## DRUG PROCESS CHIEMISTRY LABORATORY



Dr. Manjinder Singh Gill Assistant Professor

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#### Research Area

- Synthesis of therapeutically important compounds with focus on anti-diabetics, anti-mycobacterial, anticancer and anti-HIV agents
- Drug Process research and development and scale-up of drug molecules, APIs and herbals
- Synthesis and characterization of process related impurities
- Synthesis of drug-drug conjugates and complexes to enhanced therapeutic profile
- Green chemistry: The reactions in aqueous, aqueous-organic media and use of switchable solvents
- Total synthesis of therapeutically important compounds

### **Teaching Courses**

- PT-510: Industrial process and scale-up techniques (M. Tech. Sem-I)
- PT-560: Synthetic aspects of process chemistry (M. Tech. Sem-I)
- PT-610: Topics relevant to drugs and pharmaceutical industry (M. Tech. & MBA Sem-II)
- PT-630: Synthetic bulk drug technology (M. Tech. Sem-II)
- PT-820: Topics in organic process chemistry (Ph. D)

### Industrial Research Projects (selected)

- Scale-up and Development Batches for ISLLC-361 Project (7 projects)
- Scale-up and validation of Industrial product. compound (4 projects)
- Scale up studies of curcumin extraction
- Drying Trials of AZPH
- VTD drying trial of cosmetic products
- Scale-up and validation studies of APIs: Cefozopran, Entecavir, Candesartan, Pioglitazone HCl, Fondaparinux, & Valsartan
- Scale-up and validation studies of Process R&D and scale up studies of antioxidant products (2 projects)
- Process R&D and scale-up of anti-inflammatory drugs (4 projects)
- Scale-up and validation studies of Vinadifformine
- Scale-up and validation studies involving Vinadifformine conversion into Vincamine
- Scale-up and validation studies involving synthesis of Vinprocetine
- Scale-up and validation studies of hydrogenation (2 projects)
- Scale-up and development of a drug candidate
- Process development for the final step of Levosulpiride synthesis

# Research Group

### Ph.D Students



Rohani Prasad Burman (vermanniper@yahoo.com)

Design synthesis and process development of novel therapeutically important compounds and conjugates



### Dinesh Kumar Tanwar (tanwar\_dt@ymail.com)

Development of New Processes and Scale-up of Marketed Drugs, APIs and design and synthesis of Novel Antidiabetic Agents

M. Tech (2015-17)



Ravi Vaghela (vaghelaravi469@gmail.com)

Process for preparation of Cabozantinib, A tyrosine kinase inhibitor



### Vishakha Dhiman (vishakha.dhiman93@gmail.com)

Design of New synthetic pathway for substituted Hydantoins and RU58841- an antibaldness compound



Krishan Verma (krishanverma64.kv@gmail.com)

Application of Green/Switchable solvents in API synthesis



#### **Jyoti Bhatti** (jyotibhattii755@gmail.com)

To develop the synthetic route of recently isolated bioactive molecules originated from marine sources-Carpatamide-A, B and C, Sanjunolide



Divakar Singh (divakarsingh420@gmail.com)

Facile synthesis of Alogliptine and Trelagliptine



### Pinigani Vani Nikhitha (vani.nikitha0324@gmail.com)

Aspartic-acid based synthesis of anti-diabetic agents

# Alumni

## M.Tech. 2014-16

Anjali Ratan Patent Research Analyst at Stellarix Consultancy Services Pvt. Ltd., Jaipur	Process for Preparing Substituted Hydantoins and Synthesis of Antiepileptic Drug Ethotoin
Consultancy Services I vi. Etd., Janpur	
Bhawana Deshmukh	Process for the Preparation of 1,2,4-Triazol-3-One and Synthesis of its Analogues as Antidepressant Agents
Devendra Rajak	An improved and scalable synthesis of indolic enamides: Coscinamide A, B and their
Research Associate, CRD at ARK Healthcare Pvt Ltd., Derabassi	analogues
Pinninti Dileep kumar	Synthesis of Isatin semicarbazone as anticonvulsant agents
Patent Research Analyst at Stellarix Consultancy Services Pvt. Ltd., Jaipur	
Shantanu Gupta	Study toward Synthesis of Carpatamide A-B, Cytotoxic Arylamine Derivative from a
Research Associate, ARD at Biocon Limited, Bangalore	Marine Derived Streptomyces sp.
M.Tech. 2013-15	

Kamlesh Kumar Jatav	Studies on various reactions in aqueous media and switchable solvents
Anchal Singh	Synthesis and process development of raloxifene and process related impurities
M. Ravi	Synthesis and process development of OXI8006, combretastatin like indole as a vascular disrupting agents
Ripudaman	Synthesis and process development of rizatriptan and process related impurities
Bhagwana Ram	Synthesis and process development of amiodarone and their dervatives as anti leishmaniel derivatives
M. Siddardh Sai	Synthesis of novel isatin derivatives and their biological evaluation as anti-HIV compounds

## M.Tech. 2012-14

P. Sravan Kumar	Synthesis of noscapine-ascorbic acid conjugates
Nikunj Patel	Process research and development for synthesis of an anticancer drug vismodegib
Dasari Manikanta	Methodology development of regioselective ring opening of aspartic acid cyclic anhydride with various nucleophiles
Laxminarayana	Synthesis of novel drug lipid conjugates
Raghvender Reddy	Synthesis of novel sildenafil-fluoxetine hybrid molecules
Nishtha	A novel efficient approach towards the synthesis of potent insulinotropic 4-hydroxyisoleucine
Nandini Sarviya	Process development of propafenone synthesis and its analogues for MDR
Kumaraswami Musku	Synthesis and biological evaluation of novel indole derivatives as antitubercular agents

## M.Tech. 2011-13

Banu Petiwala	Synthesis of aromatase inhibitors
D. Ramya Sri	Synthesis of nitroindazole and benzimidazole derivatives
Gurpreet Kaur	Synthesis of curcumin-drug conjugates
Mukul Dhiman	Synthesis of drug-Phospholipid conjugates

## M.Tech. 2010-12

Monika Jain	Synthesis of 5-Arylidene-2-Imino-4-Thiazolidinones
Rohani Prasad Burman	An approach to Sulfonylurea from Sulfonylchloride and Urea derivatives
Neetu Dayal	Studies Towards Synthesis of (±)-Funebrine
Santosh Kumar Manupati	Synthesis of 1,2,5-substituted pyrrols

# M.Tech. 2009-11

Amit Tailor	Synthesis of Novel Imidazole and Thiazole Derivatives as Antimicrobial Agents

## Publications

- 1. **M. S. Gill** and D. K. Tanwar. A Facile Synthesis of *N*-alkyl *O*-aryl Cabamates and their Use in Synthesis of Other Therapeutically Important Compounds. Abstract published in 99th Canadian Chemistry Conference and Exhibition. Halifax, NS. June 5-9, 2016.
- 2. H. Pawar, S. K. Surapaneni, K. Tikoo, C. Singh, R. Burman, M. S. Gill, and S. Suresh: Folic acid Functionalized long-circulation co-encapsulated docetaxel and curcumin solid lipid nanoparitcles: In vitro evaluation, pharmacokinetic and biodistribution in rats. Drug Delivery, 1-16, 2016.
- 3. F. J. Bolla, P. Iyer. **M.S. Gill** and M.E. Sobhia: *In silico* Identification and Validation of Targets for Novel Scaffold, 2-Thiazolylimino-5-benzylidin-thiazolidin-4-one. *Molecular Diversity*, Volume 19 (4), 855-870, 2015.
- 4. V. Avasathi, H. Pawar, C. P. Dora, P. Bansod, **M.S. Gill** and S. Suresh: A novel nanogel formulation of Methotrexate for topical treatment of psoriasis: optimization, *in vitro* and *in vivo* evaluation. *Pharm Dev Technol*, 1-9, 2015.
- 5. M. Jain, R. P. Burman, V. S. Negi, S. S. Jhamb, A. Mital and M. S. Gill: Synthesis and Biological Evaluation of 2-thiazolylimino-5-arylidene-4- thiazolidinone Derivatives as Potent Antimycobacterial Agents. *Anti-Infective Agents*, 13, 105-113, 2015.
- 6. Charan Singh, T. D. Bhatt, **M. S. Gill**, S. Suresh: Novel rifampicin-phospholipid complex for tubercular therapy: synthesis, physicochemical characterization and in-vivo evaluation. *International Journal of Pharmaceutics* 460, 220–227, 2014.
- 7. A.K. Choubey, C. P. Dora, T. D. Bhatt, **M. S. Gill**, S. Suresh: Development and Evaluation of PEGylated Enoxaparin: a Novel Approach for Enhanced Anti-Xa Activity. *Bioorganic Chemistry*, 54, 1-6, 2014.
- 8. S. K. Jain, **M. S. Gill**, H. S. Pawar and S. Suresh: Novel Curcumin Diclofenac Conjugate Enhanced Curcumin Bioavailbility and Efficacy in Streptococcal Cel Wall-induced Arthritis. *Indian Journal of Pharmaceutical Sciences*, 415-422, 2014.
- 9. C.Singh; L. Jodave; Tara D Bhatt; **M.S. Gill**, S. Suresh. Hepatoprotective agent tethered isoniazid for the treatment of drug induced hepatotoxicity: Synthesis, biochemical and histopathological evaluation. *Toxicology Reports*, 1, 885-893, 2014.
- 10. V. Aggarwal, A. K. Angrish, A. Sharma and **M. S. Gill**: API Business in India: Opportunities and Threats. *Currents Research &Information on Pharmaceutical Sciences (CRIPS)*, Vol. 14, No. 4, 70-76, 2013.
- 11. B.M. Petiwala, and **M. S. Gill**: Dual Aromatse and Sulfatase Inhibitors (DAS(s): Structure and activity. *Currents Research &Information on Pharmaceutical Sciences (CRIPS)*, Vol. 13, No. 2, 29-33, 2012.
- 12. S. Suresh, L. Jodave, S. K. Jena, **M. S. Gill**: Antibody-Drug Conjugates: New Generation Therapeutics in Cancer Therapy. *Currents Research & Information on Pharmaceutical Sciences (CRIPS)*, Vol. 13, No. 3, 51-58, 2012.
- 13. Vivek Avasatthi, Harish Pawar, Chander Parkash, Prajeet Bansod, M. S. Gill, Sarasija Suresh. Development of Nanostructured Lipid Carrier of Methotrexate by Quality by Design Approach (*Submitted to European Journal of Pharmaceutical Sciences*) May, 2014.

## Patents

- 1. D. K. Tanwar, A. Ratan and **M. S. Gill**: "AN IMPROVED ONE POT PROCESS FOR SUBSTITUTED HYDANTOINS", Submitted for filing.
- 2. D. K. Tanwar, A. Ratan and M. S. Gill: "PROCESS FOR PREPARING SUBSTITUTED 1-SULFONYL HYDANTOINS", Submitted for filing.
- 3. D. K. Tanwar, N. Sarviya, R. P. Burman, S. Suresh and M. S. Gill: "A ONE POT PROCESS FOR SYNTHESIS OF PROPRANOLOL", Indian Patent Application No: 2643/DEL/2015, Filing date: August 26, 2015.
- 4. N. Sarviya, D. K. Tanwar, S. Suresh and **M. S. Gill**: "NOVEL PROCESS FOR SYNTHESIS OF PROPAFENONE", Indian Patent Application No: 2757/DEL/2015; Filing date: September 2, 2015.
- 5. D. K. Tanwar, A. Ratan, R. P. Burman, S. Suresh and M. S. Gill: "AN IMPROVED PROCESS FOR THE PREPARATION OF SULFONYLUREAS", Indian Patent Application No: 3386/DEL/2015; Filing date: October 10, 2015.
- 6. D. K. Tanwar, A. Ratan, R. P. Burman, S. Suresh and M. S. Gill: "IMPROVED PROCESS FOR PREPARATION OF UNSYMMETRICAL UREAS", Indian Patent Application No: 3387/DEL/2015; Filing date: October 10, 2015.
- 7. D. K. Tanwar, B. Deshmukh, P. Dileep kumar, R. P. Burman, S. Suresh and M. S. Gill: "PROCESS FOR THE PREPARATION OF SEMICARBAZIDES", Indian Patent Application No: 3385/DEL/2015; Filing date: October 10, 2015.
- 8. D. K. Tanwar, P. Dileep kumar, B. Deshmukh, R. P. Burman, S. Suresh and **M. S. Gill**: "PROCESS FOR THE PREPARATION OF MONOSUBSTITUTED UREAS", Indian Patent Application No: 3384/DEL/2015; Filing date: October 10, 2015.
- 9. A. Mital, A. Taylor and **M. S. Gill**: "A GREEN PROCESS FOR THE SYNTHESIS OF 2-IMINOTHIAZOLIDIN-4-ONE DERIVATIVES", Indian Patent Application No. 2594/DEL/2012, Filing date: August 22, 2012. Pub. No. 09/2014, Pub. Date Feburary 28, 2014.
- 10. S. K. Jain, S. Suresh, M. S. Gill: "NOVEL CURCUMIN-DRUG CONJUGATES", Indian Patent Application No. 731/DEL/2012, Filing date: March 14, 2012, Pub. No. 34/2015, Pub. Date August 21, 2015.
- 11. S. Suresh, **M. S. Gill**, C. Singh: "NOVEL NANO-FORMULATIONS OF ANTI-TUBERCULAR DRUGS", Indian Patent Application No: 1576/DEL/2013, Filing date: July 24, 2013, Pub. No. 37/2014, Pub. Date September 12, 2014.
- 12. D.I. Sherris, **M.S. Gill**, F. Ma, S.K. Bhal, R. Dunn-Default, M. Feher, P. Redden, J. Schmidt: "COMPOSITION AND METHODS TO TREAT SKIN DISEASE CHARACTERIZED BY CELLULAR PROLOFICATION AND ANIOGENSIS", PCT/US2006/040242; US11/412,618; US 14/171,217 Filed on Feburary 03, 2014; US 13/901,011 Filed on May 23, 2013.
- 13. S. Suresh, **M. S. Gill** and Charan Singh: NOVEL NANO-FORMULATIONS OF ANTI-TUBERCULAR DRUGS, Indian Patent, 1576/DEL/2013.

14. Alka Mital, Amit Taylor and **M. S. Gill**: A GREEN PROCESS FOR THE SYNTHESIS OF 2-IMINOTHIAZOLIDIN-4-ONE DERIVATIVES, Indian Patent, 2594/DEL/2012.

## Presentations

- 1. D. K. Tanwar, N. Sarviya, S. Suresh and M.S. Gill. Green Synthesis of Propafenon: An Anti-Arrhythmic Drug. National Conference on Emerging Trends in Biological and Chemical Sciences, 25-16 March, 2016, S. Govt. Sc. College of Education and Research, Jagraon, Ludhiana, Punjab, India.
- 2. **M.S. Gill**. API Synthesis and Process Chemistry. PTU-PITTTR Sponsored Faculty Development Program on "Current Scenario and Future Prospective in Pharmaceutical Research", 8-12 March, 2016, GHG Khalsa College of Pharmacy, Gurusar Sudhar, Punjab, India.
- 3. D. K. Tanwar, N. Sarviya, S. Suresh and M. S. Gill. Novel Process of Synthesis for Propafenone. Professor Ram Chand Paul National Symposium on Progressive Trends in Chemical Sciences, January 23, 2016, Panjab University, Chandigarh, India.
- 4. **M. S. Gill**. Process Chemistry Approaches to Synthesis of Drug Molecules. 9<sup>th</sup> Chandigarh Science Congress, 25-27 February, 2015, Panjab University, Chandigarh, India.
- 5. **M. S. Gill**. Drug Conjugates as Novel Therapeutics. 8<sup>th</sup> Chandigarh Science Congress, 26-28 February, 2014, Panjab University, Chandigarh, India.
- 6. **M. S. Gill**. Synthesis of Drug Conjugates. Prof. R.C. Paul National Symposium on New Visions in Chemical Sciences, 15-15 February, 2014, Panjab University, Chandigarh, India.
- 7. A. Mital, A. Tailor, V. S. Negi, Monika Jain, D. Ramya Sri and **M. S. Gill**. A Green Process for the Synthesis of 2-imino-thiazolidine-4-one Derivatives. DAV National Congress on Science and Technology, DAV Institute of Engineering and Technology, Jalandhar, Punjab, 30-31 May 2013.
- 8. S. K. Jain, Gurpreet Kaur, M. S. Gill and S. Suresh. Synthesis of Curcumin-Diclofenac Conjugate for increased curcumin bioavailability, DAV National Congress on Science and Technology, DAV Institute of Engineering and Technology, Jalandhar, India, 30-31 May 2013.