Clinical Research Made Easy

Clinical Research Made Easy

A Guide to Publishing in Medical Literature

Editors

Mohit Bhandari MD MSc FRCSC Department of Surgery Division of Orthopaedic Surgery McMaster University 293 Wellington Street North, Suite 110 Hamilton, Ontario, L8L 8E7 Canada

Parag Sancheti MS(Ortho) DNB(Ortho) MCh(Ortho) UK Sancheti Institute for Orthopaedics and Rehabilitation 16 Shivajinagar Pune 411005, India

Managing Editors

Farrah Hussain BSc (Cand) Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North, Suite 110 Hamilton, Ontario, L8L 8E7 Canada

Sheila Sprague MSc Department of Clinical Epidemiology and Biostatistics McMaster University 293 Wellington Street North, Suite 110 Hamilton, Ontario, L8L 8E7 Canada

Foreword

KH Sancheti



JAYPEE BROTHERS MEDICAL PUBLISHERS (P) LTD

Mumbai • New Delhi • St Louis (USA) • Panama City (Panama) • London (UK) • Ahmedabad Bengaluru • Chennai • Hyderabad • Kochi • Kolkata • Lucknow • Nagpur

Published by Jitendar P Vij Jaypee Brothers Medical Publishers (P) Ltd

Corporate Office

4838/24 Ansari Road, Daryaganj, **New Delhi** - 110002, India Phone: +91-11-43574357, Fax: +91-11-43574314

Registered Office

B-3 EMCA House, 23/23B Ansari Road, Daryaganj, **New Delhi** - 110 002, India Phones: +91-11-23272143, +91-11-23272703, +91-11-23282021 +91-11-23245672, Rel: +91-11-32558559, Fax: +91-11-23276490, +91-11-23245683 e-mail: jaypee@jaypeebrothers.com, Website: www.jaypeebrothers.com

Offices in India

- Ahmedabad, Phone: Rel: +91-79-32988717, e-mail: ahmedabad@jaypeebrothers.com
- Bengaluru, Phone: Rel: +91-80-32714073, e-mail: bangalore@jaypeebrothers.com
- Chennai, Phone: Rel: +91-44-32972089, e-mail: chennai@jaypeebrothers.com
- Hyderabad, Phone: Rel:+91-40-32940929, e-mail: hyderabad@jaypeebrothers.com
- Kochi, Phone: +91-484-2395740, e-mail: kochi@jaypeebrothers.com
- · Kolkata, Phone: +91-33-22276415, e-mail: kolkata@jaypeebrothers.com
- Lucknow, Phone: +91-522-3040554, e-mail: lucknow@jaypeebrothers.com
- Mumbai, Phone: Rel: +91-22-32926896, e-mail: mumbai@jaypeebrothers.com
- Nagpur, Phone: Rel: +91-712-3245220, e-mail: nagpur@jaypeebrothers.com

Overseas Offices

- North America Office, USA, Ph: 001-636-6279734
 e-mail: jaypee@jaypeebrothers.com, anjulav@jaypeebrothers.com
- Central America Office, Panama City, Panama, Ph: 001-507-317-0160
 e-mail: cservice@jphmedical.com, Website: www.jphmedical.com
- Europe Office, UK, Ph: +44 (0) 2031708910 e-mail: dholman@ipmedical.biz

Clinical Research Made Easy—A Guide to Publishing in Medical Literature

© 2010, Jaypee Brothers Medical Publishers

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

This book has been published in good faith that the material provided by contributors is original. Every effort is made to ensure accuracy of material, but the publisher, printer and editors will not be held responsible for any inadvertent error (s). In case of any dispute, all legal matters are to be settled under Delhi jurisdiction only.

First Edition: 2010

ISBN 978-81-8448-890-6

Typeset at JPBMP typesetting unit

Printed at

Dedicated to

our Parents Their endless love and support have been a constant source of inspiration to us

our Patients Without their cooperation we would not have been able to improve health care standards through evidence-based medicine

Foreword

I am very glad and congratulate the authors for writing such an informative book on a subject so close to my heart. In my opinion this book will be able to satisfy the needs of every clinician, medical student and clinical research personnel who is interested in conducting and publishing a scientific research.

My clinical practice has now spanned over decades and my keen interest in clinical research has helped me to keep a finger on the pulse of the research scenario. The last few decades have seen development of a more scientific and evidence-based approach with an inclination for collaboration and cooperation towards development of a better health care system. In India too there has been a change, however lack of infrastructure and basic education in research are a major obstruction for us. This book, even though it is guite concise, will fill this lacuna in research education to a great extent. The precise writing and accurate and practical approach that the authors were able to maintain throughout is the most unique point of this book. The simple flow of explaining the complex ideas and methods goes to indicate the excellent grasp of the authors on the topic and keeps the reader engrossed.

I am sure every clinician and medical student interested in organizing a scientific study and publishing its result will find this book extremely helpful. This book

viii

will also be a landmark in the developing clinical research scenario in India.

I wish the authors all the success.

KH Sancheti

MBBS FCPS D(Ortho) MS(Ortho) FICS FACS PhD FRCS Founder-Sancheti Institute for Orthopaedics and Rehabilitation Pune, India

Preface

Since its inception approximately 20 years ago, evidencebased medicine (EBM) has become a standard part of many educational curricula world-wide. In his original document to trainees at McMaster University, Professor Gordon Guyatt described EBM as an attitude of 'enlightened scepticism' toward the application of diagnostic, therapeutic, and prognostic technologies in their day-to-day management of patients." The concepts of EBM empower us to formulate appropriate clinical questions, appraise the literature using the hierarchy of evidence, and apply the study results to their practice. The application of evidence-based medicine is not easy. It requires a fundamental understanding of research methodology and critical appraisal. Critical concepts for health care providers include the strengths and limitations of meta-analyses, randomized trials, cohort studies and case series.

Although we may perceive that evidence-based medicine mandates a strict adherence to randomized trials, it more accurately involves informed and effective use of all types of evidence (from meta-analysis of randomized trials to individual case series and case reports). With the ever-increasing amount of available information, we must consider a shift in paradigm from traditional practice to one that involves question formulation, validity assessment of available studies and appropriate application of research evidence to individual patients.

CLINICAL RESEARCH MADE EASY

X

For those of us wishing to produce high quality evidence to guide patient care, the good research practice is paramount. From beginning to end, the development of a quality study protocol, an appropriate statistical evaluation of sample size, the use of a valid outcome measure and sound ethical practices are basic requirements.

With the ever-increasing demands to adopt evidencebased medicine in practice, health care providers require educational resources that present the concepts of the EBM, research methodology and guides to publishing medical research in a simple and easy to understand format.

Clinical Research Made Easy is one resource developed with the busy health care provider in mind. We have assembled a group of leading researchers, surgeons and physicians with the aim of providing high quality content to our readers. Whether a practising surgeon, physician, physiotherapist, student or allied health care provider, this book bridges your knowledge gaps and facilitates practical application of EBM and clinical research. Led by practising physicians and researchers, we understand the challenges in research and provide simple guides for success. We distil complexities of clinical research into simple facts. Our question and answer format uses commonly asked questions in each topic area and provides focused summaries.

In an emerging era of innovation in medicine and health care we have a responsibility to engage in the process of research acquisition, conduct and dissemination. We should no longer consider clinical

PREFACE

research as only occurring in high profile institutions around the world, but rather occurring in every health care setting in all environments. Clinical research must be made easy to become widely accessible and widely applied.

We hope this book provides one step in the right direction—and stimulates a new generation of evidence users and evidence providers!

Mohit Bhandari Parag Sancheti

xi

Contributors

Aleem Adatia BHSc(Cand) Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada adatia2@mcmaster.ca

Anil K Jain MS MAMS Professor of Orthopaedics University College of Medical Sciences University of Delhi, India Editor: Indian Journal of Orthopaedics. *dranilkjain@gmail.com*

Ashok Shyam MS (Ortho) Sancheti Institute for Orthopaedics and Rehabilitation 16 Shivajinagar, Pune 411005, India *drashokshyam@yahoo.co.uk*

AV Gurava Reddy

MB D.orth DNB orth MCh(Ortho) FRCS(Ed) FRCS(Gl) Managing Director and CEO Chief of Division of Joint Replacement Surgery Sunshine Hospitals, PG road Secunderabad 500003 guravareddy@gmail.com

Christopher Vannabouathong BSc

Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada trauma2@mcmaster.ca

CJ Thakkar MS(Ortho) DNB Hon. Professor of Orthopaedics, Maharashtra University of Health Science LTM Municipal Medical College and Hospital, Sion Mumbai Breach Candy and Lilavati Hospitals, Mumbai *cjt@rediffmail.com*

Farrah Hussain BSc(Cand) Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada farrah.hussain@utoronto.ca

Fatima Zaidi BSc(Cand) Orthopaedic Research Unit Department of Surgery McMaster University

CLINICAL RESEARCH MADE EASY

GS Kulkarni

MS(Gen) MS(Orth) FRCS Director, P G Institute of Swastiyog Pratishthan Meeraj 416410 gskorth@gmail.com

Harman Chaudhry

BHSc MD(Cand) Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada harman.chaudhry@medportal.ca

Hiba Abdul Mannan

BSc(Cand) Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada hibamannan@hotmail.com

JA Pachore MCh Director – HIP Surgery Shalby Hospital Ahmedabad

Kevin Debiparshad

BSc MSc MD Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada *kevin.debiparshad@medportal.ca*

Laura Quigley

Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada *laura_quigley@hotmail.com*

Mandeep Dhillon

MBBS MS(Ortho) MNAMS Professor and Head Dept of Orthopaedics Head Deptt of Physical Medicine and Rehabilitation, Incharge, Sports Medicine Clinic Post Graduate Institute of Medical Education and Research Chandigarh *drdhillon@gmail.com*

Meaghan Zehr BSc(Cand)

Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada *meagasis@hotmail.com*



CONTRIBUTORS

Mohit Bhandari

MD MSc FRCSC Department of Surgery Division of Orthopaedic Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada bhandam@mcmaster.ca

Nasir Hussain BSc(Cand)

Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada *nasir.hussain@utoronto.ca*

Natalie Kuurstra BA

Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada *trauma6@mcmaster.ca*

Neil Dinesh Dattani

BHSc(Cand) Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North, Suite 110 Hamilton, Ontario, L8L 8E7 Canada *dattann@muss.cis.mcmaster.ca*

Parag Sancheti

MS(ORTHO) DNB(ORTHO) MCh(ORTHO) UK Sancheti Institute for Orthopaedics and Rehabilitation 16 Shivajinagar, Pune 411005, India parag@sanchetihospital.org

Raman Mundi BHSc MD(Cand)

Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada *raman.r.mundi@learnlink. mcmaster.ca*

Roopinder Kaloty BHSc

Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada roopinder.kaloty@learnlink. mcmaster.ca

S Rajasekaran MS DNB

FRCS(Ed) MCh(Liverpool) FACS PhD Ganga Hospital 313 Mettupalayam Road Coimbatore: 641043 rajasekaran.orth@gmail.com

Sarah Resendes BSc(Cand) Orthopaedic Research Unit Department of Surgery

XV

CLINICAL RESEARCH MADE EASY

McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada sjresend@ahsmail.uwaterloo.ca

Sharan Shivaraj Patil

Chief Orthopaedic Surgeon D'Orth MS(Orth), MCh(Orth) Liverpool, UK Sparsh Hospital, No. 29/P2 Narayana Health City, Hosur Road, Bangalore-560099 *drsharanpatil@ sparshhospital.com*

Sheila Sprague MSc

Department of Clinical Epidemiology & Biostatistics McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada sprags@mcmaster.ca

Simrit Bains BSc MA(Cand) Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada s.bains3@gmail.com

SKS Marya

MS DNB MCh(UK) FICS Chairman and Chief Surgeon Max Institute of Orthopaedics and Joint Replacement 1, Press Enclave Road, Saket New Delhi – 110 017 smarya@maxhealthcare.com

Steve Rocha MS (ORTHO)

Sancheti Institute for Orthopaedics and Rehabilitation 16 Shivajinagar, Pune 411005, India *srgnrocha@gmail.com*

Tina Le Francois MD

Orthopaedic Research Unit Department of Surgery McMaster University 293 Wellington Street North Suite 110 Hamilton, Ontario, L8L 8E7 Canada *tlefrancois2007@meds.uwo.ca*

Vijay D Shetty MS(Orth) Fellow, Cambridge Hip and Knee Unit, Cambridge (UK) Hiranandani Orthopaedic Medical Education (HOME) Dr LH Hiranandani Hospital Powai,Mumbai 400 076 India vijaydshetty@gmail.com

xvi

Contents

	Section I: Evidence-based Medicine
1.	The Era of Evidence-based Medicine
	Nasir Hussain, Mohit Bhandari, Parag Sancheti
2.	Evidence-based Medicine and the Physician 11
	Nasir Hussain, Parag Sancheti
3.	The Hierarchy of Evidence
	Nasir Hussain, Ashok Shyam
4.	The Current Quality of Medical Literature
	Fatima Zaidi, Anil Jain
	Section II: Principles of Clinical Research
5.	Critical Appraisal Made Simple41
	Farrah Hussain, Nasir Hussain, Steve Rocha
6.	Level 1 Evidence: Randomized Controlled Trials45
	Hiba Abdul Mannan, Parag Sancheti
7.	Level 2 Evidence: Prospective Cohort Studies55
	Aleem Adatia, Mohit Bhandari, Gurava Reddy
8.	Level 3 Evidence: Case-control Studies68
	Harman Chaudhry, Mandeep Dhillon
9.	Level 4 Evidence: Clinical Case Series80
	Fatima Zaidi, Vijay Shetty
10.	Level 5 Evidence: Surveys
	Roopinder Kaloty, SKS Marya
11.	The Basics of Literature Reviews and Meta-analyses . 102
	Neil Dinesh Dattani, Govind Kulkarni
12.	The Basics of Economic Analysis 118
	Neil Dinesh Dattani, CJ Thakker
13.	The Basics of a Diagnostic Test Study 130
	Raman Mundi, Steve Rocha
14.	The Basics of a Reliability Study 143
	Laura Quigley, J Pachore

xviii

CLINICAL RESEARCH MADE EASY

Section	Ш÷	Practice	of	Clinical	Research
00001011		1 I douice	01	Unincar	Research

15.	Preparing a Clinical Research Proposal	7
16.	Understanding Outcomes Assessment 168 Natalie Kuurstra, Anil Jain	3
17.	Everything You Need to Know About Sample Size Calculations178 Chris Vannabouathong, Mohit Bhandari	3
18.	Preparing a Clinical Research Budget	7
19.	Research Ethics and Consent Forms	3
20.	Conducting a Randomized Controlled Trial	l
21.	How to Conduct a Cohort Study	l
22.	Conducting a Case Series 233 Meaghan Zehr, Vijay Shetty	3
23.	Common Pitfalls in Conducting a Study	5
	Section IV: Dissemination of Clinical Research	
24.	Review of Basic Statistical Principles	•
25.	Preparing a Clinical Research Presentation 273 Kevin Debiparshad, Parag Sancheti	3
26.	Preparing an Abstract for a Clinical Meeting 283 Tina Le Francois, Mandeep Dhillon	3
27.	Writing a Good Clinical Research Paper 293 Sarah Resendes, Ashok Shyam	3
28.	The Future of Clinical Research304Farrah Hussain, Sheila Sprague, Anil Jain	4
Glo	ssary	1
Inde	əx	7