

**More Information** 

# Manual of Sperm Retrieval and Preparation in Human Assisted Reproduction



More Information

# Cambridge Laboratory Manuals in Assisted Reproductive Technology

#### Titles in the series:

Manual of Oocyte Retrieval and Preparation in Human Assisted Reproduction, Rachel Cutting & Mostafa Metwally

Manual of Sperm Function Testing in Human Assisted Reproduction, Ashok Agarwal, Ralf Henkel & Ahmad Majzoub

Manual of Intracytoplasmic Sperm Injection in Human Assisted Reproduction: With Other Advanced Micromanipulation Techniques to Edit the Genetic and Cytoplasmic Content of the Oocyte, Gianpiero D. Palermo & Peter Nagy

Manual of Embryo Selection in Human Assisted Reproduction, Catherine Racowsky, Jacques Cohen & Nick Macklon

Manual of Embryo Culture in Human Assisted Reproduction, Kersti Lundin & Aisling Ahlström Manual of Cryopreservation in Human Assisted Reproduction, Sally Catt, Kiri Beilby & Denny Sakkas



Manual of Sperm Retrieval and Preparation in Human Assisted Reproduction

Edited by

#### Ashok Agarwal

The Cleveland Clinic Foundation, Cleveland, OH, USA

#### **Ahmad Majzoub**

Hamad Medical Corporation, Doha, Qatar

#### Sandro C. Esteves

ANDROFERT - Andrology & Human Reproduction Clinic, and University of Campinas (UNICAMP), Campinas, Brazil





# **CAMBRIDGE**UNIVERSITY PRESS

University Printing House, Cambridge CB2 8BS, United Kingdom One Liberty Plaza, 20th Floor, New York, NY 10006, USA 477 Williamstown Road, Port Melbourne, VIC 3207, Australia 314–321, 3rd Floor, Plot 3, Splendor Forum, Jasola District Centre, New Delhi – 110025. India

79 Anson Road, #06-04/06, Singapore 079906

Cambridge University Press is part of the University of Cambridge.

It furthers the University's mission by disseminating knowledge in the pursuit of education, learning, and research at the highest international levels of excellence.

www.cambridge.org Information on this title: www.cambridge.org/9781108792158 DOI: 10.1017/9781108867245

© Cambridge University Press 2021

This publication is in copyright. Subject to statutory exception and to the provisions of relevant collective licensing agreements, no reproduction of any part may take place without the written permission of Cambridge University Press.

First published 2021

Printed in Singapore by Markono Print Media Pte Ltd

A catalogue record for this publication is available from the British Library.

ISBN 978-1-108-79215-8 Paperback

Cambridge University Press has no responsibility for the persistence or accuracy of URLs for external or third-party internet websites referred to in this publication and does not guarantee that any content on such websites is, or will remain, accurate or appropriate.

Every effort has been made in preparing this book to provide accurate and up-to-date information that is in accord with accepted standards and practice at the time of publication. Although case histories are drawn from actual cases, every effort has been made to disguise the identities of the individuals involved. Nevertheless, the authors, editors, and publishers can make no warranties that the information contained herein is totally free from error, not least because clinical standards are constantly changing through research and regulation. The authors, editors, and publishers therefore disclaim all liability for direct or consequential damages resulting from the use of material contained in this book. Readers are strongly advised to pay careful attention to information provided by the manufacturer of any drugs or equipment that they plan to use.



To my father Professor RC Agarwal (late) for instilling the virtues of honesty, dedication, and hard work. To my wonderful wife, Meenu, sons, Rishi and Neil-Yogi, for their unconditional love and support. To Professor Kevin Loughlin (Harvard Medical School), Professor Anthony Thomas (late) (Cleveland Clinic), and Professor Edmund Sabanegh (Cleveland Clinic) for their friendship, guidance, and support, and for making an indelible positive impression on my life. To my associates at work, large number of researchers and students, and most importantly the patients who placed their trust in our work.

Ashok Agarwal

I would like to dedicate this book to my wife, Zeinab, and our two lovely daughters, Sarah and Tala, for their love and support; to my mentors, Dr. Edmund Sabanegh and Professor Ashok Agarwal (Cleveland Clinic), for their guidance and support and for all the opportunities they made possible.

Ahmad Majzoub

To my father, Waldemar Esteves (late), for instilling the virtues of integrity, perseverance, and enthusiasm. To my wife, Fabiola, my sons, Alexandre and Catarina, for their devotion, love, and support. To my team at ANDROFERT Clinic for their unconditional support and hard work. To my mentors Professor Nelson Rodrigues Netto Jr. (late) (UNICAMP), Dr. Anthony Thomas Jr. (late) (Cleveland Clinic), Professor Agnaldo Cedenho (Federal University of São Paulo), and Dr. Ashok Agarwal (Cleveland Clinic) for their guidance, support, and for being giants who take their apprentices by their hands and teach them how to climb; once on their shoulders, you have shown us what to see . . . helping us to see better and farther. To my colleague Dr. Ahmad Majzoub for sharing the enthusiasm in the study and practice of andrology and male infertility. And lastly, to all my patients who put their trust in my work and from whom I learn more and more daily.

Sandro C. Esteves

V



# **Contents**

List of Contributors viii Editor Biographies x Preface xi

#### Part I Introduction

- 1 Anatomy and Physiology of the Male Reproductive System 1 Ryan T. Scribner and Nicholas N. Tadros
- Evaluation of Candidates for
   Sperm Retrieval 9
   Darren J. Bryk, Neel V. Parekh, and Sarah C. Vij
- Testicular Histopathology and the Role of Testis Biopsy 16
   Gianmartin Cito

### Part II Sperm Retrieval

- 4 History of Surgical Sperm Retrieval Techniques 20 Rupin Shah
- 5 Epididymal Sperm Retrieval: Indications,
   Surgical Protocol, and Outcomes 25
   Jason P. Akerman and R. Matthew Coward
- 6 Testicular Sperm Retrieval: Indications, Surgical Protocol, and Outcomes 36 Ahmad Majzoub, Chak Lam Cho, and Sandro C. Esteves
- 7 Surgical and Nonsurgical Sperm Retrieval Techniques in Patients with Ejaculatory Dysfunctions 44 Omer Raheem, Mohamed Adnan, and Laith M. Alzweri
- Sperm Retrieval in Non-azoospermic Men:
   Indications, Protocol, and Outcomes 56
   Sandro C. Esteves and Ahmad Majzoub

- 9 Predictors of Positive Surgical Sperm Retrieval in Azoospermic Males: Evaluation of Clinical, Laboratory, and Histopathologic Factors 75 Kareim Khalafalla, Ahmed H. Almalki, Ahmad Majzoub, and Sandro C. Esteves
- 10 Methods for Enhancing Surgical Sperm Retrieval Success 86Jordan A. Cohen and Ranjith Ramasamy

# Part III Laboratory Handling of Retrieved Sperm

- 11 **Critical Factors for Optimizing Sperm Handling and ICSI Outcomes** 90
  Edson Borges Jr., Amanda Souza Setti, and Daniela Paes de Almeida Ferreira Braga
- 12 **Sperm Cryopreservation** 99 Rakesh Sharma, Kruyanshi Master, and Ashok Agarwal
- Future Directions in Sperm Handling for ART 117
   Catalina Barbarosie, Manesh Kumar Panner Selvam, and Ashok Agarwal

Indox	- 11	31

vii



# **Contributors**

#### **Mohamed Adnan**

Department of Urology, Tulane University, New Orleans, Louisiana, USA

#### Daniela Paes de Almeida Ferreira Braga

Fertility Medical Group – São Paulo, Brazil

Sapientiae Insitute - São Paulo, Brazil

#### **Ashok Agarwal**

Andrology Center, Department of Urology, American Center for Reproductive Medicine, Cleveland Clinic, Cleveland, OH, USA

#### Jason P. Akerman

University of North Carolina, Department of Urology, Chapel Hill, NC, USA

#### Ahmed H. Almalki

Urology Department, Hamad Medical Corporation, Doha, Qatar

#### Laith M. Alzweri

Department of Urology, University of Texas Medical branch, Galveston Texas, USA

#### Catalina Barbarosie

American Center for Reproductive Medicine, Cleveland Clinic, Cleveland, OH, USA Faculty of Biology, University of Bucharest, Bucharest, Romania

#### **Edson Borges**

Fertility Medical Group – São Paulo, Brazil Sapientiae Insitute – São Paulo, Brazil

#### Darren J. Bryk

Department of Urology, Glickman Urological and Kidney Institute, Cleveland Clinic Foundation, Cleveland, OH, USA

#### Chak-Lam Cho

Department of Surgery, Union Hospital, Shatin, Hong Kong

#### **Gianmartin Cito**

Department of Urology, Careggi Hospital, University of Florence, Florence, Italy

#### Jordan A. Cohen

Department of Urology, University of Miami, Miami, FL, USA

#### R. Matthew Coward

UNC Fertility, Raleigh, NC, USA

#### Sandro C. Esteves

ANDROFERT, Andrology & Human Reproduction Clinic, Campinas, Brazil Department of Surgery (Division of Urology),

University of Campinas (UNICAMP), Campinas, Brazil

Faculty of Health, Department of Clinical Sciences, Aarhus University, Aarhus, Denmark

#### Kareim Khalafalla

Urology Department, Hamad Medical corporation, Doha, Qatar

#### Manesh Kumar Panner Selvam

American Center for Reproductive Medicine, Cleveland Clinic, Cleveland, OH, USA

#### Neel V. Parekh

Department of Urology, Glickman Urological and Kidney Institute, Cleveland Clinic Foundation, Cleveland, OH, USA

#### Ahmad Majzoub

Department of Urology, Hamad Medical Corporation, Doha, Qatar Weill Cornell Medicine – Qatar, Doha, Qatar

viii



**More Information** 

#### List of Contributors

#### Kruyanshi Master

American Center for Reproductive Medicine, Cleveland Clinic, Cleveland, OH, USA

#### **Omer Raheem**

Department of Urology, Tulane University, New Orleans, Louisiana, USA

#### Ranjith Ramasamy

Department of Urology, University of Miami, FL, USA

#### Ryan Scribner

Memorial Medical Center, Springfield, IL, USA

#### **Rupin Shah**

Lilavati Hospital & Research Centre, Mumbai, India

#### Rakesh Sharma

American Center for Reproductive Medicine, Cleveland Clinic, Cleveland, OH, USA

#### Amanda Souza Setti

Fertility Medical Group – São Paulo, Brazil Sapientiae Insitute – São Paulo, Brazil

#### Nick N. Tadros

Department of Surgery, SIU School of Medicine, Springfield, IL, USA

#### Sarah C. Vij

Department of Urology, Glickman Urological and Kidney Institute, Cleveland Clinic Foundation, Cleveland, OH, USA



# **Editor Biographies**

Dr. Ashok Agarwal is the Head of Andrology Center and Director of Research at the American Center for Reproductive Medicine. He holds these positions at the Cleveland Clinic Foundation, where he is Professor of Surgery (Urology) at the Lerner College of Medicine of Case Western Reserve University. Ashok was trained in male infertility and andrology at the Brigham and Women's Hospital and Harvard Medical School, and has over 28 years of experience in directing busy male infertility diagnostic and fertility preservation services. He is an editor of over 40 medical textbooks on male infertility, ART, fertility preservation, DNA damage, and antioxidants.

Dr. Majzoub is a consultant at the Department of Urology and Program Director of the Andrology and Male Infertility Fellowship at Hamad Medical Corporation, Doha, Qatar. He is an Arab Board Certified Urologist and has undergone Clinical and Research Fellowship in Andrology at the world-famous Glickman Urological and Kidney Institute and the American Center for Reproductive Medicine at Cleveland Clinic Foundation, Cleveland, USA.

Dr. Majzoub is actively involved in the field of medical research, with over 150 research publications in peer-reviewed journals and several book chapters mainly focusing on andrology and men's health. He has also co-edited several books and special issues on various aspects of male infertility.

Dr. Sandro C. Esteves is Medical Director of ANDROFERT, a referral fertility center for male reproduction in Brazil. Dr. Esteves is also Collaborating Professor in the Department of Surgery (Division of Urology) at the University of Campinas (UNICAMP, Brazil) and Honorary Professor of Reproductive Endocrinology at the Faculty of Health, Aarhus University, Denmark. Professor Esteves has published over 300 peerreviewed scientific papers, authored over 80 book chapters, and served as an editor of 10 textbooks related to male infertility and assisted reproductive technology. He serves on the editorial board of several journals and is Associate Editor of International Brazilian Journal of Urology and Frontiers in Endocrinology (Reproduction).



# **Preface**

The field of male infertility has witnessed major clinical advancements in recent years, and perhaps the most important of these was the development of testicular sperm retrieval procedures that allowed men with azoospermia to father their biological children. Epididymal sperm retrieval procedures were first performed in the 1980s for men with obstructive azoospermia. The realization that men with nonobstructive azoospermia may indeed have focal areas of testicular sperm production together with the documented fertilizing ability of testicular spermatozoa allowed the development of testicular sperm retrieval procedures in the 1990s. Subsequently, testicular sperm retrieval underwent further refinement with the introduction of microsurgery, which improved the sperm retrieval rate and at the same time reduced the potential adverse impact of surgery on testicular parenchyma. Extensive research has been conducted in attempts to study the predictors of positive sperm retrieval, hoping to increase the outcome of surgical sperm retrieval procedures.

This manual presents recent advancements in the surgical management of azoospermia patients. It is divided into three parts: Part I serves as an introduction presenting important anatomic and physiologic aspects of the reproductive tract and demonstrating the ideal methods for evaluating candidates of sperm retrieval. Part II elaborates on the surgical techniques of sperm retrieval in a variety of clinical scenarios.

Moreover, it investigates the predictors of successful sperm retrieval and explores methods for enhancing sperm retrieval outcomes. Finally, Part III focuses on the laboratory handling of retrieved sperm and sperm cryopreservation, and explores future directions aimed at optimizing embryologists' work in the lab.

We are confident that our book will be a useful guide for reproductive surgeons, IVF specialists, embryologists, and other healthcare workers practicing reproductive medicine. In addition, it will be a valuable resource for students and researchers wishing to learn more about this subject. We are greatly thankful to large number of experts who worked hard to contribute the latest, well written, and well researched articles; this book would not be possible without their active support. We wish to express our deep gratitude to the superb organizational and management skills of Camille Lee-Own, publishing assistant at Cambridge University Press, and the overall support and supervision of this project by Nick Dunton, publisher at Cambridge University Press. This book is dedicated to our parents, families, mentors, and patients.

#### **Editors:**

Professor Ashok Agarwal, Cleveland Clinic, USA Dr. Ahmad Majzoub, Hamad Medical Corporation, Oatar

Professor Sandro Esteves, Androfert, Brazil