

ENGINEERING MATHEMATICS-III
(2019 Pattern) (Semester-IV) (207003)

(Max. Marks : 70)

Time : 75 Minutes

Instructions to the Candidates:

- 1) Q. No. 1 is compulsory.
- 2) Answers Q. No. 2, 3, 4 or Q. 5, 6 or Q. 7, 8 or Q. 9.
- 3) Your solutions must be drawn wherever necessary.
- 4) Figures to the right indicate full marks.
- 5) Use of electronic pocket calculator is allowed.
- 6) Assume suitable data, if necessary.

Q.1) Choose the correct option of the following.

a) If the first four central moments of a distribution are 0, 2.5, 0.7 and 18.75 then the coefficient of Kurtosis β_1 is _____ [2]

- a) 0
- b) 1
- c) 2
- d) 3

b) The probability distribution of x is

x	1	2	3	4
P(x)	1/2	1/4	1/8	1/8

The mathematical expectation E(x) is _____

- a) 13/8
- b) 11/8
- c) 9/8
- d) 15/8

c) A root of the equation $x^2 - 4x - 6 = 0$ lies between _____ [2]

- a) 1 and 2
- b) 2 and 4
- c) 0 and 1
- d) 2 and 3

P.T.O.

M3 Question Paper June 2013

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Drug Discovery and Evaluation: Safety and Pharmacokinetic Assays Franz J. Hock, Michael K. Pugsley, 2024-10-21

Many aspects of drug safety have become an outstanding and even persistent issue and may occur during the process of both drug discovery and development. Until 15 years ago, drug discovery and evaluation was primarily a sequential process starting with the selection of the most pharmacologically active compound from a series of newly synthesized small molecule chemical series by means of distinctive pharmacological assays. Safety aspects were addressed by evaluation of the selected compound at high doses in a series of specific studies directed at indications other than the intended indication of the new compound. These tests are then followed by pharmacokinetic studies which are primarily conducted to confirm whether the selected compound possesses a suitable half life for sufficient exposure and efficacy and whether it has the desired properties specific to the intended route of administration. Safety aspects relied predominantly on the conduct of single and repeat toxicology dose studies which inform changes in organ structure rather than organ function. Both toxicological and pharmacokinetic studies are adapted to the progress of studies in clinical pharmacology and clinical trials. The new edition of this well and broadly accepted reference work contains several innovative and distinguished chapters. This sequential strategy has been abandoned with this new version of the book for several reasons. Of the possible multitude of negative effects that novel drugs may impart on organ function, e.g. ventricular tachy arrhythmia, many are detected too late in non-clinical studies to inform clinicians. On the other hand, negative findings in chronic toxicity studies in animals may turn out to be irrelevant for human beings. New scientific approaches, e.g. high throughput screening, human pluripotent stem cells, transgenic animals, knock out animals, in silico models, pharmacogenomics and pharmacoproteomics, as well as Artificial Intelligence (AI) methods, offered new possibilities. There are several examples that show that the druggability of compounds was considerably underestimated when the probability of success of a new project was assessed. The success rate in the pharmaceutical industry and the introduction of new chemical entities to the market per year dropped dramatically, whereas the development time for a new compound increased, sometimes exceeding the patent protection. Research and development scientists, involving the following changes, therefore adopted a change of strategy: Parallel instead of sequential involvement of the various disciplines, multidimensional compound optimization. The term Safety Pharmacology was coined. The International Conference on Harmonization (ICH) founded a Safety Pharmacology Working Group and the Safety Pharmacology Society (SPS) was launched. The discipline provided for evaluation, development and validation of a multitude of safety tests outlined in the Core Battery of Studies. Characterizing the exposure profile of a drug by conducting pharmacokinetic studies that evaluates the absorption, distribution, metabolism and excretion should to be investigated at an early stage of development as results contribute to the selection of a compound for further development. Advancements in Toxicology were achieved by the introduction of new methods, e.g. in silico methods, genetic toxicology, computational

toxicology and AI The book is a landmark in the continuously changing world of drug research and developments As such it is essential reading for many groups not only for all students of pharmacology and toxicology but also for industry scientists and physicians especially those involved in clinical trials of drugs and for pharmacists who must know the safety requirements of drugs The book is essential for scientists and managers in the pharmaceutical industry who are involved in drug discovery drug development and decision making in the development process In particular the book will be of use to government institutions and committees working on official guidelines for drug evaluation worldwide

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implications on marginalized peoples at home and abroad Included are political economy approaches that draw on the work of theorists such as David Harvey important interventions from Indigenous and settler colonial studies feminist approaches using the work of scholars such as Silvia Federici and the concept environmental racism which draws on both critical race theory and environmental justice literature

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