# ORGANIC CHEMISTRY

WITH BIOLOGICAL TOPICS

FIFTH EDITION





Janice Gorzynski Smith Heidi R. Vollmer-Snarr

# Periodic Table of the Elements

	<del>-</del>	α	က	4	2	9	
8A	Helium 4.0026	Neon 20.1797	Argon 39.948	36 Krypton 83.80	54 Xenon Xenon 131.29	Radon (222)	118 
	7A	9 Fluorine 18.9984	Chlorine 35.4527	35 <b>Dr</b> Bromine 79.904	53 Iodine 126.9045	At Astatine (210)	117 — — — — — — — — — — — — — — — — — —
	6A	Oxygen 15.9994	32.066	Selenium 78.96	52 Tellurium 127.60	Polonium (209)	Livermorium (293)
	5A	Nitrogen 14.0067	15 Phosphorus 30.9738	33 <b>AS</b> Arsenic 74.9216	Sb Antimony 121.760	83 Bismuth 208.9804	115
	44 4	Carbon 12.011	Silicon 28.0855	Germanium 72.64	SO Tin 118.710	Pb   Lead   207.2	114 Flerovium (289)
	3A	Boron 10.811	13 Aluminum 26.9815	<b>Gallium</b> 69.723	49 Indium 114.82	Thallium 204.3833	113
			2B	30 Zinc 65.41	Oddmium 112.411	HOF WEIGHTY 200.59	Copernicium (285)
			1B	29 Copper 63.546	Ag Silver 107.8682	AU Gold 196.9665	Roentgenium (280)
		ŧ	8B	28 Nickel 58.693	Pd Palladium 106.42	Platinum	Darmstadtium (281)
	Symbol	- Atomic weight	8B	Cobalt 58.9332	45 Rhodium 102.9055	77 <b>  </b>   <b>  </b>   <b> </b>   <b> </b>	Meitnerium (276)
	1	Holmium 164.9303 ← At An element	8B	26 Iron 55.845	Ruthenium	Osmium 190.2	108 Hassium (270)
		Name 164 An e	7B	25 Manganese 54.9380	Technetium (98)	75 <b>Renium</b> 186.207	Bohrium (272)
	Atomic number	Ž	6B	Ortonium 51.9961	42 Molybdenum 95.94	74 Tungsten 183.84	Seaborgium (271)
	∢		5B	23 Vanadium 50.9415	Niobium 92.9064	73 Tantalum 180.9479	105 Db Dubnium (268)
			4B	22 Titanium 47.88	Zirconium 91.224	72 Hafnium 178.49	Pt Rutherfordium (267)
			3B	Scandium 44.9559	39 Yttrium 88.9059	Lanthanum 138.9055	89 <b>AC</b> Actinium (227)
	2A	Beryllium 9.0122	Magnesium 24.3050	20 Calcium 40.078	Strontium 87.62	56 <b>Ba</b> 137.327	<b>Ba</b> Radium (226)
Group number → 1A	Hydrogen 1.0079	3 Lithium 6.941	Na Sodium 22:9898	19 Potassium 39.0983	Rubidium 85.4678	SS Cesium 132.9054	87 Francium (223)
ber -	<u> </u>	N	ო	4	Ŋ	9	7
num (	ا ا						
Group	Period						

· ·	)	
۲	Lutetium 174.967	103 <b>Lr</b> Lawrencium (260)
R Q	Ytterbium 173.04	Nobelium (259)
a H	Thulium 168.9342	Mendelevium (258)
<sub>89</sub> П	Erbium 167.26	Fermium (257)
P 6	Holmium 164.9303	Einsteinium (252)
§	Dysprosium 162.50	Of Californium (251)
Tp Cs	Terbium 158.9253	Berkelium (247)
₽	Gadolinium 157.25	Om Curium (247)
Eu Eu	Europium 151.964	Am Americium (243)
Sm	Samarium 150.36	Putonium (244)
Pa	Promethium (145)	Neptunium (237)
<sup>©</sup> Z	Neodymium 144.24	92 Uranium 238.0289
®₽	Praseodymium 140.9076	Pa Protactinium 231.0359
Seg	Cerium 140.115	90 Thorium 232.0381
<u></u>	1	
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# Organic Chemistry with Biological Topics

Fifth Edition

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University of Hawai'i at Mānoa

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#### ORGANIC CHEMISTRY WITH BIOLOGICAL TOPICS, FIFTH EDITION

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## **Contents in Brief**

	Prologue 1					
1	Structure and Bonding 7					
2	Acids and Bases 61					
3	Introduction to Organic Molecules and Functional Groups 91					
4	Alkanes 134					
5	Stereochemistry 180					
6	Understanding Organic Reactions 219					
7	Alkyl Halides and Nucleophilic Substitution 255					
8	Alkyl Halides and Elimination Reactions 305					
9	Alcohols, Ethers, and Related Compounds 339					
10	Alkenes 391					
11	Alkynes 434					
12	Oxidation and Reduction 463					
13	Mass Spectrometry and Infrared Spectroscopy 503					
14	Nuclear Magnetic Resonance Spectroscopy 535					
15	Radical Reactions 578					
16	Conjugation, Resonance, and Dienes 612					
17	Benzene and Aromatic Compounds 649					
18	Reactions of Aromatic Compounds 686					
19	Carboxylic Acids and the Acidity of the O–H Bond 738					
20	Introduction to Carbonyl Chemistry; Organometallic Reagents; Oxidation and Reduction 774					
21	Aldehydes and Ketones—Nucleophilic Addition 827					
22	Carboxylic Acids and Their Derivatives—Nucleophilic Acyl Substitution 878					
23	Substitution Reactions of Carbonyl Compounds at the $\alpha$ Carbon 934					
24	Carbonyl Condensation Reactions 972					
25	Amines 1010					
<b>26</b>	Amino Acids and Proteins 1063					
<b>27</b>	Carbohydrates 1109					
28	Lipids 1155					
29	Carbon–Carbon Bond-Forming Reactions in Organic Synthesis 1185					
30	Pericyclic Reactions 1212					
31	Synthetic Polymers 1242 (Available online)					
	Appendices A-1					
	Glossary G-1					
	Credits C-1					
	Index I-1					

#### **Contents**

Preface xiii

Acknowledgments xxi

List of How To's xxiii

List of Mechanisms xxiv

List of Selected Applications xxvii

#### Prologue 1

What Is Organic Chemistry? 1

Some Representative Organic Molecules 2

Organic Chemistry and Malaria 4

#### **1** Structure and Bonding

- 1.1 The Periodic Table 8
- **1.2** Bonding 11
- **1.3** Lewis Structures 13
- **1.4** Isomers 18
- **1.5** Exceptions to the Octet Rule 19
- **1.6** Resonance 19
- 1.7 Determining Molecular Shape 25
- **1.8** Drawing Organic Structures 30
- **1.9** Hybridization 36
- 1.10 Ethane, Ethylene, and Acetylene 40
- 1.11 Bond Length and Bond Strength 45
- 1.12 Electronegativity and Bond Polarity 47
- **1.13** Polarity of Molecules 49
- 1.14 L-Dopa—A Representative Organic Molecule 50

  Key Concepts 52

  Problems 53

#### 2 Acids and Bases 61

- **2.1** Brønsted–Lowry Acids and Bases 62
- **2.2** Reactions of Brønsted–Lowry Acids and Bases 63
- 2.3 Acid Strength and p $K_a$  66
- **2.4** Predicting the Outcome of Acid–Base Reactions 68
- 2.5 Factors That Determine Acid Strength 70
- **2.6** Common Acids and Bases 78

#### **2.7** Aspirin 80

2.8 Lewis Acids and Bases 81

Key Concepts 84

Problems 85

# Introduction to OrganicMolecules and FunctionalGroups 91



- 3.1 Functional Groups 92
- 3.2 An Overview of Functional Groups 93
- 3.3 Intermolecular Forces 101
- 3.4 Physical Properties 105
- 3.5 Application: Vitamins 111
- 3.6 Application of Solubility: Soap 112
- 3.7 Application: The Cell Membrane 114
- 3.8 Functional Groups and Reactivity 117
- 3.9 Biomolecules 119

  Key Concepts 125

  Problems 126

#### 4 Alkanes 134

- **4.1** Alkanes—An Introduction 135
- 4.2 Cycloalkanes 138
- **4.3** An Introduction to Nomenclature 138
- **4.4** Naming Alkanes 139
- 4.5 Naming Cycloalkanes 144
- 4.6 Common Names 147
- **4.7** Fossil Fuels 147
- **4.8** Physical Properties of Alkanes 149
- **4.9** Conformations of Acyclic Alkanes—Ethane 150
- 4.10 Conformations of Butane 154
- 4.11 An Introduction to Cycloalkanes 157
- 4.12 Cyclohexane 158
- 4.13 Substituted Cycloalkanes 162
- **4.14** Oxidation of Alkanes 167
- **4.15** Lipids—Part 1 170 Key Concepts 172 Problems 173



#### Stereochemistry 180

- 5.1 Starch and Cellulose 181
- The Two Major Classes of 5.2 Isomers 183
- 5.3 Looking Glass Chemistry—Chiral and Achiral Molecules 184
- 5.4 Stereogenic Centers 187
- 5.5 Stereogenic Centers in Cyclic Compounds 189
- Labeling Stereogenic Centers with R or S 5.6
- 5.7 Diastereomers 196
- 5.8 Meso Compounds 199
- R and S Assignments in Compounds with Two or 5.9 More Stereogenic Centers 200
- 5.10 Disubstituted Cycloalkanes 201
- 5.11 Isomers—A Summary 202
- 5.12 Physical Properties of Stereoisomers
- **5.13** Chemical Properties of Enantiomers 208 Key Concepts 210 Problems 211

#### 6 Understanding Organic Reactions 219

- 6.1 Writing Equations for Organic Reactions 220
- 6.2 Kinds of Organic Reactions 221
- 6.3 Bond Breaking and Bond Making 223
- 6.4 Bond Dissociation Energy 227
- 6.5 Thermodynamics 230
- Enthalpy and Entropy 235 6.6
- 6.7 Energy Diagrams 236
- 6.8 Energy Diagram for a Two-Step Reaction Mechanism 239
- 6.9 Kinetics 241
- 6.10 Catalysts 244
- Enzymes 245 6.11 Key Concepts 247 Problems 248

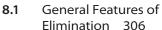
#### 7 Alkyl Halides and Nucleophilic Substitution 255

- 7.1 Introduction to Alkyl Halides 256
- 7.2 Nomenclature 257
- 7.3 Physical Properties 258



- 7.4 Interesting Alkyl Halides 259
- 7.5 The Polar Carbon–Halogen Bond 260
- 7.6 General Features of Nucleophilic Substitution 261
- 7.7 The Leaving Group 263
- 7.8 The Nucleophile 265
- 7.9 Possible Mechanisms for Nucleophilic Substitution 269
- 7.10 Two Mechanisms for Nucleophilic Substitution 270
- 7.11 The S<sub>N</sub>2 Mechanism 271
- 7.12 The S<sub>N</sub>1 Mechanism 277
- 7.13 Carbocation Stability 281
- 7.14 The Hammond Postulate 283
- 7.15 When Is the Mechanism  $S_N 1$  or  $S_N 2$ ?
- 7.16 Biological Nucleophilic Substitution 291
- 7.17 Vinyl Halides and Aryl Halides 294
- 7.18 Organic Synthesis 294 Key Concepts 296 Problems 298

#### **Alkyl Halides** and Elimination Reactions 305



- 8.2 Alkenes—The Products of Elimination Reactions 307
- 8.3 The Mechanisms of Elimination 311
- The E2 Mechanism 311 8.4
- 8.5 The Zaitsev Rule 316
- 8.6 The E1 Mechanism 318
- 8.7 S<sub>N</sub>1 and E1 Reactions 321
- Stereochemistry of the E2 Reaction 322 8.8
- 8.9 When Is the Mechanism E1 or E2? 325
- 8.10 E2 Reactions and Alkyne Synthesis 326
- 8.11 When Is the Reaction  $S_N1$ ,  $S_N2$ , E1, or E2? 327 Key Concepts 331 Problems 333

#### Alcohols, Ethers, and **Related Compounds** 339

- 9.1 Introduction 340
- 9.2 Structure and Bonding 341
- 9.3 Nomenclature 342
- 9.4 Physical Properties 345







9.5	Interesting Alcohols, Ethers, and Epoxides 346			
9.6	Preparation of Alcohols, Ethers, and Epoxides 349			
9.7	General Features—Reactions of Alcohols, Ethers, and Epoxides 351			
9.8	Dehydration of Alcohols to Alkenes 353			
9.9	Carbocation Rearrangements 356			
9.10	Dehydration Using POCl₃ and Pyridine 359			
9.11	Conversion of Alcohols to Alkyl Halides with HX 360			
9.12	Conversion of Alcohols to Alkyl Halides with $SOCl_2$ and $PBr_3$ 364			
9.13	Tosylate—Another Good Leaving Group 367			
9.14	Reaction of Ethers with Strong Acid 370			
9.15	Thiols and Sulfides 372			
9.16	Reactions of Epoxides 375			
9.17	Application: Epoxides, Leukotrienes, and Asthma 379			
9.18	Application: Benzo[a]pyrene, Epoxides, and Cancer 381			
	Key Concepts 381			
	Problems 384			
10	Alkenes 391			
<b>10</b> 10.1	Alkenes 391 Introduction 392			
10.1	Introduction 392			
10.1	Introduction 392 Calculating Degrees of			
10.1 10.2	Introduction 392 Calculating Degrees of Unsaturation 393			
10.1 10.2 10.3	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395			
10.1 10.2 10.3 10.4	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399			
10.1 10.2 10.3 10.4 10.5	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399			
10.1 10.2 10.3 10.4 10.5 10.6	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401			
10.1 10.2 10.3 10.4 10.5 10.6	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403			
10.1 10.2 10.3 10.4 10.5 10.6 10.7	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403 Introduction to Addition Reactions 404 Hydrohalogenation—Electrophilic Addition of HX 405			
10.1 10.2 10.3 10.4 10.5 10.6 10.7 10.8 10.9	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403 Introduction to Addition Reactions 404 Hydrohalogenation—Electrophilic Addition of HX 405			
10.1 10.2 10.3 10.4 10.5 10.6 10.7 10.8 10.9	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403 Introduction to Addition Reactions 404 Hydrohalogenation—Electrophilic Addition of HX 405 Markovnikov's Rule 408 Stereochemistry of Electrophilic Addition of HX 410			
10.1 10.2 10.3 10.4 10.5 10.6 10.7 10.8 10.9	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403 Introduction to Addition Reactions 404 Hydrohalogenation—Electrophilic Addition of HX 405 Markovnikov's Rule 408 Stereochemistry of Electrophilic Addition of HX 410 Hydration—Electrophilic Addition of Water 412			
10.1 10.2 10.3 10.4 10.5 10.6 10.7 10.8 10.9 10.10 10.11	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403 Introduction to Addition Reactions 404 Hydrohalogenation—Electrophilic Addition of HX 405 Markovnikov's Rule 408 Stereochemistry of Electrophilic Addition of HX 410 Hydration—Electrophilic Addition of Water 412			
10.1 10.2 10.3 10.4 10.5 10.6 10.7 10.8 10.9 10.10 10.11 10.12 10.13	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403 Introduction to Addition Reactions 404 Hydrohalogenation—Electrophilic Addition of HX 405 Markovnikov's Rule 408 Stereochemistry of Electrophilic Addition of HX 410 Hydration—Electrophilic Addition of Water 412 Halogenation—Addition of Halogen 413			
10.1 10.2 10.3 10.4 10.5 10.6 10.7 10.8 10.9 10.10 10.11 10.12 10.13 10.14	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403 Introduction to Addition Reactions 404 Hydrohalogenation—Electrophilic Addition of HX 405 Markovnikov's Rule 408 Stereochemistry of Electrophilic Addition of HX 410 Hydration—Electrophilic Addition of Water 412 Halogenation—Addition of Halogen 413 Stereochemistry of Halogenation 414 Halohydrin Formation 416			
10.1 10.2 10.3 10.4 10.5 10.6 10.7 10.8 10.9 10.10 10.11 10.12 10.13 10.14 10.15	Introduction 392 Calculating Degrees of Unsaturation 393 Nomenclature 395 Physical Properties 399 Interesting Alkenes 399 Lipids—Part 2 401 Preparation of Alkenes 403 Introduction to Addition Reactions 404 Hydrohalogenation—Electrophilic Addition of HX 405 Markovnikov's Rule 408 Stereochemistry of Electrophilic Addition of HX 410 Hydration—Electrophilic Addition of Water 412 Halogenation—Addition of Halogen 413 Stereochemistry of Halogenation 414 Halohydrin Formation 416			

Key Concepts 426

Problems 427

# **11** Alkynes 434 11.1 Introduction 435

11.2 Nomenclature 436

11.3 Physical Properties 437

11.4 Interesting Alkynes 438

11.5 Preparation of Alkynes 439

11.6 Introduction to Alkyne Reactions 440

11.7 Addition of Hydrogen Halides 442

11.8 Addition of Halogen 444

11.9 Addition of Water 445

11.10 Hydroboration-Oxidation 447

11.11 Reaction of Acetylide Anions 449

11.12 Synthesis 452 Key Concepts 455 Problems 456

# **12** Oxidation and Reduction 463



12.2 Reducing Agents 465

12.3 Reduction of Alkenes 466

12.4 Application: Hydrogenation of Oils 469

12.5 Reduction of Alkynes 471

**12.6** The Reduction of Polar C – X  $\sigma$  Bonds 474

12.7 Oxidizing Agents 475

12.8 Epoxidation 477

**12.9** Dihydroxylation 480

12.10 Oxidative Cleavage of Alkenes 482

12.11 Oxidative Cleavage of Alkynes 484

12.12 Oxidation of Alcohols 484

12.13 Green Chemistry 487

12.14 Biological Oxidation 489

12.15 Sharpless Epoxidation 490

Key Concepts 493

Problems 495

# 13 Mass Spectrometry and Infrared Spectroscopy 503



13.1 Mass Spectrometry 504

13.2 Alkyl Halides and the M + 2 Peak 508

**13.3** Fragmentation 509

**13.4** Other Types of Mass Spectrometry 512



13.5 13.6 13.7 13.8	Electromagnetic Radiation 514 Infrared Spectroscopy 516 IR Absorptions 518 IR and Structure Determination 525	15.14	P K P
	Key Concepts 527 Problems 528	16	C
	Trodicina 525		a
		16.1	C
14	Nuclear Magnetic	16.2	R
	Resonance		
	Spectroscopy 535	16.3 16.4	T
14.1	An Introduction to NMR	16.4	E
14.2	Spectroscopy 536 <sup>1</sup> H NMR: Number of Signals 539	10.5	G
14.3	<sup>1</sup> H NMR: Position of Signals 543	16.6	C
14.4	The Chemical Shift of Protons on sp <sup>2</sup> and	16.7	lr
	sp Hybridized Carbons 547	16.8	T B
14.5	<sup>1</sup> H NMR: Intensity of Signals 549	16.9	S
14.6 14.7	<sup>1</sup> H NMR: Spin–Spin Splitting 550 More Complex Examples of Splitting 554	16.10	
14.7	Spin–Spin Splitting in Alkenes 557		1
14.9	Other Facts About <sup>1</sup> H NMR Spectroscopy 559	16.11	K
14.10	Using <sup>1</sup> H NMR to Identify an Unknown 561	16.12	
14.11	<sup>13</sup> C NMR Spectroscopy 564	16.13	S
14.12	Magnetic Resonance Imaging (MRI) 568	16.14	
	Key Concepts 569	16.15	
	Problems 569		Κ
			Ρ
15	Radical Reactions 578		
15.1	Introduction 579	<b>17</b>	В
15.2	General Features of Radical		C
15.3	Reactions 580	17.1	В
15.4	Halogenation of Alkanes 582 The Mechanism of Halogenation 583	17.2	Т
15.5	Chlorination of Other Alkanes 586	17.3	Ν
15.6	Chlorination Versus Bromination 586		
15.7	Halogenation as a Tool in Organic Synthesis 589	17.4	S
15.8	The Stereochemistry of Halogenation	17.5 17.6	B T
	Reactions 590	17.0	E
15.9	Application: The Ozone Layer and CFCs 592	17.8	Α
15.10 15.11	Radical Halogenation at an Allylic Carbon 593 Application: Oxidation of Unsaturated	17.9	٧
13.11	Lipids 596	17.10	Т
15.12	Application: Antioxidants 597		Α
	Radical Addition Reactions to Double	17.11	A
	Bonds 598		E

#### **15.14** Polymers and Polymerization 601 *Key Concepts* 603 *Problems* 604

# 16 Conjugation, Resonance, and Dienes 612



**16.2** Resonance and Allylic Carbocations 615

- 16.3 Common Examples of Resonance 616
- 16.4 The Resonance Hybrid 618
- **16.5** Electron Delocalization, Hybridization, and Geometry 620
- **16.6** Conjugated Dienes 621
- 16.7 Interesting Dienes and Polyenes 622
- **16.8** The Carbon–Carbon σ Bond Length in Buta-1,3-diene 622
- **16.9** Stability of Conjugated Dienes 623
- **16.10** Electrophilic Addition: 1,2- Versus 1,4-Addition 624
- **16.11** Kinetic Versus Thermodynamic Products 626
- 16.12 The Diels-Alder Reaction 629
- **16.13** Specific Rules Governing the Diels–Alder Reaction 631
- **16.14** Other Facts About the Diels–Alder Reaction 635
- 16.15 Conjugated Dienes and Ultraviolet Light 638Key Concepts 640Problems 642

# 17 Benzene and Aromatic Compounds 649



- 17.2 The Structure of Benzene 651
- 17.3 Nomenclature of Benzene Derivatives 653
- **17.4** Spectroscopic Properties 655
- 17.5 Benzene's Unusual Stability 656
- 17.6 The Criteria for Aromaticity—Hückel's Rule 657
- **17.7** Examples of Aromatic Compounds 660
- 17.8 Aromatic Heterocycles 664
- 17.9 What Is the Basis of Hückel's Rule? 669
- **17.10** The Inscribed Polygon Method for Predicting Aromaticity 672
- 17.11 Application: Aromatase Inhibitors for Estrogen-Dependent Cancer Treatment 674Key Concepts 676Problems 677



# **18** Reactions of Aromatic Compounds 686

- **18.1** Electrophilic Aromatic Substitution 687
- 18.2 The General Mechanism 688
- 18.3 Halogenation 690
- 18.4 Nitration and Sulfonation 691
- **18.5** Friedel–Crafts Alkylation and Friedel–Crafts Acylation 693
- 18.6 Substituted Benzenes 700
- **18.7** Electrophilic Aromatic Substitution of Substituted Benzenes 703
- **18.8** Why Substituents Activate or Deactivate a Benzene Ring 705
- **18.9** Orientation Effects in Substituted Benzenes 707
- **18.10** Limitations on Electrophilic Substitution
  Reactions with Substituted Benzenes 710
- 18.11 Disubstituted Benzenes 712
- 18.12 Synthesis of Benzene Derivatives 714
- **18.13** Nucleophilic Aromatic Substitution 715
- 18.14 Halogenation of Alkyl Benzenes 718
- **18.15** Oxidation and Reduction of Substituted Benzenes 720
- **18.16** Multistep Synthesis 724 *Key Concepts 727 Problems 730*

#### 19 Carboxylic Acids and the Acidity of the O-H Bond 738

- **19.1** Structure and Bonding 739
- 19.2 Nomenclature 739
- 19.3 Physical Properties 742
- 19.4 Spectroscopic Properties 743
- **19.5** Interesting Carboxylic Acids 745
- **19.6** Aspirin, Arachidonic Acid, and Prostaglandins 745
- **19.7** Preparation of Carboxylic Acids 747
- **19.8** Reactions of Carboxylic Acids—General Features 748
- **19.9** Carboxylic Acids—Strong Organic Brønsted–Lowry Acids 749
- **19.10** The Henderson–Hasselbalch Equation 752
- **19.11** Inductive Effects in Aliphatic Carboxylic Acids 754
- 19.12 Substituted Benzoic Acids 756



- **19.13** Extraction 758
- **19.14** Organic Acids Containing Sulfur and Phosphorus 760
- **19.15** Amino Acids 761 Key Concepts 765 Problems 766

# 20 Introduction to Carbonyl Chemistry; Organometallic Reagents; Oxidation and Reduction 774



- 20.1 Introduction 775
- 20.2 General Reactions of Carbonyl Compounds 776
- **20.3** A Preview of Oxidation and Reduction 779
- 20.4 Reduction of Aldehydes and Ketones 781
- **20.5** The Stereochemistry of Carbonyl Reduction 783
- 20.6 Enantioselective Carbonyl Reductions 784
- **20.7** Reduction of Carboxylic Acids and Their Derivatives 787
- 20.8 Oxidation of Aldehydes 792
- 20.9 Organometallic Reagents 792
- **20.10** Reaction of Organometallic Reagents with Aldehydes and Ketones 796
- **20.11** Retrosynthetic Analysis of Grignard Products 800
- 20.12 Protecting Groups 802
- **20.13** Reaction of Organometallic Reagents with Carboxylic Acid Derivatives 804
- **20.14** Reaction of Organometallic Reagents with Other Compounds 807
- **20.15** α,β-Unsaturated Carbonyl Compounds 809
- **20.16** Summary—The Reactions of Organometallic Reagents 812
- 20.17 Synthesis 812 Key Concepts 815 Problems 818

#### 21 Aldehydes and Ketones—Nucleophilic Addition 827

- 000
- 21.1 Introduction 828
- 21.2 Nomenclature 829
- 21.3 Physical Properties 832
- 21.4 Spectroscopic Properties 833
- 21.5 Interesting Aldehydes and Ketones 835

Х	Contents
21.6	Preparation of Aldehydes and Ketones 836
21.7	Reactions of Aldehydes and Ketones— General Considerations 838
21.8	Nucleophilic Addition of H <sup>-</sup> and R <sup>-</sup> —A Review 841
21.9	Nucleophilic Addition of <sup>-</sup> CN 843
21.10	The Wittig Reaction 845
21.11	Addition of 1° Amines 850
	Addition of 2° Amines 852
	Addition of H <sub>2</sub> O—Hydration 854
21.14	Addition of Alcohols—Acetal Formation 857
21.15	Acetals as Protecting Groups 861
	Cyclic Hemiacetals 862
21.17	An Introduction to Carbohydrates 865
	Key Concepts 866
	Problems 868
22	Carboxylic Acids and Their Derivatives— Nucleophilic Acyl
	- State Koloni (act
<b>22 1</b>	Substitution 878
22.1	Substitution 878 Introduction 879
22.2	Substitution 878
22.2 22.3	Substitution 878  Introduction 879  Structure and Bonding 881  Nomenclature 883
22.2	Substitution 878 Introduction 879 Structure and Bonding 881 Nomenclature 883 Physical Properties 888
22.2 22.3 22.4	Substitution 878  Introduction 879  Structure and Bonding 881  Nomenclature 883
22.2 22.3 22.4 22.5	Substitution 878  Introduction 879  Structure and Bonding 881  Nomenclature 883  Physical Properties 888  Spectroscopic Properties 889
22.2 22.3 22.4 22.5 22.6 22.7	Substitution 878  Introduction 879  Structure and Bonding 881  Nomenclature 883  Physical Properties 888  Spectroscopic Properties 889  Interesting Esters and Amides 891  Introduction to Nucleophilic Acyl  Substitution 892
22.2 22.3 22.4 22.5 22.6	Substitution 878  Introduction 879  Structure and Bonding 881  Nomenclature 883  Physical Properties 888  Spectroscopic Properties 889  Interesting Esters and Amides 891  Introduction to Nucleophilic Acyl Substitution 892  Reactions of Acid Chlorides 896
22.2 22.3 22.4 22.5 22.6 22.7 22.8 22.9	Substitution 878  Introduction 879  Structure and Bonding 881  Nomenclature 883  Physical Properties 888  Spectroscopic Properties 889  Interesting Esters and Amides 891  Introduction to Nucleophilic Acyl Substitution 892  Reactions of Acid Chlorides 896  Reactions of Anhydrides 897
22.2 22.3 22.4 22.5 22.6 22.7 22.8 22.9 22.10	Substitution 878  Introduction 879  Structure and Bonding 881  Nomenclature 883  Physical Properties 888  Spectroscopic Properties 889  Interesting Esters and Amides 891  Introduction to Nucleophilic Acyl Substitution 892  Reactions of Acid Chlorides 896  Reactions of Anhydrides 897  Reactions of Carboxylic Acids 898
22.2 22.3 22.4 22.5 22.6 22.7 22.8 22.9 22.10 22.11	Substitution 878  Introduction 879  Structure and Bonding 881  Nomenclature 883  Physical Properties 888  Spectroscopic Properties 889  Interesting Esters and Amides 891  Introduction to Nucleophilic Acyl Substitution 892  Reactions of Acid Chlorides 896  Reactions of Anhydrides 897  Reactions of Carboxylic Acids 898  Reactions of Esters 903
22.2 22.3 22.4 22.5 22.6 22.7 22.8 22.9 22.10 22.11 22.12	Substitution 878  Introduction 879 Structure and Bonding 881 Nomenclature 883 Physical Properties 888 Spectroscopic Properties 889 Interesting Esters and Amides 891 Introduction to Nucleophilic Acyl Substitution 892 Reactions of Acid Chlorides 896 Reactions of Anhydrides 897 Reactions of Carboxylic Acids 898 Reactions of Esters 903 Application: Lipid Hydrolysis 905
22.2 22.3 22.4 22.5 22.6 22.7 22.8 22.9 22.10 22.11 22.12 22.13	Introduction 879 Structure and Bonding 881 Nomenclature 883 Physical Properties 888 Spectroscopic Properties 889 Interesting Esters and Amides 891 Introduction to Nucleophilic Acyl Substitution 892 Reactions of Acid Chlorides 896 Reactions of Anhydrides 897 Reactions of Carboxylic Acids 898 Reactions of Esters 903 Application: Lipid Hydrolysis 905 Reactions of Amides 908
22.2 22.3 22.4 22.5 22.6 22.7 22.8 22.9 22.10 22.11 22.12 22.13 22.14	Substitution 878  Introduction 879 Structure and Bonding 881 Nomenclature 883 Physical Properties 888 Spectroscopic Properties 889 Interesting Esters and Amides 891 Introduction to Nucleophilic Acyl Substitution 892 Reactions of Acid Chlorides 896 Reactions of Anhydrides 897 Reactions of Carboxylic Acids 898 Reactions of Esters 903 Application: Lipid Hydrolysis 905 Reactions of Amides 908 Application: The Mechanism of Action of β-Lactam Antibiotics 909
22.2 22.3 22.4 22.5 22.6 22.7 22.8 22.9 22.10 22.11 22.12 22.13 22.14	Introduction 879 Structure and Bonding 881 Nomenclature 883 Physical Properties 888 Spectroscopic Properties 889 Interesting Esters and Amides 891 Introduction to Nucleophilic Acyl Substitution 892 Reactions of Acid Chlorides 896 Reactions of Anhydrides 897 Reactions of Carboxylic Acids 898 Reactions of Esters 903 Application: Lipid Hydrolysis 905 Reactions of Amides 908 Application: The Mechanism of Action
22.2 22.3 22.4 22.5 22.6 22.7 22.8 22.9 22.10 22.11 22.12 22.13 22.14	Substitution 878  Introduction 879 Structure and Bonding 881 Nomenclature 883 Physical Properties 888 Spectroscopic Properties 889 Interesting Esters and Amides 891 Introduction to Nucleophilic Acyl Substitution 892 Reactions of Acid Chlorides 896 Reactions of Anhydrides 897 Reactions of Carboxylic Acids 898 Reactions of Esters 903 Application: Lipid Hydrolysis 905 Reactions of Amides 908 Application: The Mechanism of Action of β-Lactam Antibiotics 909 Summary of Nucleophilic Acyl Substitution

Acylation Reactions 914

Key Concepts 921

Problems 924

**22.18** Nitriles 916

#### **23** Substitution Reactions of Carbonyl Compounds at the $\alpha$ Carbon 934



- Introduction 935
- 23.2 Enols 936
- Enolates 938 23.3
- **Enolates of Unsymmetrical Carbonyl** Compounds 944
- Racemization at the  $\alpha$  Carbon 946 23.5
- **23.6** A Preview of Reactions at the  $\alpha$  Carbon 947
- **23.7** Halogenation at the  $\alpha$  Carbon 947
- 23.8 Direct Enolate Alkylation 952
- 23.9 Malonic Ester Synthesis 955
- 23.10 Acetoacetic Ester Synthesis 959 Key Concepts 962 Problems 963

#### **24** Carbonyl Condensation Reactions 972



- The Aldol Reaction 973
- 24.2 Crossed Aldol Reactions 978
- 24.3 Directed Aldol Reactions 981
- 24.4 Intramolecular Aldol Reactions 984
- 24.5 The Claisen Reaction 986
- The Crossed Claisen and Related Reactions 987
- 24.7 The Dieckmann Reaction 990
- **Biological Carbonyl Condensation** 24.8 Reactions 991
- 24.9 The Michael Reaction 994
- 24.10 The Robinson Annulation 996 Key Concepts 1000 Problems 1001

#### 25 Amines 1010

- 25.1 Introduction 1011
- Structure and Bonding 1011 25.2
- 25.3 Nomenclature 1013
- 25.4 Physical Properties 1015
- 25.5 Spectroscopic Properties 1016
- Interesting and Useful Amines
- 25.7 Preparation of Amines 1021
- Reactions of Amines—General Features 1028 25.8

- 25.9 Amines as Bases 1028
- **25.10** Relative Basicity of Amines and Other Compounds 1030
- 25.11 Amines as Nucleophiles 1036
- 25.12 Hofmann Elimination 1038
- 25.13 Reaction of Amines with Nitrous Acid 1041
- **25.14** Substitution Reactions of Aryl Diazonium Salts 1043
- **25.15** Coupling Reactions of Aryl Diazonium Salts 1048
- 25.16 Application: Synthetic Dyes and Sulfa Drugs 1050 Key Concepts 1052 Problems 1055

# 26 Amino Acids and Proteins 1063

- 26.1 Amino Acids 1064
- 26.2 Synthesis of Amino Acids 1067
- 26.3 Separation of Amino Acids 1070
- 26.4 Enantioselective Synthesis of Amino Acids 1074
- **26.5** Peptides 1075
- 26.6 Peptide Sequencing 1080
- 26.7 Peptide Synthesis 1083
- 26.8 Automated Peptide Synthesis 1088
- 26.9 Protein Structure 1090
- **26.10** Important Proteins 1097

Key Concepts 1100 Problems 1102

#### 27 Carbohydrates 1109

- 27.1 Introduction 1110
- 27.2 Monosaccharides 1111
- 27.3 The Family of D-Aldoses 1116
- 27.4 The Family of D-Ketoses 1118
- 27.5 Physical Properties of Monosaccharides 1119
- **27.6** The Cyclic Forms of Monosaccharides 1119
- **27.7** Glycosides 1127
- **27.8** Reactions of Monosaccharides at the OH Groups 1130
- **27.9** Reactions at the Carbonyl Group—Oxidation and Reduction 1131
- **27.10** Reactions at the Carbonyl Group—Adding or Removing One Carbon Atom 1134



27.12 Polysaccharides 1141

Problems 1150

27.13 Other Important Sugars and Their Derivatives 1143Key Concepts 1147

#### **28** Lipids 1155

- 28.1 Introduction 1156
- 28.2 Waxes 1157
- 28.3 Triacylglycerols 1158
- 28.4 Phospholipids 1162
- 28.5 Fat-Soluble Vitamins 1165
- 28.6 Eicosanoids 1166
- **28.7** Terpenes 1169
- 28.8 Steroids 1174

  Key Concepts 1179

  Problems 1180

#### 29 Carbon–Carbon Bond-Forming Reactions in Organic Synthesis 1185

- **29.1** Coupling Reactions of Organocuprate Reagents 1186
- 29.2 Suzuki Reaction 1188
- 29.3 Heck Reaction 1192
- 29.4 Carbenes and Cyclopropane Synthesis 1194
- 29.5 Simmons-Smith Reaction 1197
- 29.6 Metathesis 1198

  Key Concepts 1203

  Problems 1204

#### 30 Pericyclic Reactions 1212

- **30.1** Types of Pericyclic Reactions 1213
- **30.2** Molecular Orbitals 1214
- 30.3 Electrocyclic Reactions 1217
- 30.4 Cycloaddition Reactions 1223
- 30.5 Sigmatropic Rearrangements 1227
- **30.6** Summary of Rules for Pericyclic Reactions 1233 *Key Concepts* 1234 *Problems* 1235







# 31 Synthetic Polymers (Available online) 1242

- **31.1** Introduction 1243
- **31.2** Chain-Growth Polymers—Addition Polymers 1244
- 31.3 Anionic Polymerization of Epoxides 1251
- **31.4** Ziegler–Natta Catalysts and Polymer Stereochemistry 1252
- 31.5 Natural and Synthetic Rubbers 1254
- **31.6** Step-Growth Polymers—Condensation Polymers 1255
- **31.7** Polymer Structure and Properties 1260
- 31.8 Green Polymer Synthesis 1261
- 31.9 Polymer Recycling and Disposal 1264

  Key Concepts 1267

  Problems 1268



**Appendix A** pK<sub>a</sub> Values for Selected Compounds A-1

Appendix B Nomenclature A-3

**Appendix C** Bond Dissociation Energies for Some Common Bonds  $[A-B \rightarrow A \cdot + \cdot B]$  A-7

**Appendix D** Reactions That Form Carbon–Carbon Bonds A-8

**Appendix E** Characteristic IR Absorption Frequencies A-9

Appendix F Characteristic NMR Absorptions A-10

**Appendix G** General Types of Organic Reactions A-12

**Appendix H** How to Synthesize Particular Functional Groups A-14

Glossary G-1 Credits C-1

Index I-1

#### **Preface**

Since the publication of *Organic Chemistry* in 2005, chemistry has witnessed a rapid growth in its understanding of the biological world. The molecular basis of many complex biological processes is now known with certainty, and can be explained by applying the basic principles of organic chemistry. Because of the close relationship between chemistry and many biological phenomena, *Organic Chemistry with Biological Topics* presents an approach to traditional organic chemistry that incorporates the discussion of biological applications that are understood using the fundamentals of organic chemistry.

#### The Basic Features

Organic Chemistry with Biological Topics continues the successful student-oriented approach used in Organic Chemistry by Janice Gorzynski Smith. This text uses less prose and more diagrams and bulleted summaries for today's students, who rely more heavily on visual imagery to learn than ever before. Each topic is broken down into small chunks of information that are more manageable and easily learned. Sample Problems illustrate stepwise problem solving, and relevant examples from everyday life are used to illustrate topics. New concepts are introduced one at a time so that the basic themes are kept in focus.

The organization of *Organic Chemistry with Biological Topics* provides the student with a logical and accessible approach to an intense and fascinating subject. The text begins with a healthy dose of review material in Chapters 1 and 2 to ensure that students have a firm grasp of the fundamentals. Stereochemistry, the three-dimensional structure of molecules, is introduced early (Chapter 5) and reinforced often. Certain reaction types with unique characteristics and terminology are grouped together. These include acid—base reactions (Chapter 2), oxidation and reduction (Chapters 12 and 20), radical reactions (Chapter 15), and reactions of organometallic reagents (Chapter 20). Each chapter ends with Key Concepts, end-of-chapter summaries that succinctly organize the main concepts and reactions.

#### New to Organic Chemistry with Biological Topics

While there is no shortage of biological applications that can be added to an organic chemistry text, we have chosen to concentrate on the following areas.

- Chapter 3 on functional groups now includes an expanded section on four types of biomolecules—amino acids and proteins, monosaccharides and carbohydrates, nucleotides and nucleic acids, and lipids. This material augments the discussions of vitamins and the cell membrane, topics already part of *Organic Chemistry* in past editions. Phosphorus-containing compounds such as ATP (adenosine triphosphate), the key intermediate used in energy transfer in cells, are also introduced in this chapter.
- Chapter 6 now uses biological examples to illustrate the basic types of organic reactions, and the energetics of coupled reactions in metabolism is presented. The discussion of enzymes as biological catalysts is expanded, and a specific example of an enzyme's active site is shown.
- Chapter 17 now applies the discussion of aromatic heterocycles to the bases in DNA, the high molecular weight molecule that holds the encrypted genetic instructions for our development and cellular processes. In addition, new material has been added on the synthesis of female sex hormones with the aromatase enzyme, which has resulted in the development of drugs used to treat estrogen-dependent breast cancers.

- Chapter 19 contains a section on the Henderson–Hasselbalch equation, a mathematical expression that allows us to tell whether a compound exists as an uncharged compound or ion at the cellular pH of 7.4. A section on phosphoric acid esters has been added, and the ionization of amino acids is now explained using the Henderson–Hasselbalch equation.
- Chapter 22 contains additional material on two common carboxylic acid derivatives—acyl phosphates and thioesters. The role of these functional groups in the biosynthesis of amino acids and the metabolism of fatty acids is discussed.
- Chapter 24 contains a new section on biological carbonyl condensation reactions. Topics
  include the biological aldol reaction in the citric acid cycle, the retro-aldol reaction in the
  metabolism of glucose, and the biological Claisen reaction in the biosynthesis of fatty acids.

In addition, the later chapters of the text are now reorganized to emphasize the connection of biomolecules to prior sections. The chapter on Amino Acids and Proteins (Chapter 26) now directly follows the chapter on Amines (Chapter 25), followed by the remaining chapters on biomolecules, Carbohydrates (Chapter 27) and Lipids (Chapter 28).

#### **Tools to Make Learning Organic Chemistry Easier**

#### Illustrations

Organic Chemistry with Biological Topics is supported by a well-developed illustration program. Besides traditional skeletal (line) structures and condensed formulas, there are numerous ball-and-stick molecular models and electrostatic potential maps to help students grasp the three-dimensional structure of molecules (including stereochemistry) and to better understand the distribution of electronic charge.

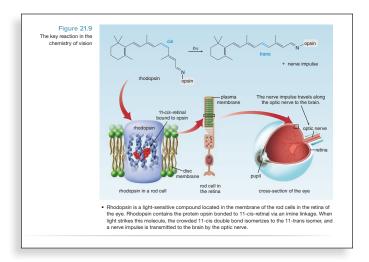


Figure 12.4 Partial hydrogenation of the double bonds in a vegetable of

#### Micro-to-Macro Illustrations

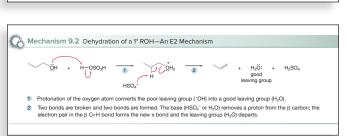
Unique to *Organic Chemistry with Biological Topics* are microto-macro illustrations, where line art and photos combine with chemical structures to reveal the underlying molecular structures giving rise to macroscopic properties of common phenomena. Examples include starch and cellulose (Chapter 5), adrenaline (Chapter 7), partial hydrogenation of vegetable oil (Chapter 12), and dopamine (Chapter 25).

# Spectra

Over 100 spectra created specifically for *Organic Chemistry* with *Biological Topics* are presented throughout the text. The spectra are color-coded by type and generously labeled. Mass spectra are green; infrared spectra are red; and proton and carbon nuclear magnetic resonance spectra are blue.

#### **Mechanisms**

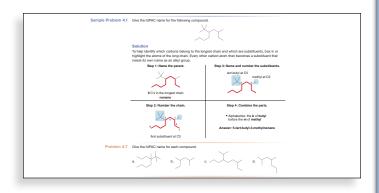
Curved arrow notation is used extensively to help students follow the movement of electrons in reactions.



#### **Problem Solving**

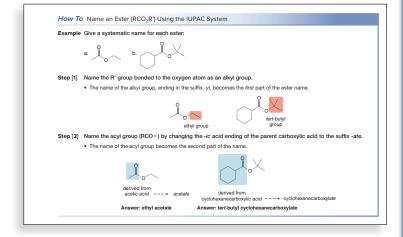
#### **Sample Problems**

Sample Problems show students how to solve organic chemistry problems in a logical, stepwise manner. More than 800 follow-up problems are located throughout the chapters to test whether students understand concepts covered in the Sample Problems.



#### How To's

*How To*'s provide students with detailed instructions on how to work through key processes.



# **Applications and Summaries Key Concept Summaries**

Succinct summary tables reinforcing important principles and concepts are provided at the end of each chapter.

#### **Margin Notes**

Margin notes are placed carefully throughout the chapters, providing interesting information relating to topics covered in the text. Some margin notes are illustrated with photos to make the chemistry more relevant.



All soaps are salts of fatty acids. The main difference between soaps is the addition of other ingredients that do not alter their cleaning properties: dyes for color, scents for a pleasing odor, and oils for lubrication. Soaps that float are aerated, so that they are less dense than water.

# 



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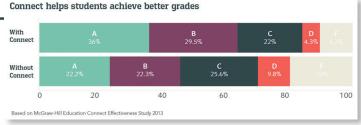
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Within the Instructor's Presentation Tools, instructors have access to editable PowerPoint lecture outlines, which appear as ready-made presentations that combine art and lecture notes for each chapter of the text. For instructors who prefer to create their lecture notes from scratch, all illustrations, photos, tables, *How To*'s, and Sample Problems are pre-inserted by chapter into a separate set of PowerPoint slides. They are also available as individual .jpg files.

Photos, artwork, animations, and other media types can be used to create customized lectures, visually enhanced tests and quizzes, compelling course websites, or attractive printed support materials. All assets are copyrighted by McGraw-Hill Higher Education, but can be used by instructors for classroom purposes. The visual resources in this collection include:

- **Art** Full-color digital files of all illustrations in the book can be readily incorporated into lecture presentations, exams, or custom-made classroom materials.
- **Photos** The photo collection contains digital files of photographs from the text, which can be reproduced for multiple classroom uses.
- **Tables** Every table that appears in the text has been saved in electronic form for use in classroom presentations and/or quizzes.
- **Animations** Full-color animations illustrating important processes are also provided. Harness the visual impact of concepts in motion by importing these files into classroom presentations or online course materials.

#### **Student Study Guide/Solutions Manual**

Written by Janice Gorzynski Smith and Erin R. Smith, the Student Study Guide/Solutions Manual provides step-by-step solutions to all in-chapter and end-of-chapter problems. Each chapter begins with an overview of key concepts and includes a short-answer practice test on the fundamental principles and new reactions.

#### **Acknowledgments**

Organic Chemistry with Biological Topics is an outgrowth of many fruitful discussions with McGraw-Hill personnel about how best to meld biological applications with basic organic chemistry. Special thanks go to Brand Manager Andrea Pellerito, an organic chemist with extensive teaching experience, who understood the need to maintain the integrity and rigor of organic chemistry in this approach, and devised a method to bring this plan to reality.

Special thanks are also due to Senior Product Developer Mary Hurley, who skillfully navigated the logistics involved with integrating a new project within the framework of an existing text. Much appreciation also goes to Production Manager Sherry Kane, who managed an aggressive but workable production schedule. In truth, this new text is the result of an entire team of publishing professionals, beginning with manuscript preparation and culminating with publication of the completed text that is brought to the chemistry community through the dedicated work of the marketing and sales team. Our sincere appreciation goes out to all of them.

JGS: I especially thank my husband Dan and the other members of my immediate family, who have experienced the day-do-day demands of living with a busy author. The joys and responsibilities of the family have always kept me grounded during the rewarding but sometimes all-consuming process of writing a textbook. This book, like prior editions of *Organic Chemistry*, is dedicated to my wonderful daughter Megan, who passed away after a nine-year battle with cystic fibrosis.

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Among the many others that go unnamed but who have profoundly affected this work are the thousands of students we have been lucky to teach over many years. We have learned so much from our daily interactions with them, and we hope that the wider chemistry community can benefit from this experience.

This edition has evolved based on the helpful feedback of many people who reviewed the fourth edition text and digital products, class-tested the book, and attended focus groups or symposiums. These many individuals have collectively provided constructive improvements to the project.

Listed below are the reviewers of the *Organic Chemistry*, fourth edition text:

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Although every effort has been made to make this text and its accompanying Student Study Guide/Solutions Manual as error-free as possible, some errors undoubtedly remain. Please feel free to email one of the authors about any inaccuracies, so that subsequent editions may be further improved.

With much aloha,

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## List of How To's

*How To* boxes provide detailed instructions for key procedures that students need to master. Below is a list of each *How To* and where it is presented in the text.

Chapter 1	Structure and Bonding How To Draw a Lewis Structure 14
	How To Interpret a Skeletal Structure 33
Chapter 2	Acids and Bases
Chapter 4	How To Determine the Relative Acidity of Protons 77  Alkanes
Chapter 4	How To Name an Alkane Using the IUPAC System 141
	How To Name a Cycloalkane Using the IUPAC System 145
	How To Draw a Newman Projection 151
	How To Draw the Chair Form of Cyclohexane 160
	How To Draw the Two Conformations for a Substituted Cyclohexane 162
	How To Draw Two Conformations for a Disubstituted Cyclohexane 165
Chapter 5	Stereochemistry
	How To Assign R or S to a Stereogenic Center 193
	How To Find and Draw All Possible Stereoisomers for a Compound with Two Stereogenic Centers 197
Chapter 7	Alkyl Halides and Nucleophilic Substitution
	How To Name an Alkyl Halide Using the IUPAC System 257
Chapter 9	Alcohols, Ethers, and Related Compounds
<b>.</b>	How To Name an Alcohol Using the IUPAC System 342
Chapter 10	Alkenes
	How To Name an Alkene 395
	How To Assign the Prefixes E and Z to an Alkene 397
Chapter 11	Alkynes
Ch 4 4 2	How To Develop a Retrosynthetic Analysis 453
Chapter 13	Mass Spectrometry and Infrared Spectroscopy
Chamban 14	How To Use MS and IR for Structure Determination 526
Chapter 14	Nuclear Magnetic Resonance Spectroscopy  How To Use <sup>1</sup> H NMR Data to Determine a Structure 562
Chapter 16	Conjugation, Resonance, and Dienes
Chapter 10	How To Draw the Product of a Diels–Alder Reaction 630
Chapter 17	Benzene and Aromatic Compounds
	How To Use the Inscribed Polygon Method to Determine the Relative Energies of MOs for Cyclic, Completely Conjugated Compounds 672
Chapter 18	Reactions of Aromatic Compounds
	How To Determine the Directing Effects of a Particular Substituent 707
Chapter 21	Aldehydes and Ketones—Nucleophilic Addition
	How To Determine the Starting Materials for a Wittig Reaction Using Retrosynthetic Analysis 848
Chapter 22	Carboxylic Acids and Their Derivatives—Nucleophilic Acyl Substitution
	How To Name an Ester (RCO <sub>2</sub> R') Using the IUPAC System 884
	How To Name a Thioester (RCOSR') Using the IUPAC System 884
	How To Name a 2° or 3° Amide 885
Chapter 24	Carbonyl Condensation Reactions
	How To Synthesize a Compound Using the Aldol Reaction 978
Ch + 2.5	How To Synthesize a Compound Using the Robinson Annulation 999
Chapter 25	Amines  Henry To Name 2° and 2° Amines with Different Allyd Crowns 1012
Chapter 26	How To Name 2° and 3° Amines with Different Alkyl Groups 1013  Amino Acids and Proteins
Chapter 26	
	<ul> <li>How To Use (R)-α-Methylbenzylamine to Resolve a Racemic Mixture of Amino Acids 1072</li> <li>How To Synthesize a Dipeptide from Two Amino Acids 1084</li> </ul>
	How To Synthesize a Peptide Using the Merrifield Solid Phase Technique 1089
Chapter 27	Carbohydrates
Chapter 27	How To Draw a Haworth Projection from an Acyclic Aldohexose 1122

## **List of Mechanisms**

Mechanisms are the key to understanding the reactions of organic chemistry. For this reason, great care has been given to present mechanisms in a detailed, step-by-step fashion. The list below indicates when each mechanism in the text is presented for the first time.

Chapter 7	Alkyl Halides and Nucleophilic Substitution 7.1 The S <sub>N</sub> 2 Mechanism 272 7.2 The S <sub>N</sub> 1 Mechanism 277
Chapter 8	Alkyl Halides and Elimination Reactions 8.1 The E2 Mechanism 312 8.2 The E1 Mechanism 318
Chapter 9	<ul> <li>Alcohols, Ethers, and Related Compounds</li> <li>9.1 Dehydration of 2° and 3° ROH—An E1 Mechanism 354</li> <li>9.2 Dehydration of a 1° ROH—An E2 Mechanism 355</li> <li>9.3 A 1,2-Methyl Shift—Carbocation Rearrangement During Dehydration 357</li> <li>9.4 Dehydration Using POCl₃ + Pyridine—An E2 Mechanism 359</li> <li>9.5 Reaction of a 1° ROH with HX—An S<sub>N</sub>2 Mechanism 361</li> <li>9.6 Reaction of 2° and 3° ROH with HX—An S<sub>N</sub>1 Mechanism 362</li> <li>9.7 Reaction of ROH with SOCl₂ + Pyridine—An S<sub>N</sub>2 Mechanism 364</li> <li>9.8 Reaction of ROH with PBr₃—An S<sub>N</sub>2 Mechanism 365</li> <li>9.9 Mechanism of Ether Cleavage in Strong Acid— (CH₃)₃COCH₃ + HI → (CH₃)₃CI + CH₃I + H₂O 371</li> </ul>
Chapter 10	Alkenes  10.1 Electrophilic Addition of HX to an Alkene 407  10.2 Electrophilic Addition of H <sub>2</sub> O to an Alkene—Hydration 412  10.3 Addition of X <sub>2</sub> to an Alkene—Halogenation 414  10.4 Addition of X and OH—Halohydrin Formation 416  10.5 Addition of H and BH <sub>2</sub> —Hydroboration 419
Chapter 11	Alkynes  11.1 Electrophilic Addition of HX to an Alkyne 443  11.2 Addition of X <sub>2</sub> to an Alkyne—Halogenation 444  11.3 Tautomerization in Acid 446  11.4 Hydration of an Alkyne 446
Chapter 12	Oxidation and Reduction  12.1 Addition of H <sub>2</sub> to an Alkene—Hydrogenation 467  12.2 Dissolving Metal Reduction of an Alkyne to a Trans Alkene 473  12.3 Reduction of RX with LiAlH <sub>4</sub> 475  12.4 Epoxidation of an Alkene with a Peroxyacid 477  12.5 Oxidation of an Alcohol with CrO <sub>3</sub> 486  12.6 Oxidation of a 1° Alcohol to a Carboxylic Acid 486
Chapter 15	Radical Reactions  15.1 Radical Halogenation of Alkanes 584  15.2 Allylic Bromination with NBS 594  15.3 Radical Addition of HBr to an Alkene 599  15.4 Radical Polymerization of CH <sub>2</sub> =CHZ 603
Chapter 16	Conjugation, Resonance, and Dienes  16.1 Biological Formation of Geranyl Diphosphate 616  16.2 Electrophilic Addition of HBr to a 1,3-Diene—1,2- and 1,4-Addition 625
Chapter 18	Reactions of Aromatic Compounds  18.1 General Mechanism—Electrophilic Aromatic Substitution 688  18.2 Bromination of Benzene 690  18.3 Formation of the Nitronium Ion (*NO <sub>2</sub> ) for Nitration 691  18.4 Formation of the Electrophile *SO <sub>2</sub> H for Sulfonation 692

	18.5 Formation of the Electrophile in Friedel–Crafts Alkylation—Two Possibilities 694 18.6 Friedel–Crafts Alkylation Using a 3° Carbocation 694 18.7 Formation of the Electrophile in Friedel–Crafts Acylation 695 18.8 Friedel–Crafts Alkylation Involving Carbocation Rearrangement 696 18.9 A Rearrangement Reaction Beginning with a 1° Alkyl Chloride 697 18.10 Nucleophilic Aromatic Substitution by Addition–Elimination 716 18.11 Nucleophilic Aromatic Substitution by Elimination–Addition: Benzyne 717 18.12 Benzylic Bromination 719
Chapter 20	Introduction to Carbonyl Chemistry; Organometallic Reagents;
	<ul> <li>Oxidation and Reduction</li> <li>20.1 Nucleophilic Addition—A Two-Step Process 777</li> <li>20.2 Nucleophilic Substitution—A Two-Step Process 778</li> <li>20.3 LiAlH<sub>4</sub> Reduction of RCHO and R<sub>2</sub>C=O 782</li> <li>20.4 Reduction of RCOCl and RCOOR' with a Metal Hydride Reagent 788</li> <li>20.5 Reduction of an Amide to an Amine with LiAlH<sub>4</sub> 790</li> <li>20.6 Nucleophilic Addition of R"MgX to RCHO and R<sub>2</sub>C=O 796</li> <li>20.7 Reaction of R"MgX or R"Li with RCOCl and RCOOR' 805</li> <li>20.8 Carboxylation—Reaction of RMgX with CO<sub>2</sub> 808</li> <li>20.9 1,2-Addition to an α,β-Unsaturated Carbonyl Compound 810</li> <li>20.10 1,4-Addition to an α,β-Unsaturated Carbonyl Compound 810</li> </ul>
Chapter 21	Aldehydes and Ketones—Nucleophilic Addition  21.1 General Mechanism—Nucleophilic Addition 839  21.2 General Mechanism—Acid-Catalyzed Nucleophilic Addition 839  21.3 Nucleophilic Addition of CN—Cyanohydrin Formation 843  21.4 The Wittig Reaction 847  21.5 Imine Formation from an Aldehyde or Ketone 851  21.6 Enamine Formation from an Aldehyde or Ketone 853  21.7 Base-Catalyzed Addition of H <sub>2</sub> O to a Carbonyl Group 856  21.8 Acid-Catalyzed Addition of H <sub>2</sub> O to a Carbonyl Group 856  21.9 Acetal Formation 859  21.10 Acid-Catalyzed Cyclic Hemiacetal Formation 863  21.11 A Cyclic Acetal from a Cyclic Hemiacetal 864
Chapter 22	Carboxylic Acids and Their Derivatives—Nucleophilic Acyl Substitution  22.1 General Mechanism—Nucleophilic Acyl Substitution 892  22.2 Conversion of Acid Chlorides to Anhydrides 896  22.3 Conversion of an Anhydride to an Amide 898  22.4 Conversion of Carboxylic Acids to Acid Chlorides 899  22.5 Fischer Esterification—Acid-Catalyzed Conversion of Carboxylic Acids to Esters 900  22.6 Conversion of Carboxylic Acids to Amides with DCC 902  22.7 Acid-Catalyzed Hydrolysis of an Ester to a Carboxylic Acid 904  22.8 Base-Promoted Hydrolysis of an Ester to a Carboxylic Acid 904  22.9 Amide Hydrolysis in Base 908  22.10 Biological Conversion of a Carboxylate to an Acyl Phosphate 911  22.11 Biological Conversion of an Acyl Phosphate to a Thioester 913  22.12 Hydrolysis of a Nitrile in Base 918  22.13 Reduction of a Nitrile with LiAlH <sub>4</sub> 919  22.14 Reduction of Grignard and Organolithium Reagents (R-M) to Nitriles 920
Chapter 23	Substitution Reactions of Carbonyl Compounds at the Carbon 23.1 Tautomerization in Acid 937 23.2 Tautomerization in Base 937 23.3 Acid-Catalyzed Halogenation at the α Carbon 948 23.4 Halogenation at the α Carbon in Base 949 23.5 The Haloform Reaction 950
Chapter 24	<ul> <li>Carbonyl Condensation Reactions</li> <li>24.1 The Aldol Reaction 974</li> <li>24.2 The Retro-Aldol Reaction 976</li> <li>24.3 Dehydration of β-Hydroxy Carbonyl Compounds with Base 977</li> <li>24.4 The Intramolecular Aldol Reaction 984</li> <li>24.5 The Claisen Reaction 986</li> </ul>

	24.6	The Dieckmann Reaction 990
	24.0	The Michael Reaction 995
	24.7	The Robinson Annulation 997
Chapter 25	Amine	
	25.1	The E2 Mechanism for the Hofmann Elimination 1039
	25.2	Formation of a Diazonium Salt from a 1° Amine 1042
	25.3	Formation of an <i>N</i> -Nitrosamine from a 2° Amine 1043
	25.4	Azo Coupling 1048
Chapter 26	Amino	Acids and Proteins
	26.1	Formation of an α-Amino Nitrile 1070
	26.2	Edman Degradation 1081
Chapter 27	Carbo	hydrates
	27.1	Glycoside Formation 1128
	27.2	Glycoside Hydrolysis 1129
Chapter 28	Lipids	
•	28.1	Biological Formation of Farnesyl Diphosphate 1172
	28.2	Isomerization of Geranyl Diphosphate to Neryl Diphosphate 1173
Chapter 29	Carbo	n-Carbon Bond-Forming Reactions in Organic Synthesis
	29.1	Suzuki Reaction 1191
	29.2	Heck Reaction 1194
	29.3	Formation of Dichlorocarbene 1195
	29.4	Addition of Dichlorocarbene to an Alkene 1196
	29.5	Simmons–Smith Reaction 1198
	29.6	Olefin Metathesis: $2 \text{ RCH} = \text{CH}_2 \rightarrow \text{RCH} = \text{CHR} + \text{CH}_2 = \text{CH}_2$ 1200
Chapter 31	Synthe	etic Polymers (Available online)
	31.1	Radical Polymerization of CH <sub>2</sub> =CHPh 1245
	31.2	Forming Branched Polyethylene During Radical Polymerization 1247
	31.3	Cationic Polymerization of CH <sub>2</sub> =CHZ 1248
	31.4	Anionic Polymerization of CH <sub>2</sub> =CHZ 1250
	31.5	Ziegler–Natta Polymerization of CH <sub>2</sub> =CH <sub>2</sub> 1253

### **List of Selected Applications**

Applications make any subject seem more relevant and interesting—for nonmajors and majors alike. The following is a list of the biological, medicinal, and environmental applications that have been integrated throughout *Organic Chemistry with Biological Topics*. Each chapter opener showcases an interesting and current application relating to the chapter's topic. (Code: G = general; M = medicinal; B = biological; E = environmental)

#### **Prologue**

- G Methane, the main component of natural gas
- G Ethanol, the alcohol in beverages
- E Trichlorofluoromethane, a CFC responsible for destroying the stratospheric ozone layer
- M Amoxicillin, a widely used antibiotic
- M Fluoxetine, the antidepressant Prozac
- M AZT, a drug used to treat HIV
- M Capsaicin, a compound found in topical pain relief creams
- E DDT, a nonspecific pesticide that persists in the environment
- M The antimalarial drugs quinine, chloroquine, and artemisinin

#### Chapter 1 Structure and Bonding

- M L-Dopa, a drug used to treat Parkinson's disease (Chapter opener and Section 1.14)
- M Alendronic acid (Fosamax), a drug used to prevent osteoporosis (Section 1.5)
- B Enanthotoxin, a poisonous compound isolated from hemlock water dropwort (Section 1.7)
- G Vanillin, the principal component in the extract of the vanilla bean (Section 1.8B)
- M Structures of active ingredients in common sunscreens (Section 1.8B)
- G Ethane, a component of natural gas (Section 1.10A)
- G Ethylene, a hydrocarbon used to make the plastic polyethylene (Section 1.10B)
- G Acetylene, a gas used in welding torches (Section 1.10C)
- G Cucumber aldehyde, the compound responsible for the odor of freshly cut cucumbers (Section 1.10C)
- M Sinemet, a drug used to treat Parkinson's disease that combines L-dopa and carbidopa (Section 1.14)
- B Vitamin B<sub>6</sub> (Section 1.14)

#### Chapter 2 Acids and Bases

- M Aspirin, a common analgesic and antipyretic (Chapter opener and Section 2.7)
- M The acid-base chemistry of morphine (Section 2.1)
- M The nasal decongestant pseudoephedrine (Section 2.5, Problem 2.17)
- M Glycolic acid, an α-hydroxy acid used in skin care products (Section 2.5, Problem 2.20)
- E Sulfuric acid, a major contributor to acid rain (Section 2.6)
- M Salicin, an analgesic found in willow bark (Section 2.7)

#### **Chapter 3** Introduction to Organic Molecules and Functional Groups

- 3 Vitamin C, a water-soluble vitamin that is important in the formation of collagen (Chapter opener and Section 3.5B)
- E Hemibrevetoxin B, a neurotoxin produced by algal blooms ("red tides") (Section 3.2B)
- M Diethyl ether, the first common general anesthetic (Section 3.2B)
- B Sucrose and the antibiotic amoxicillin (Section 3.2B, Problem 3.3)
- M Dexamethasone, a synthetic steroid (Section 3.2B, Problem 3.5)
- B Spermine, isolated from semen, and meperidine, the narcotic Demerol (Section 3.2B, Problem 3.6)
- M The anticancer agent doxorubicin (Adriamycin) (Section 3.2B, Problem 3.7)
- M Thyrotropin-releasing hormone (Section 3.2C, Problem 3.8)
- M Tamiflu, an antiviral drug used to treat influenza (Section 3.2C, Problem 3.9)
- B Pyruvic acid, lipoic acid, and folic acid as examples of biological molecules with multiple functional groups (Section 3.2C, Problem 3.10)
- B Biological phosphorus compounds (Section 3.2D)
- G How geckos use van der Waals forces to stick to walls (Section 3.3B)
- B Ionic, water-soluble biological compounds: isopentenyl diphosphate and acetylcholine (Section 3.4C)
- G MTBE, a high-octane additive in unleaded gasoline, and 4,4'-dichlorobiphenyl, a PCB (Section 3.4C)
- B Phenylalanine and 11-cis-retinal (Section 3.4C, Sample Problem 3.4)

- B Adrenaline and estrone (Section 3.4C, Problem 3.17)
- B Progesterone and testosterone (Section 3.4C, Sample Problem 3.5)
- B Norethindrone, an oral contraceptive, and arachidonic acid, a fatty acid (Section 3.4C, Problem 3.18)
- B Vitamin A (retinol), a fat-soluble vitamin found in the vision receptors of the eyes (Section 3.5A)
- B β-Carotene, a precursor to vitamin A (Section 3.5A)
- B Vitamin  $B_3$  and vitamin  $K_1$  (Section 3.5B, Problem 3.19)
- B Avocados as a source of pantothenic acid, vitamin B<sub>5</sub> (Section 3.5B, Problem 3.20)
- M Morphine and heroin (Section 3.7A, Problem 3.23)
- M The antibiotics nonactin and valinomycin (Section 3.7B)
- B The reactive features of isopentenyl diphosphate and pyruvic acid (Section 3.8)
- B The nucleophilic thiol of coenzyme A (Section 3.8)
- B Methionine, ATP, and S-adenosylmethionine (Section 3.8, Problem 3.28)
- B Amino acids and proteins (Section 3.9A)
- B Monosaccharides and carbohydrates (Section 3.9B)
- B Nucleotides and nucleic acids (Section 3.9C)
- B Lipids (Section 3.9D)
- M, B End-of-chapter problems: 3.33–3.35, 3.37, 3.38, 3.40, 3.48–3.57, 3.60, 3.61, 3.63–3.65, and 3.67

#### Chapter 4 Alkanes

- E Oil slicks that result from crude petroleum being spilled into the ocean from oil tankers or oil wells (Chapter opener)
- B The cockroach pheromone undecane (Section 4.1)
- B Cyclohexane, one component of mangoes (Section 4.1)
- B Allicin, a compound responsible for the odor of garlic (Section 4.3)
- M Systematic names, generic names, and trade names in over-the-counter drugs like Motrin (Section 4.3)
- G Fossil fuels such as natural gas and petroleum (Section 4.7)
- E The combustion of alkanes and how it contributes to climate change (Section 4.14B)
- B Lipids such as fat-soluble vitamins, phospholipids, waxes, prostaglandins, and steroids (Section 4.15)
- B Pristane, a high molecular weight alkane found in shark liver oil (Section 4.15, Problem 4.33)
- B End-of-chapter problems: 4.66 and 4.69

#### Chapter 5 Stereochemistry

- M, B Paclitaxel (Taxol), a drug used to treat ovarian, breast, and other cancers (Chapter opener)
  - B How differences in the three-dimensional structure of starch and cellulose affect their shape and function (Section 5.1)
- M, B Identifying stereogenic centers in Darvon (an analgesic), ephedrine (a decongestant), and fructose (a simple sugar) (Section 5.4A)
  - M The three-dimensional structure of thalidomide, an anti-nausea drug that caused catastrophic birth defects (Section 5.5)
- M, B Identifying stereogenic centers in paclitaxel (anticancer agent) and sucrose (Section 5.5)
  - M Identifying stereogenic centers in gabapentin (a drug used to treat seizures and chronic pain), gabapentin enacarbil, cholesterol, and Zocor (cholesterol-lowering drug) (Section 5.5, Problems 5.9 and 5.10)
  - M Assigning R and S configurations in the drugs Plavix and Zestril (Section 5.6, Problems 5.14 and 5.15)
  - B The sweetener sorbitol (Section 5.9, Problem 5.24)
  - B The specific rotation of MSG, a common flavor enhancer (Section 5.12D, Problem 5.32)
  - M Chiral drugs and how mirror image isomers can have drastically different properties—the analgesic ibuprofen, the antidepressant fluoxetine, and the anti-inflammatory agent naproxen (Section 5.13A)
  - B The sense of smell and how mirror image isomers (e.g., carvone and celery ketone) can smell different (Section 5.13B and Problem 5.35)
- M, B End-of-chapter problems: 5.36, 5.43, 5.49, 5.50, 5.53, 5.55, 5.60, and 5.65–5.71

#### **Chapter 6** Understanding Organic Reactions

- B Entropy changes in the metabolism of glucose (Chapter opener and Section 6.4)
- B A biological substitution reaction: the hydrolysis of a triacylglycerol to glycerol + fatty acids (Section 6.2A)
- B A biological elimination reaction in the citric acid cycle (Section 6.2B)
- B A biological addition reaction with a thioester, a key step in fatty acid metabolism (Section 6.2C)
- B Four enzyme-catalyzed steps in the citric acid cycle (Section 6.2C, Problem 6.2)
- B The air oxidation of vegetable oils (Section 6.3C, Sample Problem 6.1)
- B Examples of exothermic reactions: the hydrolysis of ATP and the oxidation of glucose (Section 6.4)
- B Coupled reactions in metabolism (Section 6.5C)
- G The reaction of gasoline with  $O_2$  (Section 6.9A)
- G Refrigeration and spoilage (Section 6.9A)
- B Enzymes, biological catalysts (Section 6.11)
- B End-of-chapter problems: 6.27, 6.28, 6.32, 6.39, 6.41, 6.52, and 6.56