaluation of garlic extract mouthwash and chlorhexidine mouthwash on salivary *Streptococcus mutans* count-An in vivo study, *Oral Health and Preventive Dentistry* 8(4) (2010) 369-374.
- [70] K. Barot Hitesh, G. Mallika, B. Sutariya Bhavin, S. Jitendra, L.V.G. Nargund, Synthesis of Nitrogen Mustards of fluoro- benzothiazoles of pharmacological interest, *Research Journal of Pharmaceutical, Biological and Chemical Sciences* 1(1) (2010) 124-129.



- [71] B.V. Akbari, B.P. Patel, R.B. Dholakiya, B.G. Shiyani, D.J. Lodhiya, Development and evaluation of taste masked suspension of prokinetic agent by using ion exchange resin, *International Journal of PharmTech Research* 2(1) (2010) 240-245.
- [72] B.V. Akbari, R.B. Dholakiya, B.G. Shiyani, D.J. Lodhiya, Design, development and characterization of mouth dissolving tablets of cinnarizine using super-disintegrants, *International Journal of PharmTech Research* 2(1) (2010) 97-105.
- [73] N.R. Vekariya, G.F. Patel, H.S. Bhatt, M.B. Patel, R.B. Dholakiya, G.K. Ramani, Application of TLC-densitometry method for simultaneous estimation of telmisartan and amlodipine besylate in pharmaceutical dosage form, *International Journal of PharmTech Research* 1(4) (2009) 1644-1649.
- [74] G.F. Patel, N.R. Vekariya, H.S. Bhatt, Application of TLC-densitometry method for simultaneous determination of lopinavir and ritonavir in capsule dosage form, *Oriental Journal of Chemistry* 25(3) (2009) 727-730.
- [75] D.J. Lodhiya, D.J. Mukherjee, R.B. Dholakiya, B.V. Akbari, B.G. Shiyani, H.N. Lathiya, Gastroretentive system of atenolol using HPMC K15, *International Journal of PharmTech Research* 1(4) (2009) 1616-1620.
- [76] A. Paharia, A.K. Yadav, G. Rai, S.K. Jain, S.S. Pancholi, G.P. Agrawal, Eudragit-coated pectin microspheres of 5-Fluorouracil for colon targeting, *AAPS PharmSciTech* 8(1) (2007) E1-E7.
- [77] K.L. Krishna, M. Paridhavi, S.S. Agrawal, Physico-chemical standardization of Sufoof-e-Suzak Qawi an Unani polyherbomineral formulation, *Indian Drugs* 44(3) (2007) 220-223.
- [78] D. Gogna, S.K. Jain, A.K. Yadav, G.P. Agrawal, Microsphere based improved sunscreen formulation of ethylhexyl methoxycinnamate, *Current Drug Delivery* 4(2) (2007) 153-159.
- [79] S.K. Jain, G.P. Agrawal, N.K. Jain, A novel calcium silicate based microspheres of repaglinide: In vivo investigations, *Journal of Controlled Release* 113(2) (2006) 111-116.
- [80] A.K. Jain, S.K. Jain, A. Yadav, G.P. Agrawal, Controlled release calcium silicate based floating granular delivery system of ranitidine hydrochloride, *Current Drug Delivery* 3(4) (2006) 367-372.



Research collaboration outcomes

RSC Advances



PAPER



Cite this: *RSC Adv.*, 2016, 6, 79166

Exploring the anti-breast cancer potential of flavonoid analogs†

Vanrajsinh Thakor,^a Mayur Poddar,^a Sumit Dey,^b S. N. Manjula,^b SubbaRao V. Madhunapantula,^c Rahul Pawara,^d Harun M. Patel^{*d} and Malleshappa N. Noolvi^{*a}

In the course of our search for new antitumor agents for breast cancer, novel flavone derivatives were synthesized, characterized and examined for their antitumor activities against breast cancer cell lines. In initial screening, analogs **7a** [3-(5-amino-1,3,4-thiadiazol-2-yl)methoxy-2-phenyl-4*H*-chromen-4-one] and **7b** [3-(5-amino-1,3,4-thiadiazol-2-yl)methoxy-2-(4-methoxyphenyl)-4*H*-chromen-4-one] were found to be effective against the estrogen receptor negative cell line (MDA-MB 453), which was followed by their evaluation in five dose assays. In addition, mechanistic studies of **7a** and **7b** were performed by cytometric analysis and electrophoretic studies and it was observed that apoptosis is a mechanism of cell death, confirmed morphologically by acridine orange/ethidium bromide double staining and TUNEL analysis. Further *in vivo* evaluation of the anti-tumor activity of compound **7a** and **7b** by Ehrlich Ascites Carcinoma (EAC) model and related studies confirms the anti-breast cancer potential of flavonoid analogs.

Received 3rd June 2016
Accepted 3rd August 2016
DOI: 10.1039/c6ra14428d
www.rsc.org/advances

1. Introduction

Breast cancer is the most commonly diagnosed malignancy among women with more than one million new cases diagnosed per year throughout the world.^{1,2} Despite advances in the early detection of breast cancer and the advent of novel targeted therapies, breast cancer still remains a significant public health problem due to the involvement of multiple aberrant and redundant signaling pathways in the tumorigenesis and the development of resistance to the existing therapeutic agents. The currently available breast cancer therapies achieve meaningful clinical results in only 30–40% of the patients.³ The efficacy of current chemotherapeutics is low and undesirable side effects are still unacceptably high.⁴ Hence, the development of novel, efficient, and less toxic anti-breast cancer agents remains an important and challenging goal of medicinal chemists worldwide.

The female hormone estrogen stimulates breast cell division leading to the increase in risk of permanent damage to DNA.⁵ Compounds that can regulate the apoptosis of cancer cells are of a high medical significance.⁶ Natural products (NPs) have

played a valuable role in the drug discovery and development.^{7–9} Newman and Cragg¹⁰ reported that in the case of cancer around 79% of FDA-approved drugs during a period of 1981–2010 are either natural products or their based/mimicked-compounds. NPs are chosen through evolutionary process *via* lead optimization to interact with various enzymes/proteins and thus represent biologically relevant regions of the vast chemical space.^{11–13} Flavopiridol, a semisynthetic flavones analog, acts as CDK9 inhibitor, is FDA-approved orphan drug for acute myeloid leukaemia. It has been reported that myricetin, (flavonoid compound) could decrease pancreatic cancer growth *via* induction of cell apoptosis.¹⁴ LY294002 (flavonoid analogue) entered clinical trials as a potential antineoplastic agent.¹⁵ Effects of phytoestrogens in cancer prevention have been reported for decades.^{16–18} Since then many molecular mechanisms underlying these effects have been identified. Targets of phytoestrogens comprise steroid receptors, steroid metabolising enzymes, elements of signal transduction and apoptosis pathways, and even the DNA processing machinery.¹⁸ Phytoestrogens include chalcones (A), flavones (B) and isoflavones (C) which are non-steroidal compounds possessing anti-estrogenic activity (Fig. 1).¹⁹

In light of these findings and in continuation of our research for novel anti-cancer agents^{20–23} in the present study, new series of flavone derivatives has been synthesized and screened *in vitro* for cytotoxicity by sulphorhodamine B assay. Five dose assay in estrogen receptor negative cell line (MDA-MB 453) and determination of IC₅₀ by SRB assay was also performed. In addition, mechanistic study was done with cytometric analysis and electrophoretic determination of apoptosis. Further *in vivo* activity

^aDepartment of Pharmaceutical Chemistry, Shree Dharmantary Pharmacy College, Kim (Surat)-394110, Gujarat, India. E-mail: mnoolvi@yahoo.co.uk

^bDepartment of Pharmacology, JSS College of Pharmacy, Mysore-570015, Karnataka, India

^cDepartment of Biochemistry, JSS Medical College, Mysore-570015, Karnataka, India

^dDepartment of Pharmaceutical Chemistry, R. C. Patel Institute of Pharmaceutical Education and Research, Shitpur, District Dhule-425 405, Maharashtra, India

† Electronic supplementary information (ESI) available. See DOI: 10.1039/c6ra14428d



REVIEW ARTICLE

Aromatase Inhibitors Evolution as Potential Class of Drugs in the Treatment of Postmenopausal Breast Cancer Women

Stephen Paul Avvaru¹, Malleshappa N. Noolvi^{1*}, Tejraj M. Aminbhavi¹, Sudipta Chkraborty², Ashutosh Dash² and Shyam S. Shukla³

¹Department of Pharmaceutical Chemistry, Shree Dhanvantary Pharmacy College, Kim, Surat, 394110, India; ²Radiopharmaceuticals Division, Bhabha Atomic Research Centre, India; ³Department of Chemistry and Biochemistry, Lamar University, Beaumont, Texas 77710, USA

Abstract: Aromatase inhibitors are class of drugs that inhibit aromatase, a rate limiting enzyme in the biosynthesis of estrogens from their corresponding androgens. Estrogens play a vital role in the development and growth of breast tumors especially in postmenopausal women apart from their important functions in cell homeostasis. The reduction of estrogen physiological concentration through aromatase inhibition is one of the most important therapeutic strategies against this cancer type. The third-generation aromatase inhibitors are now used as first-line therapy in the treatment of early and metastatic breast cancer in postmenopausal women. However the quest for new class of drugs still stays indispensable to evade the danger of conceivable rising resistances to existing drugs, toxicity and unwanted side effects due to chronic treatment. The current review deals with recent advances in understanding of aromatase, its mechanism and research in the development of various novel chemotypes as aromatase inhibitors. The new challenges and the fast changing trends in bringing rational approach in aromatase inhibitors to a different level like research in dual/multiple target enzyme inhibition strategies, radiolabeling of aromatase inhibitors as theranostic agents; the development of new computational models for complete understanding of aromatase enzyme and its substrate/ligand interactions will bring in holistic approach to comprehensive inhibition of aromatase and other relevant enzymes for effective treatment and monitoring of postmenopausal breast cancer.

ARTICLE HISTORY

Received: September 06, 2016
Revised: December 15, 2016
Accepted: February 06, 2017

DOI:
10.2174/1389557517666171101100902

Keywords: Aromatase inhibitors, estrogen, breast cancer, dual inhibitors, molecular modeling, radiolabeling.

1. INTRODUCTION

An estimated 1.67 million new breast cancer cases were diagnosed in 2012 which makes it the second most common cancer among women in the world [1]. Around 70-80 per cent of postmenopausal breast cancer patients are found to be estrogen-dependent breast cancer cases [2]. Endogenous estrogens affect the cellular roles of genes involved in cell division, protein expression, cell communication [3] and play a vital role in the development and growth of breast tumors [4]. Ovaries and placenta are the major sources of estrogen biosynthesis and to a lesser degree in testes, liver, adrenal glands, breasts and fat cells. Androgens are known to be precursors for the synthesis of Estrogens.

1.1. Aromatase

Estrogens are biosynthesized from their corresponding androgens by aromatase or estrogen synthetase (CYP19A1; EC 1.14.14.1) a rate-limiting enzyme [5]. It is a microsomal

membrane-bound cytochrome p-450 monooxygenase complex comprising of aromatase cytochrome p-450 arom and NADPH-cytochrome p-450 reductase. Cytochrome p-450 arom is a heme protein that binds the steroid substrate, molecular oxygen and catalyses oxidation. The reductase is a flavoprotein, found ubiquitously in endoplasmic reticulum, and responsible for transferring reducing equivalents from NADPH to cytochrome p-450 arom [4, 6]. The gene CYP19 expresses cytochrome p-450 arom, and the gene is located on chromosome 15 in humans. The human CYP19 gene includes nine coding exons, 2-10. Tissue-specific regulation of the aromatase gene in various tissues is determined by tissue-specific promoters. In placenta, p-450 arom transcripts contain promoter 1.1 or 1.2. Ovarian transcripts comprise of promoter PII [7, 8]. Aromatase activity in human adipose and ovarian granulosa cells is associated to complex multifactorial regulation and changes in aromatase activity are correlated with changes in the levels of mRNA encoding p-450 arom [9].

Aromatase cytochrome P450 is the rate limiting enzyme that catalyzes androgens conversion to estrogens in vertebrates [10]. It is the hallmark androgenic specificity that is C19 steroids with 4-ene-3-one system that sets aromatase

*Address correspondence to this author at the Department of Pharmaceutical Chemistry, Shree Dhanvantary Pharmacy College, Kim, Surat, 394110, India; Tel/Fax: +912621233344; E-mail: noolvimalleshappa@gmail.com



Cite this: *RSC Adv.*, 2017, 7, 28313

Small hybrid heteroaromatics: resourceful biological tools in cancer research

Vikrant Abbot,^a Poonam Sharma,^{id}^a Saurabh Dhiman,^a Mallehappa N. Noolvi,^b Harun M. Patel^c and Varun Bhardwaj^{*a}

Nowadays, hybrid drugs containing two or more covalently linked known potential pharmacophores are designed to simultaneously modulate multiple targets of multifactorial diseases to overcome the side effects associated with a single drug. In this review, an overview of the design strategies employed by various scientists over the past 20 years has been presented. The overview includes the synthesis of different chemical structure-based anticancer hybrids using molecular hybridization techniques. To tackle one of the world's most devastating diseases such as cancer, researchers have exploited the molecular hybridization (MH) technique to synthesize different anticancer hybrids, which include hybrids based on azole, camptothecin, chalcone, pyrrolobenzodiazepine (PBD), coumarin, colchicine, platinum, and some miscellaneous structures. The selection of two or more moieties for generating the hybrid drug is generally aided by the observed (or anticipated) synergistic or additive pharmacological activities of each single moiety. This eventually leads to the identification of novel and better active chemical entities with a superior profile as compared to the parent moieties. In addition to the design strategies, this review also highlights the structure–activity relationship (SAR), mechanism of action, and key features of the synthesized anticancer hybrids.

Received 3rd October 2016
Accepted 17th April 2017

DOI: 10.1039/c6ra24662a
rsc.li/rsc-advances

^aDepartment of Biotechnology, Bioinformatics and Pharmacy, Jaypee University of Information Technology, Wanknaghat, Solan, Himachal Pradesh 173234, India. E-mail: varunmilton@yahoo.com; varun.bhardwaj@mail.juit.ac.in; Tel: +91-94189-05865; +91-8427653284

^bDepartment of Pharmaceutical Chemistry, Shree Dhanvantary Pharmacy College, Kim, Surat, 3941110, Gujarat, India

^cDepartment of Pharmaceutical Chemistry, R.C. Patel Institute of Pharmaceutical Education and Research, Shirpur, Dhule, 425405, Maharashtra, India

1. Introduction

Cancer is a cell cycle disease characterized by rapid, uncontrolled, and pathological proliferation of abnormal cells. It is one of the most formidable diseases in the world.^{1,2} The most frequent forms of cancer are lung cancer, breast cancer, colorectal cancer, stomach cancer, and prostate cancer; among these, prostate cancer (male) and breast cancer (female)



Vikrant Abbot obtained his Bachelor's degree in Pharmacy from the Punjab Technical University in 2009. He started his professional career working as a Production Chemist at Comed Pharmaceuticals Ltd. Baddi for 2.5 years. Thereafter, he completed his Master's degree in Medicinal Chemistry from JUIT, Wanknaghat in 2014. Then, he joined academics and is currently working as an

Assistant Professor at ISF College of Pharmacy, Moga, Punjab. His present research interests include design and synthesis of small molecules with anticancer and antimicrobial profiles.



Poonam Sharma obtained her Master's in Philosophy in 2003 with a gold medal and completed her PhD in chemistry in 2006 from Himachal Pradesh University (India). She was awarded a UGC project fellowship for her PhD research. She was also awarded the Fast Track Young Scientist research project by DST in 2010. Presently, she is working as an Assistant Professor at the Jaypee Univer-

sity of Information Technology, Wanknaghat, Solan (India). Her research revolves around physicochemical drug interactions, thermodynamics, heterocyclic bioactive analogs and topical drug delivery.



Chapter contents Book contents

Outline

Abstract

Keywords

1. Introduction
2. Physicochemical properties
3. Regulatory status
4. Applications in drug delivery
5. Concluding remarks and future directions

References

Show full outline

Figures (14)



Natural Polysaccharides in Drug Delivery and Biomedical Applications

2019, Pages 59-100



Chapter 3 - Sodium alginate in drug delivery and biomedical areas

Kiran Chaturvedi¹, Kuntal Ganguly¹, Uttam A. More², Kakarla Raghava Reddy³, Tanavi Dugge⁴, Balaram Naik⁴, Tejraj M. Aminabhavi¹, Malleshappa N. Noolvi²

- ¹ Department of Pharmaceutical Engineering and Polymer Science, SET's College of Pharmacy, Dharwad, India
- ² Department of Pharmaceutical Chemistry, Shree Dhanvantary Pharmacy College, Kim, Surat, India
- ³ School of Chemical and Biomolecular Engineering, The University of Sydney, Sydney, NSW, Australia
- ⁴ Shri Dharmasthala Manjunath Dental College, Dharwad, India

Available online 26 July 2019.

Show less

Outline

Highlights

Abstract

Graphical abstract

Keywords

1. Introduction
2. Molecular modeling/docking studies
3. Results and discussion
4. Experimental section
5. Biological activity
6. Conclusion

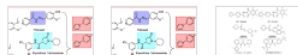
Acknowledgements

Appendix A. Supplementary material

References

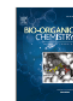
Show full outline

Figures (8)



Bioorganic Chemistry

Volume 75, December 2017, Pages 181-200



Short communication

Chemical synthesis and *in silico* molecular modeling of novel pyrrolyl benzohydrazide derivatives: Their biological evaluation against enoyl ACP reductase (InhA) and *Mycobacterium tuberculosis*

Shrinivas D. Joshi^{a,*,} Uttam A. More^{a, b,} Sheshagiri R. Dixit^c, Sunil V. Balmi^a, Basavaraj G. Kulkarni^a, Geeta Ullagaddi^a, Christian Lherbet^{c, d,} Tejraj M. Aminabhavi^a

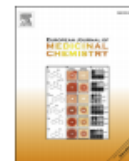
- ^a Novel Drug Design and Discovery Laboratory, Department of Pharmaceutical Chemistry, Soniya Education Trust's, College of Pharmacy, Sangolli Rayanna Nagar, Dharwad 580 002, India
- ^b Shree Dhanvantary Pharmacy College, Kim, Surat, India
- ^c Université de Toulouse, UPS, Laboratoire de Synthèse et Physico-chimie de Molécules d'Intérêt Biologique, LSPCMIB, 118 Route de Narbonne, F-31062 Toulouse Cedex 9, France
- ^d ITAV-USR3505, Université de Toulouse, CNRS, UPS, F-31106 Toulouse, France



Contents lists available at ScienceDirect

European Journal of Medicinal Chemistry

journal homepage: <http://www.elsevier.com/locate/ejmech>



Original article

Design, synthesis and evaluation of small molecule imidazo[2,1-*b*][1,3,4]thiadiazoles as inhibitors of transforming growth factor- β type-I receptor kinase (ALK5)



Harun M. Patel ^a, Baljeet Sing ^b, Varun Bhardwaj ^c, Mahesh Palkar ^a,
Mahamadhanif S. Shaikh ^a, Rajesh Rane ^a, Wesam S. Alwan ^a, Andanappa K. Gadad ^d,
Malleappa N. Noolvi ^{e,*}, Rajshekhar Karpoormath ^{a,**}

^a Department of Pharmaceutical Chemistry, College of Health Sciences, University of KwaZulu-Natal (Westville Campus), Private Bag X54001, Durban 4000, South Africa

^b Department of Pharmaceutical Chemistry, ASBASJSM College of Pharmacy, Bela (Ropar) 140111, Punjab, India

^c Department of Biotechnology, Bioinformatics and Pharmacy, Jaypee University of Information Technology, Waknaghat, Solan 173234, Himachal Pradesh, India

^d School Pharmacy, Faculty of Medical Sciences, EWMS Complex, Mt. Hope, The University of the West Indies, Trinidad and Tobago, West Indies

^e Department of Pharmaceutical Chemistry, Shree Dhanvantary Pharmacy College, Kim (Surat) 3941110, Gujarat, India

ARTICLE INFO

Article history:

Received 30 May 2014

Received in revised form

21 August 2014

Accepted 2 September 2014

Available online 3 September 2014

Keywords:

Synthesis imidazo [2,1-*b*][1,3,4]thiadiazole

ALK5

XP docking

Lipinski's rule

ABSTRACT

A new series of imidazo[2,1-*b*][1,3,4]thiadiazoles **5(a–g)**, **6(a–g)**, **9(a–i)** and **12(a–h)** were synthesized as transforming growth factor- β (TGF- β) type I receptor (also known as activin receptor-like kinase 5 or ALK5) inhibitors. These compounds were evaluated for their ALK5 inhibitory activity in an enzyme assay and their TGF- β -induced Smad2/3 phosphorylation inhibitory activity in a cell-based assay. Compound **6d**, 2-(5-((2-cyclopropyl-6-(4-fluorophenyl) imidazo [2,1-*b*][1,3,4]thiadiazol-5-yl)methylene)-4-oxo-2-thioxothiazolidin-3-yl) acetic acid, shows prominent ALK5 inhibition ($IC_{50} = 0.0012 \mu\text{M}$) and elective inhibition (91%) against the P38zkinase at $10 \mu\text{M}$. The binding mode of compound **6d** by XP docking studies shows that it fits well into the active site cavity of ALK5 by forming broad and tight interactions. Lipinski's rule and *in silico* ADME pharmacokinetic parameters are within the acceptable range defined for human use thereby indicating their potential as a drug-like molecules.

© 2014 Elsevier Masson SAS. All rights reserved.

1. Introduction

Transforming growth factor-beta (TGF- β) is a ubiquitous cytokine that affects various biological processes such as regulation of cell proliferation, immune responses, growth, differentiation, angiogenesis, and apoptosis of different cell types [1]. TGF- β 1 transduces signals through two highly conserved single transmembrane serine/threonine kinases, the type I and type II TGF- β receptors (T β R-I and T β R-II, respectively) [2]. T β R-II activates T β R-I upon formation of the ligand–receptor complex by hyperphosphorylating serine/threonine residues in the GS region of the T β R-I or activin-like kinase (ALK5), which creates a binding site for Smad proteins. The activated T β R-I in turn phosphorylates Smad2/

Smad3 proteins at the C-terminal SSXS-motif thereby causing dissociation from the receptor and heteromeric complex formation with the Smad4 [3–5]. Smad complexes translocate to the nucleus, assemble with specific DNA-binding co-factors and co-modulators to finally activate transcription of an extracellular matrix component, and inhibitors of matrix-degrading proteases [6]. Therefore, it becomes evident that inhibition of ALK5 phosphorylation of Smad2/Smad3 could reduce TGF- β 1-induced excessive accumulation of the extracellular matrix. Small molecules inhibitors of TGF- β R1 offer an attractive way to regulate the TGF- β pathway and can consequently find applications in the treatment of various diseases, especially, cancer [7]. Our on-going interest in the design and synthesis of novel anti-cancer agents [8–13], and recent reports by Hoelzemann and collaborators [14] suggesting the imidazo[2,1-*b*][1,3,4]thiadiazoles scaffold as a template to the design of inhibitors of ALK5; inspired us to synthesize and *in vitro* evaluated imidazo [2,1-*b*][1,3,4]thiadiazoles **5(a–g)**, **6(a–g)**, **9(a–i)** and **12(a–h)** for the ALK5 inhibitory activity in an enzyme assay and their TGF- β

* Corresponding author.

** Corresponding author.

E-mail addresses: malleesh7301@rediffmail.com, mnoolvi@yahoo.co.uk (M.N. Noolvi), karpoormath@ukzn.ac.za, rvk2006@gmail.com (R. Karpoormath).

Patent Details

Sr. No.	Product / Process	Country	Year	Patent Application Number
1	In situ forming micro particle of Gosereline	India	2016	201621030794
2	Development of gosereline acetate long acting parenteral depot formulation	India	2016	201621030797
3	New benzothiazole coumarin derivatives A Process and user thereof	India	2016	201621034608
4	New coumarin derivatives A Process and user thereof	India	2016	201621034607
5	Formulation and evaluation of sodium alginate and HPMC K4M based in situ gel containing trifluridine	India	2018	201821010721
6	New 2,4-diaminopyrimidine derivatives, A process and uses thereof	India	2018	201821010739
7	New indole derivatives, A process and uses thereof	India	2018	201821010737
8	New 7-chloro-2-furanyl quinazolin-4(3H)-one derivatives and preparation thereof	India	2018	201821010736
9	New 7-chloro-2-methyl quinazolin-4(3H)-one derivatives and preparation thereof	India	2018	201821010735
10	New 2-furanyl quinazolin-4(3H)-one derivatives and preparation thereof	India	2018	201821010730
11	Finasteride loaded nanoemulsion for treatment of androgenic alopecia	India	2018	201821010727
12	Polysaccharide and cellulose derivatives based hydrogels for agricultural needs thereof	India	2018	201821010724