



June 2001

C31-A2

Ionized Calcium Determinations: Precollection Variables, Specimen Choice, Collection, and Handling; Approved Guideline—Second Edition



This document addresses preanalytical considerations, such as patient condition, specimen choice, collection, and handling—that can influence the accuracy and clinical utility of ionized calcium measurements.

A guideline for global application developed through the Clinical and Laboratory Standards Institute consensus process.

Clinical and Laboratory Standards Institute

Setting the standard for quality in clinical laboratory testing around the world.

The Clinical and Laboratory Standards Institute (CLSI) is a not-for-profit membership organization that brings together the varied perspectives and expertise of the worldwide laboratory community for the advancement of a common cause: to foster excellence in laboratory medicine by developing and implementing clinical laboratory standards and guidelines that help laboratories fulfill their responsibilities with efficiency, effectiveness, and global applicability.

Consensus Process

Consensus—the substantial agreement by materially affected, competent, and interested parties—is core to the development of all CLSI documents. It does not always connote unanimous agreement, but does mean that the participants in the development of a consensus document have considered and resolved all relevant objections and accept the resulting agreement.

Commenting on Documents

CLSI documents undergo periodic evaluation and modification to keep pace with advancements in technologies, procedures, methods, and protocols affecting the laboratory or health care.

CLSI's consensus process depends on experts who volunteer to serve as contributing authors and/or as participants in the reviewing and commenting process. At the end of each comment period, the committee that developed the document is obligated to review all comments, respond in writing to all substantive comments, and revise the draft document as appropriate.

Comments on published CLSI documents are equally essential, and may be submitted by anyone, at any time, on any document. All comments are addressed according to the consensus process by a committee of experts.

Appeals Process

If it is believed that an objection has not been adequately addressed, the process for appeals is documented in the CLSI Administrative Procedures.

All comments and responses submitted on draft and published documents are retained on file at CLSI and are available upon request.

Get Involved—Volunteer!

Do you use CLSI documents in your workplace? Do you see room for improvement? Would you like to get involved in the revision process? Or maybe you see a need to develop a new document for an emerging technology? CLSI wants to hear from you. We are always looking for volunteers. By donating your time and talents to improve the standards that affect your own work, you will play an active role in improving public health across the globe.

For further information on committee participation or to submit comments, contact CLSI.

Clinical and Laboratory Standards Institute
950 West Valley Road, Suite 2500
Wayne, PA 19087 USA
P: 610.688.0100
F: 610.688.0700
www.clsi.org
standard@clsi.org

NOTE: This document is no longer being reviewed as part of the CLSI consensus process. However, because of its usefulness to segments of the health care community, it is available for its informational content.

ISBN 1-56238-436-8
ISSN 0273-3099

C31-A2
Vol. 21 No. 10
Replaces C31-A
Vol. 15 No. 20

Ionized Calcium Determinations: Precollection Variables, Specimen Choice, Collection, and Handling; Approved Guideline— Second Edition

Volume 21 Number 10

Paul D’Orazio, Ph.D.
John G. Toffaletti, Ph.D.
Jesper Wandrup, M.D., Ph.D.

Abstract

Ionized Calcium Determinations: Precollection Variables, Specimen Choice, Collection, and Handling; Approved Guideline—Second Edition (CLSI document C31-A2) is a guideline for specimen collection for ionized calcium determinations. The primary audience for this publication is personnel responsible for ionized calcium determinations. This document discusses the reasons for *in vivo* (nonpathologic) and *in vitro* changes in ionized calcium concentrations, and it presents recommendations for avoiding or minimizing these effects.

Clinical and Laboratory Standards Institute (CLSI). *Ionized Calcium Determinations: Precollection Variables, Specimen Choice, Collection, and Handling; Approved Guideline—Second Edition*. CLSI document C31-A2 (ISBN 1-56238-436-8). Clinical and Laboratory Standards Institute, 950 West Valley Road, Suite 2500, Wayne, Pennsylvania 19087, 2001.

The Clinical and Laboratory Standards Institute consensus process, which is the mechanism for moving a document through two or more levels of review by the health care community, is an ongoing process. Users should expect revised editions of any given document. Because rapid changes in technology may affect the procedures, methods, and protocols in a standard or guideline, users should replace outdated editions with the current editions of CLSI documents. Current editions are listed in the CLSI catalog and posted on our website at www.clsi.org. If your organization is not a member and would like to become one, and to request a copy of the catalog, contact us at: Telephone: 610.688.0100; Fax: 610.688.0700; E-Mail: customerservice@clsi.org; Website: www.clsi.org.



Copyright ©2001 Clinical and Laboratory Standards Institute. Except as stated below, any reproduction of content from a CLSI copyrighted standard, guideline, companion product, or other material requires express written consent from CLSI. All rights reserved. Interested parties may send permission requests to permissions@clsi.org.

CLSI hereby grants permission to each individual member or purchaser to make a single reproduction of this publication for use in its laboratory procedure manual at a single site. To request permission to use this publication in any other manner, e-mail permissions@clsi.org.

Suggested Citation

CLSI. *Ionized Calcium Determinations: Precollection Variables, Specimen Choice, Collection, and Handling; Approved Guideline—Second Edition*. CLSI document C31-A2. Wayne, PA: Clinical and Laboratory Standards Institute; 2001.

Proposed Guideline

November 1993

Approved Guideline

December 1995

Approved Guideline—Second Edition

June 2001

ISBN 1-56238-436-8
ISSN 0273-3099

Committee Membership

Area Committee on Clinical Chemistry and Toxicology

W. Gregory Miller, Ph.D.
Chairholder

Virginia Commonwealth University
Richmond, Virginia

Gary L. Myers, Ph.D.
Vice-Chairholder

Centers for Disease Control and Prevention
Atlanta, Georgia

Paul D'Orazio, Ph.D.

Instrumentation Laboratory
Lexington, Massachusetts

Basil T. Doumas, Ph.D.

Medical College of Wisconsin
Milwaukee, Wisconsin

John H. Eckfeldt, M.D., Ph.D.

Fairview-University Medical Center
Minneapolis, Minnesota

Susan A. Evans, Ph.D.

Dade Behring Inc.
Deerfield, Illinois

Gary A. Graham, Ph.D., DABCC

Big Sandy, Texas

Patrick J. Parsons, Ph.D.

New York State Department of Health
Albany, New York

Noel V. Stanton, M.S.

WI State Laboratory of Hygiene
Madison, Wisconsin

Advisors

George N. Bowers, Jr., M.D.

Hartford Hospital
Hartford, Connecticut

Robert W. Burnett, Ph.D.

Hartford Hospital
Hartford, Connecticut

Mary F. Burritt, Ph.D.

Mayo Clinic
Rochester, Minnesota

Kevin D. Fallon, Ph.D.

Instrumentation Laboratory
Lexington, Massachusetts

Carl C. Garber, Ph.D.

Quest Diagnostics, Incorporated
Teterboro, New Jersey

Harvey W. Kaufman, M.D.

Quest Diagnostics, Incorporated
Teterboro, New Jersey

Richard R. Miller, Jr.

Dade Behring Inc.
Newark, Delaware

Advisors (Continued)

Robert F. Moran, Ph.D., FCCM, FAIC

mvi Sciences
Methuen, Massachusetts

Bette Seamonds, Ph.D.

National Academy of Clinical Biochemistry
Swarthmore, Pennsylvania

Beth Ann Wise, M.T.(ASCP), M.S.Ed.
Staff Liaison

NCCLS
Wayne, Pennsylvania

Patrice E. Polgar
Editor

NCCLS
Wayne, Pennsylvania

Donna M. Wilhelm
Assistant Editor

NCCLS
Wayne, Pennsylvania

Acknowledgements

The Area Committee on Clinical Chemistry and Toxicology extends its appreciation to Paul D. D’Orazio, Ph.D., Co-Chairholder of the former Subcommittee on Electrolytes, John G. Toffaletti, Ph.D., and Jesper Wandrup, M.D., Ph.D. for their help and advice in preparing the second edition of this approved-level guideline.

In addition, the area committee would also like to recognize the valuable contributions of the members and advisors of the Subcommittee on Electrolytes that developed the first approved edition of this guideline.

Paul D’Orazio, Ph.D., Co-Chairholder
Gary A. Graham, Ph.D., Co-Chairholder
Carolyn Bergkuist, M.S.
Alan D. Cormier, Ph.D.
Sharon Ehrmeyer, Ph.D.
William F. Koch, Ph.D.
Ioannis Laios, Ph.D.
Arthur Malenfant, Ph.D.
Richard R. Miller
John G. Toffaletti, Ph.D.
Jesper Wandrup, M.D., Ph.D.

Advisors

George N. Bowers, Jr., M.D.
Robert W. Burnett, Ph.D.
Roger R. Calam, Ph.D.
Kent C. Dooley, Ph.D.
Richard A. Durst, Ph.D.
Wayne J. Gilli, BSEE
Neil Greenberg, Ph.D.
Jack H. Ladenson, Ph.D.
Robert F. Moran, Ph.D., FCCM, FAIC
Kathleen O’Connell, Ph.D.
Anthony O. Okorodudu, Ph.D.
Charles Sachs, M.D.
James L. Seago, Ph.D.
Salvador Sena, Ph.D., DABCC
Lawrence Uhteg, M.D.
Paul A. Van Dreal, Ph.D.
Francesco Zoppi

Active Membership (as of 1 April 2001)

Sustaining Members

Abbott Laboratories
 American Association for
 Clinical Chemistry
 Bayer Corporation
 Beckman Coulter, Inc.
 BD and Company
 bioMérieux, Inc.
 College of American Pathologists
 Dade Behring Inc.
 GlaxoSmithKline
 Nippon Becton Dickinson Co., Ltd.
 Ortho-Clinical Diagnostics, Inc.
 Pfizer Inc
 Roche Diagnostics, Inc.

Professional Members

AISAR-Associazione Italiana per lo
 Studio degli
 American Academy of Family
 Physicians
 American Association for
 Clinical Chemistry
 American Association for
 Respiratory Care
 American Chemical Society
 American Medical Technologists
 American Public Health Association
 American Society for Clinical
 Laboratory Science
 American Society of Hematology
 American Society for Microbiology
 American Society of
 Parasitologists, Inc.
 American Type Culture
 Collection, Inc.
 Asociacion de Laboratorios de Alta
 Complejidad
 Asociación Española Primera de
 Socorros (Uruguay)
 Asociacion Mexicana de
 Bioquimica Clinica A.C.
 Assn. of Public Health Laboratories
 Assoc. Micro. Clinici Italiani-
 A.M.C.L.I.
 Australasian Association of
 Clinical Biochemists
 British Society for Antimicrobial
 Chemotherapy
 CADIME-Camara De Instituciones
 De Diagnostico Medico

Canadian Society for Medical
 Laboratory Science—Société
 Canadienne de Science de
 Laboratoire Médical
 Canadian Society of Clinical
 Chemists
 Clinical Laboratory Management
 Association
 COLA
 College of American Pathologists
 College of Medical Laboratory
 Technologists of Ontario
 College of Physicians and
 Surgeons of Saskatchewan
 Fundación Bioquímica Argentina
 International Association of Medical
 Laboratory Technologists
 International Council for
 Standardization in Haematology
 International Federation of
 Clinical Chemistry
 Italian Society of Clinical
 Biochemistry
 Japan Society of Clinical Chemistry
 Japanese Committee for Clinical
 Laboratory Standards
 Joint Commission on Accreditation
 of Healthcare Organizations
 National Academy of Clinical
 Biochemistry
 National Society for
 Histotechnology, Inc.
 Ontario Medical Association
 Quality Management Program-
 Laboratory Service
 RCPA Quality Assurance Programs
 PTY Limited
 Sociedade Brasileira de Analises
 Clinicas
 Sociedade Brasileira de
 Patologia Clinica
 Sociedad Espanola de Bioquimica
 Clinica y Patologia Molecular

Government Members

Association of Public Health
 Laboratories
 Armed Forces Institute of Pathology
 BC Centre for Disease Control
 Centers for Disease Control and
 Prevention
 Chinese Committee for Clinical
 Laboratory Standards

Commonwealth of Pennsylvania
 Bureau of Laboratories
 Department of Veterans Affairs
 Deutsches Institut für Normung
 (DIN)
 FDA Center for Devices and
 Radiological Health
 FDA Center for Veterinary
 Medicine
 FDA Division of Anti-Infective
 Drug Products
 Health Care Financing
 Administration/CLIA Program
 Health Care Financing
 Administration
 Iowa State Hygienic Laboratory
 Massachusetts Department of
 Public Health Laboratories
 National Association of Testing
 Authorities – Australia
 National Center of Infectious
 and Parasitic Diseases (Bulgaria)
 National Institute of Standards
 and Technology
 Ohio Department of Health
 Ontario Ministry of Health
 Saskatchewan Health-Provincial
 Laboratory
 Scientific Institute of Public Health;
 Belgium Ministry of Social
 Affairs, Public Health and the
 Environment
 South African Institute for Medical
 Research
 Swedish Institute for Infectious
 Disease Control
 Thailand Department of Medical
 Sciences

Industry Members

AB Biodisk
 Abbott Laboratories
 Abbott Laboratories, MediSense
 Products
 Accumetrics, Inc.
 Agilent Technologies, Inc.
 Ammirati Regulatory Consulting
 Assessor
 AstraZeneca
 Aventis
 Avocet Medical, Inc.
 Bayer Corporation – Elkhart, IN

Bayer Corporation – Tarrytown, NY
 Bayer Corporation – West Haven, CT
 Bayer Medical Ltd.
 BD
 BD Biosciences – San Jose, CA
 BD Consumer Products
 BD Diagnostic Systems
 BD Italia S.P.A.
 BD VACUTAINER Systems
 Beckman Coulter, Inc.
 Beckman Coulter, Inc. Primary Care Diagnostics
 Beckman Coulter K.K. (Japan)
 Bio-Development SRL
 Bio-Inova Life Sciences International
 Bio-Inova Life Sciences North America
 BioMedia Laboratories Sdn Bhd
 bioMérieux, Inc.
 Biometrology Consultants
 Bio-Rad Laboratories, Inc.
 Bio-Rad Laboratories, Inc. - France
 Biotest AG
 Bristol-Myers Squibb Company
 Canadian External Quality Assessment Laboratory
 Capital Management Consulting, Inc.
 Checkpoint Development Inc.
 Clinical Design Group Inc.
 Clinical Laboratory Improvement Consultants
 COBE Laboratories, Inc.
 Community Medical Center (NJ)
 Control Lab (Brazil)
 Copan Diagnostics Inc.
 Cosmetic Ingredient Review
 Cubist Pharmaceuticals
 Cytometrics, Inc.
 Dade Behring Inc. - Deerfield, IL
 Dade Behring Inc. - Glasgow, DE
 Dade Behring Inc. - Marburg, Germany
 Dade Behring Inc. - Sacramento, CA
 Dade Behring Inc. - San Jose, CA
 DAKO A/S
 Diagnostic Products Corporation
 Eiken Chemical Company, Ltd.
 Enterprise Analysis Corporation
 Fort Dodge Animal Health
 General Hospital Vienna (Austria)
 Gen-Probe
 GlaxoSmithKline
 Greiner Bio-One Inc.
 Health Systems Concepts, Inc.
 Helena Laboratories
 Home Diagnostics, Inc.

I-STAT Corporation
 International Technidyne Corporation
 Kendall Sherwood-Davis & Geck
 LAB-Interlink, Inc.
 Labtest Diagnostica S.A.
 LifeScan, Inc. (a Johnson & Johnson Company)
 Lilly Research Laboratories
 Medical Device Consultants, Inc.
 Medtronic, Inc.
 Merck & Company, Inc.
 mvi Sciences (MA)
 Nabi
 Neometrics is.
 Nichols Institute Diagnostics (Div. of Quest Diagnostics, Inc.)
 Nissui Pharmaceutical Co., Ltd.
 Nippon Becton Dickinson Co., Ltd.
 Norfolk Associates, Inc.
 Organon Teknika Corporation
 Ortho-Clinical Diagnostics, Inc. (Raritan, NJ)
 Ortho-Clinical Diagnostics, Inc. (Rochester, NY)
 Oxoid Inc.
 Pfizer Inc
 Pharmacia Corporation
 Premier Inc.
 Procter & Gamble Pharmaceuticals, Inc.
 The Product Development Group
 Quintiles, Inc.
 Radiometer America, Inc.
 Radiometer Medical A/S
 David G. Rhoads Associates, Inc.
 Roche Diagnostics GmbH
 Roche Diagnostics, Inc.
 Roche Laboratories (Div. Hoffmann-La Roche Inc.)
 The R.W. Johnson Pharmaceutical Research Institute
 Sarstedt, Inc.
 SARL Laboratoire Carron (France)
 Schering Corporation
 Schleicher & Schuell, Inc.
 Second Opinion
 Showa Yakuhin Kako Company, Ltd.
 Streck Laboratories, Inc.
 SurroMed, Inc.
 Sysmex Corporation (Japan)
 Sysmex Corporation (Long Grove, IL)
 The Toledo Hospital (OH)
 Trek Diagnostic Systems, Inc.
 Vetoquinol S.A.
 Visible Genetics, Inc.
 Vysis, Inc.

Wallac Oy
 Wyeth-Ayerst
 Xyletech Systems, Inc.
 YD Consultant
 Yeongdong Pharmaceutical Corporation

Trade Associations

AdvaMed
 Association of Medical Diagnostic Manufacturers
 Japan Association Clinical Reagents Ind. (Tokyo, Japan)
 Medical Industry Association of Australia

Associate Active Members

20th Medical Group (SC)
 67th CSH Wuerzburg, GE (NY)
 121st General Hospital (CA)
 Academisch Ziekenhuis-VUB (Belgium)
 Acadiana Medical Laboratories, LTD (LA)
 Adena Regional Medical Center (OH)
 Advocate Laboratories (IL)
 The Aga Khan Hospital & Medical College, Karachi (Pakistan)
 Akershus Central Hospital and AFA (Norway)
 Albany Medical Center Hospital (NY)
 Albemarle Hospital (NC)
 Allegheny General Hospital (PA)
 Allegheny University of the Health Sciences (PA)
 Allina Laboratories (MN)
 Alton Ochsner Medical Foundation (LA)
 American Medical Laboratories (VA)
 Arkansas Department of Health
 ARUP at University Hospital (UT)
 Armed Forces Research Institute of Medical Science (APO, AP)
 Associated Regional & University Pathologists (UT)
 Aurora Consolidated Laboratories (WI)
 Bay Medical Center (MI)
 Baystate Medical Center (MA)
 Bbaguas Duzen Laboratories (Turkey)
 Bo Ali Hospital (Iran)
 Bonnyville Health Center (Alberta, Canada)

Boulder Community Hospital (CO)	Durham Regional Hospital (NC)	Huddinge University Hospital (Sweden)
Brantford General Hospital (Ontario, Canada)	Dynacare Laboratories - Eastern Region (Ottawa, ON, Canada)	Hurley Medical Center (MI)
Brazileiro De Promocao (Brazil)	Dynacare Memorial Hermann Laboratory Services (TX)	Indiana State Board of Health
Brookdale Hospital Medical Center (NY)	E.A. Conway Medical Center (LA)	Indiana University
Brooke Army Medical Center (TX)	Eastern Maine Medical Center	Instituto Scientifico HS. Raffaele (Italy)
Brooks Air Force Base (TX)	East Side Clinical Laboratory (RI)	International Health Management Associates, Inc. (IL)
Broward General Medical Center (FL)	Elyria Memorial Hospital (OH)	Jersey Shore Medical Center (NJ)
Calgary Laboratory Services	Emory University Hospital (GA)	Joel T. Boone Branch Medical Clinic (VA)
Carilion Consolidated Laboratory (VA)	Esoterix Center for Infectious Disease (TX)	John F. Kennedy Medical Center (NJ)
Cathay General Hospital (Taiwan)	Fairfax Hospital (VA)	John Randolph Hospital (VA)
CB Healthcare Complex (Sydney, NS, Canada)	Fairview-University Medical Center (MN)	Kaiser Permanente (CA)
Central Kansas Medical Center	Foothills Hospital (Calgary, AB, Canada)	Kaiser Permanente (MD)
Central Texas Veterans Health Care System	Fort St. John General Hospital (Fort St. John, BC, Canada)	Kantonsspital (AG, Switzerland)
Centro Diagnostico Italiano (Milano, Italy)	Fox Chase Cancer Center (PA)	Kenora-Rainy River Regional Laboratory Program (Ontario, Canada)
Champlain Valley Physicians Hospital (NY)	Franklin Square Hospital Center (MD)	Kern Medical Center (CA)
Chang Gung Memorial Hospital (Taiwan)	Fresenius Medical Care/Spectra East (NJ)	King Fahad National Guard Hospital (Saudi Arabia)
Children's Hospital (LA)	Fresno Community Hospital and Medical Center	King Khalid National Guard Hospital (Saudi Arabia)
Children's Hospital (NE)	Gambro Healthcare Laboratory (FL)	Kings County Hospital Center (NY)
Children's Hospital & Clinics (MN)	GDS Technology, Inc (IN)	Klinični Center (Slovenia)
Children's Hospital King's Daughters (VA)	Geisinger Medical Center (PA)	LabCorp (NC)
Children's Hospital Medical Center (Akron, OH)	Grady Memorial Hospital (GA)	Laboratories at Bonfils (CO)
Children's Hospital of Philadelphia (PA)	Guthrie Clinic Laboratories (PA)	Laboratório Fleury S/C Ltda. (Brazil)
Clarian Health-Methodist Hospital (IN)	Harris Methodist Erath County (TX)	Laboratory Corporation of America (MO)
Clendo Lab (Puerto Rico)	Harris Methodist Fort Worth (TX)	LAC and USC Healthcare Network (CA)
CLSI Laboratories (PA)	Harris Methodist Northwest (TX)	Lakeland Regional Medical Center (FL)
Columbus County Hospital (NC)	Hartford Hospital (CT)	Lancaster General Hospital (PA)
Commonwealth of Kentucky	Headwaters Health Authority (Alberta, Canada)	Langley Air Force Base (VA)
CompuNet Clinical Laboratories (OH)	Health Network Lab (PA)	LeBonheur Children's Medical Center (TN)
Covance Central Laboratory Services (IN)	Health Sciences Centre (Winnipeg, MB, Canada)	Lewis-Gale Medical Center (VA)
Danish Veterinary Laboratory (Copenhagen, Denmark)	Heartland Health System (MO)	Libero Instituto Univ. Campus BioMedico (Italy)
Danville Regional Medical Center (VA)	Highlands Regional Medical Center (FL)	Licking Memorial Hospital (OH)
Deaconess Hospital (MO)	Hoag Memorial Hospital Presbyterian (CA)	Long Beach Memorial Medical Center (CA)
Delaware Public Health Laboratory	Holmes Regional Medical Center (FL)	Louisiana State University Medical Center
Department of Health & Community Services (New Brunswick, Canada)	Holy Spirit Hospital (PA)	Maccabi Medical Care and Health Fund (Israel)
Detroit Health Department (MI)	Holzer Medical Center (OH)	Magee Womens Hospital (PA)
Diagnostic Laboratory Services, Inc. (HI)	Hospital for Sick Children (Toronto, ON, Canada)	Magnolia Regional Health Center (MS)
Duke University Medical Center (NC)	Hospital Israelita Albert Einstein (Brazil)	Martin Luther King/Drew Medical Center (CA)
	Hotel Dieu Hospital (Windsor, ON, Canada)	

Massachusetts General Hospital (Microbiology Laboratory)	North Shore – Long Island Jewish Health System Laboratories (NY)	St. Elizabeth Hospital (NJ)
Massachusetts General Hospital (Pathology Laboratory)	Northridge Hospital Medical Center (CA)	St-Eustache Hospital (Quebec, Canada)
Mayo Clinic Scottsdale (AZ)	Northwestern Memorial Hospital (IL)	St. John Hospital and Medical Center (MI)
MDS Metro Laboratory Services (Burnaby, BC, Canada)	Ohio Valley Medical Center (WV)	St. John Regional Hospital (St. John, NB, Canada)
Medical College of Virginia Hospital	O.L. Vrouwziekenhuis (Belgium)	St. Joseph Hospital (NE)
Medicare/Medicaid Certification, State of North Carolina	Ordre professionnel des technologistes médicaux du Québec	St. Joseph Mercy – Oakland (MI)
Memorial Hospital (CO)	Ospedali Riuniti (Italy)	St. Joseph’s Hospital – Marshfield Clinic (WI)
Memorial Medical Center (Napoleon Ave., New Orleans, LA)	The Ottawa Hospital (Ottawa, ON, Canada)	St. Joseph’s Medical Center (CA)
Memorial Medical Center (N. Jefferson Davis Pkwy., New Orleans, LA)	Our Lady of Lourdes Hospital (NJ)	St. Luke’s Regional Medical Center (IA)
Memorial Medical Center (IL)	Our Lady of the Resurrection Medical Center (IL)	St. Mark’s Hospital (UT)
Mercy Health System (PA)	Pathology and Cytology Laboratories, Inc. (KY)	St. Mary Medical Center (IN)
Mercy Hospital (NC)	Pathology Associates Laboratories (CA)	St. Mary of the Plains Hospital (TX)
Mercy Medical Center Des Moines (IA)	The Permanente Medical Group (CA)	St. Mary’s Hospital & Medical Center (CO)
Mescalero Indian Hospital (NM)	Pocono Hospital (PA)	St. Paul’s Hospital (Vancouver, BC, Montreal)
Methodist Hospitals of Memphis (TN)	Presbyterian Hospital of Dallas (TX)	Ste. Justine Hospital (Montreal, PQ, Canada)
Michigan Department of Community Health	Prodia Clinical Laboratory (Indonesia)	Salina Regional Health Center (KS)
Mississippi Baptist Medical Center	Providence Health System (OR)	San Francisco General Hospital (CA)
Monmouth Medical Center (NJ)	Providence Seattle Medical Center (WA)	Santa Cabrini Hospital (Montreal, PQ Canada)
Monte Tabor – Centro Italo - Brazileiro de Promocao (Brazil)	Queen Elizabeth Hospital (Prince Edward Island, Canada)	Santa Clara Valley Medical Center (CA)
Montreal Children’s Hospital (Canada)	Queensland Health Pathology Services (Australia)	Seoul Nat’l University Hospital (Korea)
Montreal General Hospital (Canada)	Quest Diagnostics, Incorporated (AZ)	Shanghai Center for the Clinical Laboratory (China)
Morton Plant Mease Health Care (FL)	Quest Diagnostics Incorporated (CA)	Shands Healthcare (FL)
Mount Sinai Hospital (NY)	Quintiles Laboratories, Ltd. (GA)	South Bend Medical Foundation (IN)
Mount Sinai Medical Center (FL)	Reading Hospital and Medical Center (PA)	Southern California Permanente Medical Group
MRL Pharmaceutical Services, Inc. (VA)	Regions Hospital	South Western Area Pathology Service (Australia)
MRL Reference Laboratory (CA)	Research Medical Center (MO)	Specialty Laboratories, Inc. (CA)
National University Hospital (Singapore)	Rex Healthcare (NC)	Stanford Hospital and Clinics (CA)
Naval Surface Warfare Center (IN)	Rhode Island Department of Health Laboratories	State of Washington Department of Health
New Britain General Hospital (CT)	Riyadh Armed Forces Hospital (Saudi Arabia)	Stormont-Vail Regional Medical Center (KS)
New England Fertility Institute (CT)	Royal Columbian Hospital (New Westminster, BC, Canada)	Sun Health-Boswell Hospital (AZ)
New England Medical Center Hospital (MA)	Saint Mary’s Regional Medical Center (NV)	Sunrise Hospital and Medical Center (NV)
New York Hospital Medical Center of Queens	St. Alexius Medical Center (ND)	T.A. Sourasky Medical Center (Israel)
New York State Department of Health	St. Anthony Hospital (CO)	Temple University Hospital (PA)
NorDx (ME)	St. Barnabas Medical Center (NJ)	Tenet Odessa Regional Hospital (TX)
North Carolina Laboratory of Public Health	St. Boniface General Hospital (Winnipeg, Canada)	The Toledo Hospital (OH)
North Mississippi Medical Center		Touro Infirmary (LA)
		Tri-City Medical Center (CA)

Trident Regional Medical Center (SC)	University of Texas M.D. Anderson Cancer Center	Vejle Hospital (Denmark)
Tripler Army Medical Center (HI)	University of Virginia Medical Center	Virginia Department of Health
Truman Medical Center (MO)	University of Washington	Viridae Clinical Sciences, Inc. (Vancouver, BC, Canada)
UCSF Medical Center (CA)	UPMC Bedford Memorial (PA)	Washoe Medical Center Laboratory (NV)
UNC Hospitals (NC)	UZ-KUL Medical Center (Belgium)	Watson Clinic (FL)
Unilab Clinical Laboratories (CA)	VA (Dayton) Medical Center (OH)	Wilford Hall Medical Center (TX)
University Hospital (Gent) (Belgium)	VA (Denver) Medical Center (CO)	William Beaumont Hospital (MI)
University Hospital (TX)	VA (Kansas City) Medical Center (MO)	Williamsburg Community Hospital (VA)
The University Hospitals (OK)	VA (Martinez) Medical Center (CA)	Winn Army Community Hospital (GA)
University of Alabama-Birmingham Hospital	VA (San Diego) Medical Center (CA)	Wishard Memorial Hospital (IN)
University of Alberta Hospitals (Canada)	VA (Tuskegee) Medical Center (AL)	Yonsei University College of Medicine (Korea)
University of Chicago Hospitals (IL)	VA Outpatient Clinic (OH)	York Hospital (PA)
University of Florida		Zale Lipshy University Hospital (TX)
University of the Ryukyus (Japan)		

OFFICERS

BOARD OF DIRECTORS

F. Alan Andersen, Ph.D., President Cosmetic Ingredient Review	Susan Blonshine, RRT, RPFT, FAARC TechEd	Tadashi Kawai, M.D., Ph.D. International Clinical Pathology Center
Donna M. Meyer, Ph.D., President Elect CHRISTUS Health	Kurt H. Davis, FCSMLS, CAE Canadian Society for Medical Laboratory Science	J. Stephen Kroger, M.D., FACP COLA
Robert F. Moran, Ph.D., F.C.C.M., F.A.I.C. Secretary mvi Sciences	Robert L. Habig, Ph.D. Newtown Square, PA	Barbara G. Painter, Ph.D. Bayer Corporation
Gerald A. Hoeltge, M.D. Treasurer The Cleveland Clinic Foundation	Thomas L. Hearn, Ph.D. Centers for Disease Control and Prevention	Emil Voelkert, Ph.D. Roche Diagnostics GmbH
William F. Koch, Ph.D., Immediate Past President National Institute of Standards and Technology	Elizabeth D. Jacobson, Ph.D. FDA Center for Devices and Radiological Health	Ann M. Willey, Ph.D. New York State Department of Health
John V. Bergen, Ph.D., Executive Director	Carolyn D. Jones, J.D., M.P.H. AdvaMed	Judith A. Yost, M.A., M.T.(ASCP) Health Care Financing Administration

Contents

Abstract..... i

Committee Membership..... iii

Active Membership..... vii

Foreword..... xv

1 Introduction..... 1

2 Scope..... 1

3 Standard Precautions..... 1

4 Definitions 1

5 Precollection Variables: Influences of Physical Activity, Posture, Meals, Ventilation Rate, and Circadian Variation..... 2

 5.1 Effect of Physical Activity..... 2

 5.2 Influences of Posture and Prolonged Bed Rest..... 2

 5.3 Effect of Meals..... 3

 5.4 Ventilation Rate 3

 5.5 Circadian Variation..... 3

 5.6 Recommendations..... 3

6 Specimen Choice 3

 6.1 Whole Blood..... 3

 6.2 Serum..... 4

 6.3 Plasma..... 5

 6.4 Recommendations..... 5

7 Specimen Collection 6

 7.1 Collection Site Selection..... 6

 7.2 Collection Devices..... 6

 7.3 Collection Techniques 7

 7.4 Recommendations..... 7

8 Specimen Transportation, Processing, and Storage 8

 8.1 Anticoagulated Whole Blood in Syringes..... 8

 8.2 Serum..... 8

 8.3 Recommendations for Transporting Specimens 9

9 Specimen Handling During Analysis..... 10

 9.1 Whole Blood..... 10

 9.2 Serum..... 10

 9.3 Recommendations..... 11

References..... 12

Appendix A. Anticoagulants..... 15

Appendix B. Specimen Type: Arterial, Venous, or Capillary Blood..... 19

Contents (Continued)

Appendix C. Aerobically Handled Tubes (pH-Adjusted Ionized Calcium)	20
Summary of Comments and Committee Responses	22
Related NCCLS Publications.....	23

Foreword

Ionized calcium determinations have proven to be clinically useful in the differential diagnosis of calcium disorders of endocrine origin, identification of hypercalcemia in various neoplasias, and managing the critically ill adult and neonatal patient. However, it is the responsibility of the laboratorian to choose which specimen is most appropriate for each clinical situation and how to collect and handle that specimen to ensure accuracy and clinical utility. This choice is complicated by the equilibrium between free (ionized) and bound calcium in blood, which is influenced by alterations in hydrogen ion and/or ligand concentrations. This guideline is designed to aid the laboratorian in determining the most appropriate specimen and its proper handling for each specific purpose.

Specifically, C31-A2 offers guidance in recognizing preanalytical factors that can affect ionized calcium determinations. The influence of patient conditions (e.g., physical activity, posture, meals, ventilation rate, and circadian variation) is considered in Section 5, while the advantages and disadvantages of whole blood, serum, and plasma are discussed in Section 6. The guideline also describes the selection of the collection site and device in Section 7. In Section 8, appropriate transportation, processing, and storage procedures are recommended.

References to pH-adjusted ionized calcium results are found throughout the guideline, and appropriate citations are provided.

Key Words

Ionized calcium, pH, preanalytical conditions, precollection variables, specimen choice, specimen collection, specimen transportation

Ionized Calcium Determinations: Precollection Variables, Specimen Choice, Collection, and Handling; Approved Guideline—Second Edition

1 Introduction

Ionized calcium is widely recognized as a better indicator of physiological calcium status in blood than total calcium. Generally, the reasons for measuring ionized calcium can be divided into three clinical categories: monitoring trends in acute or critical care, routine diagnostic care, and research. Generally, ionized calcium measurements for diagnostic purposes or research purposes require a high degree of accuracy.

This document describes the preanalytical variables for ionized calcium determinations and makes recommendations for minimizing the effects of these variables on the accuracy of ionized calcium measurements. Patient preparation and specimen handling options are presented, as well as the advantages and disadvantages of the various choices for specimen type, collection device, and technique. Recommendations are offered in each section.

2 Scope

This document addresses the preanalytical variables that can influence the accuracy and clinical utility of ionized calcium measurements.

3 Standard Precautions

Because it is often impossible to know which specimens might be infectious, all human blood specimens are to be treated as infectious and handled according to “standard precautions.” Standard precautions are new guidelines that combine the major features of “universal precautions and body substance isolation” practices. Standard precautions cover the transmission of any pathogen and thus are more comprehensive than universal precautions which are intended to apply only to transmission of blood-borne pathogens. Standard precaution and universal precaution guidelines are available from the U.S. Centers for Disease Control and Prevention (*Guideline for Isolation Precautions in Hospitals*. Infection Control and Hospital Epidemiology. CDC. 1996;Vol 17;1:53-80.), [MMWR 1987;36(suppl 2S):2S-18S] and (MMWR 1988;37:377-382, 387-388). For specific precautions for preventing the laboratory transmission of blood-borne infection from laboratory instruments and materials; and recommendations for the management of blood-borne exposure, refer to NCCLS document M29—*Protection of Laboratory Workers from Instrument Biohazards and Infectious Disease Transmitted by Blood, Body Fluids, and Tissue*.

4 Definitions^a

Circadian variation/chronobiological variation, diurnal variation, *n* – Variations in physiological parameters, including blood analyte concentrations, which are related to cyclic events, i.e., time of day, season of the year, and ingestion of meals.

Ionized calcium, *n* – The portion of calcium ions in the plasma water of whole blood that is not bound by protein or other molecules; **NOTE:** This parameter has also been called “free” or “ionic” calcium.

pH-adjusted ionized calcium, *n* – A calculated result empirically based on a measured pH and ionized calcium concentration, with the ionized calcium concentration normalized to a pH of 7.40; **NOTE:** These

^a Some of these definitions are found in NCCLS document NRSCL8—*Terminology and Definitions for Use in NCCLS Documents*. For complete definitions and detailed source information, please refer to the most current edition of that document.