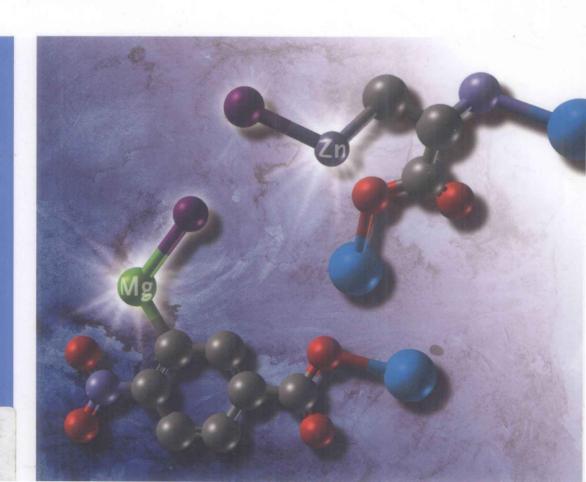
## Handbook of Functionalized Organometallics

Applications in Synthesis

Volume 1



# Handbook of Functionalized Organometallics

Applications in Synthesis

Volume 1

Edited by Paul Knochel



WILEY-VCH Verlag GmbH & Co. KGaA

#### Editor

Prof. Paul Knochel
Department of Chemistry
Ludwig-Maximilians-Universität
Butenandtstraße 5–13
Haus F
81377 München
Germany

All books published by Wiley-VCH are carefully produced. Nevertheless, authors, editors, and publisher do not warrant the information contained in these books, including this book, to be free of errors. Readers are advised to keep in mind that statements, data, illustrations, procedural details or other items may inadvertently be inaccurate.

Library of Congress Card No.: applied for

**British Library Cataloguing-in-Publication Data** A catalogue record for this book is available from the British Library.

### Bibliographic information published by Die Deutsche Bibliothek

Die Deutsche Bibliothek lists this publication in the Deutsche Nationalbibliografie; detailed bibliographic data is available in the Internet at <a href="http://dnb.ddb.de">http://dnb.ddb.de</a>>.

© 2005 WILEY-VCH Verlag GmbH & Co. KGaA, Weinheim

All rights reserved (including those of translation into other languages). No part of this book may be reproduced in any form – nor transmitted or translated into machine language without written permission from the publishers. Registered names, trademarks, etc. used in this book, even when not specifically marked as such, are not to be considered unprotected by law.

Printed in the Federal Republic of Germany. Printed on acid-free paper.

Cover Design Grafik-Design Schulz, Fußgönheim Typesetting Kühn & Weyh, Satz und Medien, Freiburg

Printing betz-druck GmbH, Darmstadt
Bookbinding Litges & Dopf Buchbinderei GmbH,
Heppenheim

**ISBN-13:** 978-3-527-31131-6 **ISBN-10:** 3-527-31131-9

## Handbook of Functionalized Organometallics

Volume 1

Edited by Paul Knochel

#### **Related Titles**

Tamaru, Y. (Ed.)

**Modern Organonickel Chemistry** 

2005, ISBN 3-527-30796-6

Bäckvall, J.-E. (Ed.)

**Modern Oxidation Methods** 

2004, ISBN 3-527-30642-0

Murahashi, S.-I. (Ed.)

**Ruthenium in Organic Synthesis** 

2004, ISBN 3-527-30692-7

Cornils, B., Herrmann, W. A. (Eds.)

Aqueous-Phase Organometallic Catalysis

**Concepts and Applications** 

2004. ISBN 3-527-30712-5

de Meijere, A., Diederich, F. (Eds.)

Metal-Catalyzed Cross-Coupling Reactions

938 pages in 2 volumes with 29 figures and 61 tables 2004, ISBN 3-527-30518-1

Marek, I. (Ed.)

Titanium and Zirconium in Organic Synthesis

2002, ISBN 3-527-30428-2

Beller, M., Bolm, C. (Eds.)

Transition Metals for Organic Synthesis

**Building Blocks and Fine Chemicals** 

2004, ISBN 3-527-30613-7

Cornils, B., Herrmann, W. A. (Eds.)

Applied Homogeneous Catalysis with Organometallic Compounds

A Comprehensive Handbook in Three Volumes

2002, ISBN 3-527-30434-7

#### **Preface**

Since the pioneering work of Frankland and Wurtz, organometallic intermediates have occupied a central position in organic synthesis. The chemical behavior of organometallic reagents depends greatly on the nature of the metal and on the carbon hybridization. Each metal has intrinsic chemical properties, which confer a specific reactivity for forming new carbon-carbon bonds to the organic moiety attached to it. The nature of the metal substituents (ligands) enables a modulation and adjustment to this reactivity of the organometallic to the organic substrate. Choosing the correct metal and ligand sphere to achieve any given transformation represents a major task for the synthetic chemist. During the course of the last thirty years, chemists have realized that this fine-tuning of the reactivity of organometallics has a number of synthetic advantages (selectivity, yields, reaction conditions, etc.). However, they have also noticed that a broad range of functionalities can be present in the organometallic intermediate itself and therefore these reagents allow for the preparation of polyfunctional molecules without the need for multiple protection and deprotection steps.

This book summarizes the synthetic knowledge available as of 2005 for preparing functionalized organometallics and the optimum conditions for their reacting with electrophilic species. It also covers main group and transition organometallics while outlining in detail the functional group compatibility for each class of organometallics in the various book chapters.

Organometallic chemistry is a field of chemistry that is constantly experiencing discoveries and is one of the motors of chemistry. Thus it can be expected that numerous new synthetic methods based on the use of functionalized organometallics will be added to the chemistry presented in this book within the next few years. An effort has been made to present the material in an attractive layout with many equations and numerous practical details, allowing for rapidly entry in the field. Therefore, this book is well suited for master and PhD students, for advanced undergraduate students, as well as industrial process and research chemists.

Munich, August 2005

Paul Knochel

Organometallics. Paul Knochel Copyright © 2005 WILEY-VCH Verlag GmbH & Co. KGaA, Weinheim ISBN: 3-527-31131-9

#### **List of Authors**

#### Oliver Baron

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5–13, Haus F 81377 München Germany

#### Gerard Cahiez

Departement de Chimie, ESCOM Université de Paris 13, Boulevard de l'Hautil 95092 Cergy-Pontoise France

#### Helena Chechik-Lankin

Department of Chemistry Technion-Israel Institute of Technology Technion City Haifa 32000 Israel

#### Karl-Heinz Dötz

Kékulé-Institut für Organische Chemie und Biochemie Gerhard-Domagk-Strasse 1 53121 Bonn Germany

#### Francisco Foubelo

Departamento de Química Orgánica Facultad de Ciencias and Instituto de Síntesis Orgánica (ISO) Universidad de Alicante Apdo. 99 03080 Alicante Spain

#### **Eric Fouquet**

Laboratoire de Chimie Organique et Organometallique Université de Bordeaux I 351, Cours de la Liberation 33405 Talence Cedex France

#### Nina Gommermann

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5–13, Haus F 81377 München Germany

#### Liu-Zhu Gong

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5–13, Haus F 81377 München Germany

Organometallics. Paul Knochel Copyright © 2005 WILEY-VCH Verlag GmbH & Co. KGaA, Weinheim ISBN: 3-527-31131-9

#### Corinne Gosmini

Laboratoire d'Electrochimie. Catalyse et Synthèse Organique 2 à 8, rue Henri-Dunant B. P. 28 94320 Thiais France

#### Agnès Herve

Laboratoire de Chimie Organique et Organometallique Université de Bordeaux I 351. Cours de la Liberation 33405 Talence Cedex France

#### Tamejiro Hiyama

Department of Material Chemistry Graduate School of Engineering **Kyoto University** Kyoto University Katsura Nishikyo-ku Kyoto 615-8510 Japan

#### Li-Fu Huang

Institute of Chemistry Academia Sinica Nangang Taipei Taiwan 125

#### Hiriyakkanavar Ila

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13, Haus F 81377 München Germany

#### Florian F. Kneisel

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13, Haus F 81377 München Germany

#### Paul Knochel

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13, Haus F 81377 München Germany

#### Alexander Koch

Kékulé-Institut für Organische Chemie und Biochemie Gerhard-Domagk-Strasse 1 53121 Bonn Germany

#### Felix Kopp

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13. Haus F 81377 München Germany

#### Tobias J. Korn

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13. Haus F 81377 München Germany

#### Arkady Krasovskiy

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13, Haus F 81377 München Germany

#### Helena Leuser

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13, Haus F 81377 München Germany

#### Tien-Yau Luh

Institute of Chemistry Academia Sinica Nangang Taipei Taiwan 125

#### Florence Mahuteau-Betzer

Departement de Chimie, ESCOM Université de Paris 13, Boulevard de l'Hautil 95092 Cergy-Pontoise France

#### Ilan Marek

Department of Chemistry Technion-Israel Institute of Technology **Technion City** Haifa 32000 Israel

#### Seijiro Matsubara

Graduate School of Engineering **Kyoto University** Kyoutodaigaku-katsura Nishikyo Kyoto 615-8510 Japan

#### Jacques Périchon

Laboratoire d'Electrochimie, Catalyse et Synthèse Organique 2 à 8, rue Henri-Dunant B. P. 28 94320 Thiais France

#### Sylvie Perrone

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13. Haus F 81377 München Germany

#### Ioannis Sapountzis

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13, Haus F 81377 München Germany

#### Masaki Shimizu

Department of Material Chemistry Graduate School of Engineering **Kyoto University** Kyoto University Katsura Nishikyo-ku Kyoto 615-8510 Japan

#### G. Richard Stephenson

Wolfson Materials and Catalysis Centre School of Chemical Sciences and Pharmacy University of East Anglia Norwich NR4 7TI United Kingdom

#### Martin Werner

Kékulé-Institut für Organische Chemie und Biochemie Gerhard-Domagk-Strasse 1 53121 Bonn Germany

#### Xiaoyin Yang

Department of Chemistry Ludwig-Maximilians-Universität Butenandtstrasse 5-13. Haus F 81377 München Germany

#### Miguel Yus

Departamento de Química Orgánica Facultad de Ciencias and Instituto de Síntesis Orgánica (ISO) Universidad de Alicante Apdo. 99 03080 Alicante Spain

#### Contents

Preface	XV
1 ICIACC	27 4

List of Authors XVII

#### Volume 1 1

2.5.4

1 Introduction 1 Paul Knochel and Felix Kopp

2	Polyfunctional Lithium Organometallics for Organic Synthesis 7  Miguel Yus and Francisco Foubelo
2.1	Introduction 7
2.2	a-Functionalized Organolithium Compounds 8
2.2.1	sp³-Hybridized <i>a</i> –Oxygenated Organolithium Compounds 8
2.2.2	sp <sup>2</sup> -Hybridized <i>a</i> -Oxygenated Organolithium Compounds 13
2.2.3	sp³-Hybridized a-Nitrogenated Organolithium Compounds 14
2.2.4	sp <sup>2</sup> -Hybridized <i>a</i> -Nitrogenated Organolithium Compounds 16
2.2.5	Other sp <sup>2</sup> -Hybridized a-Functionalized Organolithium Compounds 18
2.3	β-Functionalized Organolithium Compounds 18
2.3.1	sp <sup>3</sup> -Hybridized $\beta$ -Functionalized Organolithium Compounds 19
2.3.2	sp <sup>2</sup> -Hybridized $\beta$ -Functionalized Organolithium Compounds 22
2.4	γ-Functionalized Organolithium Compounds 24
2.4.1	γ-Functionalized Alkyllithium Compounds 24
2.4.2	γ-Functionalized Allyllithium Compounds 26
2.4.3	γ-Functionalized Benzyllithium Compounds 27
2.4.4	γ-Functionalized Akenyllithium Compounds 28
2.4.5	γ-Functionalized Alkynyllithium Compounds 30
2.5	$\delta$ -Functionalized Organolithium Compounds 31
2.5.1	$\delta$ -Functionalized Alkyllithium Compounds 31
2.5.2	$\delta$ -Functionalized Allyl and Benzyllithium Compounds 32
2.5.3	$\delta$ -Functionalized Alkenyllihium Compounds 33

 $\delta$ -Functionalized Alkynyllithium Compounds 34

Contents	
2.6	Remote Functionalized Organolithium Compounds 34
2.6.1	Remote Functionalized Alkyllithium Compounds 34
2.6.2	Remote Allyl and Benzyllithium Compounds 35
2.6.3	Remote Functionalized Alkenyl- and Alkynyllithium Compounds 36
3	Functionalized Organoborane Derivatives in Organic Synthesis  45  Paul Knochel, Hiriyakkanavar Ila, Tobias J. Korn, and Oliver Baron
3.1	Introduction 45
3.2	Preparation and Reaction of Functionalized Aryl and Heteroaryl Boranes 45
3.2.1	Preparation from Polar Organometallics 45
3.2.2	Preparation from Aryl Halides and Sulfonates by Cross-coupling 50
3.2.3	Synthesis of Functionalized Aryl Boranes by Catalytic Aromatic C–H Borylation 54
3.2.4	Synthesis of Functionalized Trifluoroborates and their Palladium- catalyzed Suzuki–Miyaura Cross-coupling Reactions 57
3.2.5	Palladium-catalyzed Suzuki–Miyaura Cross-coupling Reactions of Functionalized Aryl and Heteroaryl Boronic Esters 58
3.2.6	Copper-mediated Carbon–Heteroatom-Bond-forming Reactions with Functionalized Aryl Boronic Acids 68
3.2.7	Palladium-catalyzed Acylation of Functionalized Aryl Boronic Acids 73
3.2.8	Miscellaneous C–C-bond Formations of Functionalized Aryl Organoboranes 74
3.2.9	Miscellaneous Reactions of Functionalized Alkenyl Boronic Acids 78
3.3	Preparation and Reactions of Functionalized Alkenyl Boranes 79
3.3.1	Synthesis of Alkenyl Boronic Acids by Transmetallation of Alkenyl Grignard Reagents with Boronate Esters 79
3.3.2	Synthesis of Functionalized Alkenyl Boronic Acids by Hydroboration of Functionalized Alkynes and their Suzuki Cross-coupling Reactions 79
3.3.3	Synthesis of Functionalized Alkenyl Boronic Esters by Crossmetathesis 81
3.3.4	Synthesis and Palladium-catalyzed Cross-coupling Reactions of Functionalized Alkenyl Trifluoroborates 82
3.3.5	Palladium-catalyzed Cross-coupling of Functionalized Alkenyl Boronates with Cyclopropyl Iodides 83
3.3.6	Intermolecular Suzuki Cross-coupling Reactions of Functionalized Alkenylborane Derivates: Application in Natural Product Synthesis (Alkenyl B-Alkenyl Coupling) 83
3.3.7	Intramolecular Macrocyclization via Suzuki Cross-coupling of Functionalized Alkenyl Boronic Esters (Alkenyl B-Alkenyl Coupling) 84
3.3.8	Three-component Mannich Reaction of Functionalized Alkenyl Boronic Acids (Petasis Reaction): Synthesis of $\beta$ , $\gamma$ -Unsaturated $\alpha$ -Amino Acids 85

3.3.9	Oxidation of Functionalized Alkenyl Boronic Esters to Aldehydes with Trimethylamine Oxide 86
3.3.10	Lewis-acid-catalyzed Nucleophilic Addition of Functionalized Alkenyl Boronic Esters to Activated N-acyliminium Ions 86
3.4	Preparation and Reactions of Functionalized Alkynlboron Derivatives 87
3.5	Synthesis and Reactions of Functionalized Allylic Boronates 88
3.6	Synthesis and Reactions of Functionalized Cyclopropyl Boronic Esters 90
3.7	Synthesis and Reactions of Functionalized Alkyl Boron Derivates 91
3.7.1	Synthesis of Aminoalkyl Boranes by Hydroboration and their Suzuki Cross-coupling Reaction 91
3.7.2	Synthesis of Functionalized Alkyl Boronates by Nucleophilic 1,4-Conjugate Addition of Borylcopper Species to $a,\beta$ -Unsaturated Carbonyl Compounds 92
3.7.3	Preparation and B-alkyl-Suzuki–Miyaura Cross-coupling Reactions of Functionalized Alkyl Trifluoroborates 93
3.7.4	Silver(I)-promoted Suzuki Cross-coupling of Functionalized <i>n</i> -Alkyl Boronic Acids 94
3.7.5	Alkyl-Alkyl Suzuki Cross-coupling of Functionalized Alkyl Boranes with Alkyl Bromides, Chlorides and Tosylates 95
3.7.6	Synthesis of Natural and Unnatural Amino Acids via B-alkyl Suzuki Coupling of Functionalized Alkyl Boranes 95
3.7.7	Application of Intermolecular B-alkyl Suzuki Cross-coupling of Functionalized Alkyl Boranes in Natural Product Synthesis 96
3.8	Conclusion 104
4	Polyfunctional Magnesium Organometallics for Organic Synthesis 109 Paul Knochel, Arkady Krasovskiy, and Ioannis Sapountzis
4.1	Introduction 109
4.2	Methods of Preparation of Grignard Reagents and their Uncatalyzed Reactions 110
4.2.1	Direct Oxidative Addition of Magnesium to Organic Halides 110
4.2.2	Metalation Reactions with Magnesium Amides 111
4.2.3	The Halogen–Magnesium Exchange Reaction 113
4.2.3.1	Early Studies 113
4.2.3.2	The Preparation of Functionalized Arylmagnesium Reagents 115
4.2.3.3	Halogen–Magnesium Exchange Using Lithium
	Trialkylmagnesiates 128
4.2.3.4	The Preparation of Functionalized Heteroarylmagnesium Reagents 129
4.2.3.4 4.2.4	, 0
	The Preparation of Functionalized Heteroarylmagnesium Reagents 129 The Preparation of Functionalized Alkenylmagnesium Reagents 136 Preparation of Functionalized Alkylmagnesium Reagents 142
4.2.4	The Preparation of Functionalized Heteroarylmagnesium Reagents 129 The Preparation of Functionalized Alkenylmagnesium Reagents 136

VIII	Contents	
	4.4	Application of Functionalized Magnesium Reagents in Cross-coupling Reactions 155
	4.4.1	Palladium-catalyzed Cross-coupling Reactions 155
	4.4.2	Nickel-catalyzed Cross-coupling Reactions 157
	4.4.3	Iron-catalyzed Cross-coupling Reactions 159
	4.5	Summary and Outlook 164
	5	Polyfunctional Silicon Organometallics for Organic Synthesis 173 Masaki Shimizu and Tamejiro Hiyama
	5.1	Introduction 173
	5.2	Allylic Silanes 174
	5.2.1	Intermolecular Reactions of Polyfunctional Allylic Silanes 174
	5.2.2	Intramolecular Reactions of Polyfunctional Allylic Silanes 176
	5.2.3	Tandem Reactions of Polyfunctional Allylic Silanes 180
	5.2.4	Sequential Synthetic Reactions of Metal-containing Allylic Silanes 183
	5.3	Alkenylsilanes 189
	5.3.1	Intermolecular Reactions of Polyfunctional Alkenylsilanes 189
	5.3.2	Intramolecular Reactions of Polyfunctional Alkenylsilanes 190
	5.3.3	Synthetic Reactions of Metal-containing Alkenylsilanes 191
	5.4	Alkylsilanes 193
	5.4.1	Synthetic Reactions of Polyhalomethylsilanes 193
	5.4.2	Synthetic Reactions of Cyclopropyl, Oxiranyl, and Aziridinylsilanes 195
	5.4.3	Synthetic Reactions of Polysilylmethanes 196
	5.5	Miscellaneous Preparations and Reactions of Polyfunctional
		Organosilicon Reagents 197
	6	Polyfunctional Tin Organometallics for Organic Synthesis 203 Eric Fouquet and Agnès Herve
	6.1	Introduction 203
	6.2	Metal-Catalyzed Coupling Reactions 203
	6.2.1	The Stille Cross-Coupling Reaction 203
	6.2.1.1	Mechanism 204
	6.2.1.2	Organotins for the Stille Reaction 205
	6.2.1.3	Substrates 208
	6.2.1.4	Intermolecular Stille Cross-coupling 210
	6.2.1.5	Intramolecular Stille Cross-coupling 212
	6.2.1.6	Solid-Phase-Supported Stille Coupling 214
	6.2.1.7	Stille Coupling Catalytic in Tin 215
	6.2.2	Other Metal-Catalyzed Coupling Reactions 215
	6.2.2.1	Palladium-Catalyzed Reactions 215
	6.2.2.2	Copper-Catalyzed Reactions 215
	6.2.2.3	Nickel-Catalyzed Reactions 216
	6.2.2.4	Rhodium-Catalyzed Reactions 216
	6.3	Nucleophilic Additions 217

6.3.1	Nucleophilic Addition onto Carbonyl Compounds 217
6.3.1.1	Introduction 217
6.3.1.2	Functionalized Allyltins 217
6.3.1.3	Catalytic Use of Lewis Acid 221
6.3.1.4	Enantioselectivity 221
6.3.1.5	Others Organotin Reagents 222
6.3.2	Nucleophilic Addition onto Imines and Related Compounds 224
6.3.2.1	Reactions with Imines 224
6.3.2.2	Other Imino Substrates 225
6.3.2.3	Catalytic Enantioselective Addition 227
6.4	Radical Reactions of Organotins 227
6.4.1	Introduction 227
6.4.2	Allyltins 227
6.4.2.1	Mechanistic Overview 227
6.4.2.2	Functionalized Allyltins 229
6.4.3	Other Organotin Reagents 230
6.4.3.1	Tetraorganotins 230
6.4.3.2	Modified Organotins 231
6.4.4	The Stereoselective Approach 231
6.5	Transmetallations 232
6.5.1	Introduction 232
6.5.2	Tin-to-lithium Exchange 233
6.5.2.1	a-Heterosubstituted Alkyltins 233
6.5.2.2	Alkenyltins 235
6.5.3	Tin to Other Metal Exchanges 236
6.6	Conclusion 236
7	Polyfunctional Zinc Organometallics for Organic Synthesis 251
	Paul Knochel, Helena Leuser, Liu-Zhu Gong, Sylvie Perrone,
	and Florian F. Kneisel
7.1	Introduction 251
7.2	Methods of Preparation of Polyfunctional Organozinc Reagents 252
7.2.1	Classification 252
7.2.2	Preparation of Polyfunctional Organozinc Halides 252
7.2.2.1	Preparation by the Oxidative Addition to Zinc Metal 252
7.2.2.2	Preparation of Organozinc Halides using Transmetallation
	Reactions 261
7.2.3	Preparation of Diorganozincs 270
7.2.3.1	Preparation via an I/Zn Exchange 270
7.2.3.2	The Boron–Zinc Exchange 273
7.2.3.3	Hydrozincation of Alkenes 278
7.2.4	Diverse Methods of Preparation of Allylic Zinc Reagents 278
7.2.5	Preparation of Lithium Triorganozincates 281
7.3	Reactions of Organozinc Reagents 282
	0

۲l	Contents	
1	7.3.1 7.3.2 7.3.2.1 7.3.2.2 7.3.2.3 7.3.2.4 7.3.3 7.3.4 7.3.5	Uncatalyzed Reactions 283 Copper(I)-catalyzed Reactions 292 Substitution Reactions 293 Acylation Reactions 303 Addition Reactions 305 Michael Additions 309 Palladium- and Nickel-catalyzed Reactions 316 Reactions Catalyzed by Titanium and Zirconium(IV) Complexes 326 Reactions of Zinc Organometallics Catalyzed by Cobalt, Iron or Manganese Complexes 332 Conclusion 333
	Index	I 1
	Volume	<b>2</b> 347
	8	Polyfunctional 1,1-Organodimetallic for Organic Synthesis 347 Seijiro Matsubara
	8.1 8.2 8.2.1 8.2.2 8.2.3 8.2.4 8.2.5 8.2.6 8.2.7 8.2.8 8.2.9 8.2.10 8.3 8.3.1 8.3.2 8.3.3	Introduction 347 gem-Dizincio Compounds 348 General View 348 Methylenation with Bis(iodozincio)methane 351 gem-Dizincio Species from gem-Dihaloalkane 357 Alkenylsilane, -Germane, -and Borane Synthesis 360 Stepwise Coupling Reaction with Two Different Electrophiles 361 Reaction with Acyl Chloride and Cyanide 364 1,4-Addition of bis(iodozincio)methane to a,β-unsaturated ketones 365 Cyclopropanation Reaction 367 Pinacolone Rearrangement with Unusual Diastereospecificity 368 gem-Dizincio Reagent Working as Carbenoid 370 Chromium Compounds 371 General View 371 Alkylidenation 371 a-Halogen Atom Substituted gem-Dichromium Reagent 373 Conclusion 375
	9	Polyfunctional Organocopper Reagents for Organic Synthesis 379 Paul Knochel, Xiaoyin Yang, and Nina Gommermann
	9.1 9.2 9.2.1 9.2.2	Introduction 379 Preparation of Functionalized Organocopper Reagents 379 Preparation by the Direct Insertion of Activated Copper 379 Preparation by a Halogen-Copper Exchange Reaction 382

9.2.3	Preparation of Functionalized Copper Reagents Starting from Organolithium Reagents 386
9.2.4	Preparation of Functionalized Alkenylcopper Derivatives Starting from Organozirconium Compounds 389
9.3	Applications of Functionalized Copper Reagents 391
9.4	Conclusion 394
10	Functional Organonickel Reagents 397 Tien-Yau Luh and Li-Fu Huang
10.1	Introduction 397
10.2	Homocoupling Reactions 397
10.3	Cross-coupling Reactions 400
10.3.1	Kumada–Corriu Reactions 401
10.3.2	Negishi Reaction 403
10.3.3	Suzuki Reaction 405
10.3.4	Stille Coupling 407
10.3.5	Heck Reaction 407
10.3.6	Miscellaneous Coupling Reactions 407
10.3.7	Aliphatic Substrates 409
10.4	Carbozincation Reactions 411
10.5	Cycloadditions 413
10.5.1	[2+2] Cycloaddition 413
10.5.2	[4+2] Cycloaddition 414
10.5.3	[4 + 4] Cycloaddition 415
10.5.4	[2 + 2 + 2] Cycloaddition 416
10.5.5	[3+2+2] Cycloaddition 418
10.5.6	[4+2+1] Cycloaddition 419
10.6	Intramolecular Coupling of Enynes or Alkynes 420
10.7	Reactions of Enones with Alkynes 422
10.8	Reaction of Simple Aldehydes or Ketones with Alkynes 429
10.9	Miscellaneous Reactions 436
10.10	Conclusion 443
11	Polyfunctional Metal Carbenes for Organic Synthesis 451 Karl Heinz Dötz, Alexander Koch, and Martin Werner
11.1	Introduction 451
11.2	Chromium-Templated Cycloaddition Reactions 451
11.2.1	Cyclopropanation 452
11.2.2	Benzannulation 455
11.2.3	Cyclization of Chromium Oligoene(-yne) Carbenes 461
11.3	Reactions of Higher Nuclearity Chromium and Tungsten Carbenes 467
11.4	Metathesis Reactions Catalyzed by Group VI and VIII Metal Carbenes 473
11.5	Transmetallation 477

XII	Contents	
1	11.6	Metal Carbenes in Peptide Chemistry 481
	11.7	Stereoselective Syntheses with Sugar Metal Carbenes 483
	11.8	Sugar Metal Carbenes as Organometallic Gelators 495
	11.9	Conclusion 496
	12	Functionalized Organozirconium and Titanium in Organic Synthesis 503 Ilan Marek and Helena Chechik-Lankin
	12.1	Introduction 503
	12.2	Functionalized Organozirconocene Derivatives 503
	12.2.1	Preparation of Functionalized Alkenylzirconocene Derivatives 503
	12.2.2	Preparation of Functionalized Alkylzirconocene Derivatives 511
	12.2.3	Preparation and Reactivity of Acylzirconocene Derivatives 514
	12.2.4	Preparation of Functionalized Low-valent Zirconocene Derivatives 519
	12.3	Functionalized Organotitanium Derivatives 520
	12.3.1	Preparation of Functionalized Substrates via Titanocene
		Derivatives 521
	12.3.1.1	Intramolecular Reductive Cyclization 521
		Allenylation of Functionalized Carbonylic Compounds 525
	12.3.2	Preparation of Functionalized Substrates via Titanium (ii) Alkoxide
		Derivatives 526
	12.3.2.1	Generation of $\eta$ 2-Alkene, $\eta$ 2-Alkyne Complexes and their Utilization as
		Vicinal Dianionic Species 526
	13	Manganese Organometallics for the Chemoselective Synthesis of Polyfunctional Compounds 541 Gérard Cahiez and Florence Mahuteau-Betzer
	13.1	Introduction 541
	13.2	Preparation of Organomanganese Compounds 541
	13.2.1	Preparation of Organomanganese Compounds by
		Transmetallation 541
	13.2.2	Preparation of Organomanganese Compounds from Mn <sup>0</sup> 543
	13.3	1,2-Addition to Aldehydes and Ketones 544
	13.3.1	Chemoselective 1,2-Addition of Organomanganese Reagents to Carbonyl
		Compounds 544
	13.3.2	Manganese-Mediated Barbier- and Reformatsky-like Reactions 547
	13.4	Preparation of Ketones by Acylation of Organomanganese Reagents 548
	13.4.1	Acylation of Organomanganese Reagents 548
	13.4.2	Manganese-Catalyzed Acylation of Grignard Reagents 554

1,4-Addition of Organomanganese Reagents to Enones 555

Transition-Metal-Catalyzed Cross-coupling Reactions 559

Copper-Catalyzed Cross-coupling Reactions 559

Palladium-Catalyzed Cross-coupling Reactions 561

Iron-Catalyzed Cross-coupling Reactions 560

13.5

13.6

13.6.1

13.6.2 13.6.3