Organic Chemistry

EIGHTH EDITION

Paula Yurkanis Bruice

University Of California Santa Barbara



Brief Table of Contents

	Preface xxii
CHAPTER 1	Remembering General Chemistry: Electronic Structure and Bonding 2
CHAPTER 2	Acids and Bases: Central to Understanding Organic Chemistry 50
TUTORIAL	Acids and Bases 80
CHAPTER 3	An Introduction to Organic Compounds: Nomenclature, Physical Properties, and Structure 88
TUTORIAL	Using Molecular Models 142
CHAPTER 4	Isomers: The Arrangement of Atoms in Space 143
TUTORIAL	Interconverting Structural Representations 187
CHAPTER 5	Alkenes: Structure, Nomenclature, and an Introduction to Reactivity • Thermodynamics and Kinetics 190
TUTORIAL	Drawing Curved Arrows 225
CHAPTER 6	The Reactions of Alkenes • The Stereochemistry of Addition Reactions 235
CHAPTER 7	The Reactions of Alkynes • An Introduction to Multistep Synthesis 288
CHAPTER 8	Delocalized Electrons: Their Effect on Stability, pK_a , and the Products of a Reaction • Aromaticity and Electronic Effects: An Introduction to the Reactions of Benzene 318
TUTORIAL	Drawing Resonance Contributors 382
CHAPTER 9	Substitution and Elimination Reactions of Alkyl Halides 391
CHAPTER 10	Reactions of Alcohols, Ethers, Epoxides, Amines, and Sulfur-Containing Compounds 458
CHAPTER 11	Organometallic Compounds 508
CHAPTER 12	Radicals 532
TUTORIAL	Drawing Curved Arrows in Radical Systems 563
CHAPTER 13	Mass Spectrometry; Infrared Spectroscopy; UV/Vis Spectroscopy 567
CHAPTER 14	NMR Spectroscopy 620
CHAPTER 15	Reactions of Carboxylic Acids and Carboxylic Acid Derivatives 686

- **CHAPTER 16** Reactions of Aldehydes and Ketones More Reactions of Carboxylic Acid Derivatives **739**
- **CHAPTER 17** Reactions at the α -Carbon **801**
- **TUTORIAL** Synthesis and Retrosynthetic Analysis **854**
- CHAPTER 18 Reactions of Benzene and Substituted Benzenes 868
- CHAPTER 19 More About Amines Reactions of Heterocyclic Compounds 924
- CHAPTER 20 The Organic Chemistry of Carbohydrates 950
- CHAPTER 21 Amino Acids, Peptides, and Proteins 986
- CHAPTER 22 Catalysis in Organic Reactions and in Enzymatic Reactions 1030
- **CHAPTER 23** The Organic Chemistry of the Coenzymes, Compounds Derived from Vitamins **1063**
- CHAPTER 24 The Organic Chemistry of the Metabolic Pathways 1099
- CHAPTER 25 The Organic Chemistry of Lipids 1127
- CHAPTER 26 The Chemistry of the Nucleic Acids 1155
- CHAPTER 27 Synthetic Polymers 1182
- CHAPTER 28 Pericyclic Reactions 1212
- **APPENDICES I** pK_a Values A-1
 - II Kinetics A-3
 - III Summary of Methods Used to Synthesize a Particular Functional Group A-8
 - **v** Summary of Methods Employed to Form Carbon–Carbon Bonds A-11
 - v Spectroscopy Tables A-12
 - **vi** Physical Properties of Organic Compounds A-18
 - **vii** Answers to Selected Problems ANS-1

Glossary G-1

Photo Credits C-1

Index I-1

Complete List of In-Chapter Connection Features

Medical Connections

Fosamax Prevents Bones from Being Nibbled Away (2.8) Aspirin Must Be in its Basic Form to be Physiologically Active (2.10) Blood: A Buffered Solution (2.11) Drugs Bind to Their Receptors (3.9) Cholesterol and Heart Disease (3.16) How High Cholesterol is Treated Clinically (3.16) The Enantiomers of Thalidomide (4.17) Synthetic Alkynes Are Used to Treat Parkinson's Disease (7.0) Synthetic Alkynes Are Used for Birth Control (7.1) The Inability to Perform an S_N2 Reaction Causes a Severe Clinical Disorder (10.3) Treating Alcoholism with Antabuse (10.5) Methanol Poisoning (10.5) Anesthetics (10.6) Alkylating Agents as Cancer Drugs (10.11) S-Adenosylmethionine: A Natural Antidepressant (10.12) Artificial Blood (12.12) Nature's Sleeping Pill (15.1) Penicillin and Drug Resistance (15.12) **Dissolving Sutures (15.13)** Cancer Chemotherapy (16.17) Breast Cancer and Aromatase Inhibitors (17.12) Thyroxine (18.3) A New Cancer-Fighting Drug (18.20) Atropine (19.2) Porphyrin, Bilirubin, and Jaundice (19.7) Measuring the Blood Glucose Levels in Diabetes (20.8) Galactosemia (20.15) Why the Dentist is Right (20.16) Resistance to Antibiotics (20.17) Heparin-A Natural Anticoagulant (20.17) Amino Acids and Disease (21.2) Diabetes (21.8) Diseases Caused by a Misfolded Protein (21.15) How Tamiflu Works (22.11) Assessing the Damage After a Heart Attack (23.5) Cancer Drugs and Side Effects (23.7) Anticoagulants (23.8) Phenylketonuria (PKU): An Inborn Error of Metabolism (24.8) Alcaptonuria (24.8) Multiple Sclerosis and the Myelin Sheath (25.5) How Statins Lower Cholesterol Levels (25.8) One Drug—Two Effects (25.10) Sickle Cell Anemia (26.9) Antibiotics That Act by Inhibiting Translation (26.9) Antibiotics Act by a Common Mechanism (26.10) Health Concerns: Bisphenol A and Phthalates (27.11)

Biological Connections

Poisonous Amines (2.3) Cell Membranes (3.10) How a Banana Slug Knows What to Eat (7.2) Electron Delocalization Affects the Three-Dimensional Shape of Proteins (8.4) Naturally Occurring Alkyl Halides That Defend Against Predators (9.5) Biological Dehydrations (10.4) Alkaloids (10.9) Dalmatians: Do Not Fool with Mother Nature (15.11) A Semisynthetic Penicillin (15.12) Preserving Biological Specimens (16.9) A Biological Friedel-Crafts Alkylation (18.7) A Toxic Disaccharide (20.15) Controlling Fleas (20.16) Primary Structure and Taxonomic Relationship (21.12) Competitive Inhibitors (23.7) Whales and Echolocation (25.3) Snake Venom (25.5) Cyclic AMP (26.1) There Are More Than Four Bases in DNA (26.7)

Chemical Connections

Natural versus Synthetic Organic Compounds (1.0) Diamond, Graphite, Graphene, and Fullerenes: Substances that Contain Only Carbon Atoms (1.8) Water—A Unique Compound (1.12) Acid Rain (2.2) Derivation of the Henderson-Hasselbalch Equation (2.10) Bad-Smelling Compounds (3.7) Von Baeyer, Barbituric Acid, and Blue Jeans (3.12) Starch and Cellulose—Axial and Equatorial (3.14) Cis-Trans Interconversion in Vision (4.1) The Difference between ΔG^{\ddagger} and E_{a} (5.11) Calculating Kinetic Parameters (End of Ch 05) Borane and Diborane (6.8) Cyclic Alkenes (6.13) Chiral Catalysts (6.15) Sodium Amide and Sodium in Ammonia (7.10) Buckyballs (8.18) Why Are Living Organisms Composed of Carbon Instead of Silicon? (9.2) Solvation Effects (9.14) The Lucas Test (10.1) Crown Ethers—Another Example of Molecular Recognition (10.7) Crown Ethers Can be Used to Catalyze $S_N 2$ Reactions (10.7) Eradicating Termites (10.12) Cyclopropane (12.9) What Makes Blueberries Blue and Strawberries Red? (13.22) Nerve Impulses, Paralysis, and Insecticides (15.19) Enzyme-Catalyzed Carbonyl Additions (16.4) Carbohydrates (16.9) β -Carotene (16.13) Synthesizing Organic Compounds (16.14) Enzyme-Catalyzed Cis-Trans Interconversion (16.16) Incipient Primary Carbocations (18.7) Hair: Straight or Curly? (21.8) Right-Handed and Left-Handed Helices (21.14) β -Peptides: An Attempt to Improve on Nature (21.14) Why Did Nature Choose Phosphates? (24.1) Protein Prenylation (25.8) Bioluminescence (28.6)

Pharmaceutical Connections

Chiral Drugs (4.18) Why Are Drugs so Expensive? (7.0) Lead Compounds for the Development of Drugs (10.9) Aspirin, NSAIDs, and COX-2 Inhibitors (15.9) Penicillins in Clinical Use (15.12) Serendipity in Drug Development (16.8) Semisynthetic Drugs (16.14) Drug Safety (18.19) Searching for Drugs: An Antihistamine, a Nonsedating Antihistamine, and a Drug for Ulcers (19.7) A Peptide Antibiotic (21.2) Natural Products That Modify DNA (26.6) Using Genetic Engineering to Treat the Ebola Virus (26.13) Nanocontainers (27.9)

Historical Connections

Kekule's Dream (8.1)
Mustard Gas–A Chemical Warfare Agent (10.11)
Grubbs, Schrock, Suzuki, and Heck Receive the Nobel Prize (11.5)
The Nobel Prize (11.5)
Why Radicals No Longer Have to Be Called Free Radicals (12.2)
Nikola Tesla (1856–1943) (14.1)
The Discovery of Penicillin (15.12)
Discovery of the First Antibiotic (18.19)
Vitamin C (20.17)
Vitamin B₁ (23.0)
Niacin Deficiency (23.1)
The First Antibiotics (23.7)
The Structure of DNA: Watson, Crick, Franklin, and Wilkins (26.1)
Influenza Pandemics (26.11)

Nutritional Connections

Trans Fats (5.9) Decaffeinated Coffee and the Cancer Scare (12.11) Food Preservatives (12.11) Is Chocolate a Health Food? (12.11) Nitrosamines and Cancer (18.20) Lactose Intolerance (20.15) Acceptable Daily Intake (20.19) Proteins and Nutrition (21.1) Too Much Broccoli (23.8) Differences in Metabolism (24.0) Fats Versus Carbohydrates as a Source of Energy (24.6) Basal Metabolic Rate (24.10) Omega Fatty Acids (25.1) Olestra: Nonfat with Flavor (25.3) Melamine Poisoning (27.12) The Sunshine Vitamin (28.6) Animals, Birds, Fish—And Vitamin D (28.6)

Industrial Connections

How is the Octane Number of Gasoline Determined? (3.2) Organic Compounds That Conduct Electricity (8.7) Synthetic Polymers (15.13) The Synthesis of Aspirin (17.7) Teflon: An Accidental Discovery (27.3) Designing a Polymer (27.11)

Environmental Connections

Pheromones (5.0)
Which are More Harmful: Natural Pesticides or Synthetic Pesticides? (6.16)
Green Chemistry: Aiming for Sustainability (7.12)
The Birth of the Environmental Movement (9.0)
Environmental Adaptation (9.14)
Benzo[a]pyrene and Cancer (10.8)
Chimney Sweeps and Cancer (10.8)
Resisting Herbicides (26.13)
Recycling Symbols (27.3)

General Connections

A Few Words About Curved Arrows (5.5) Grain Alcohol and Wood Alcohol (10.1) Blood Alcohol Concentration (10.5) Natural Gas and Petroleum (12.1) Fossil Fuels: A Problematic Energy Source (12.1) Mass Spectrometry in Forensics (13.8) The Originator of Hooke's Law (13.13) Ultraviolet Light and Sunscreens (13.19) Structural Databases (14.24) What Drug-Enforcement Dogs Are Really Detecting (15.16) Butanedione: An Unpleasant Compound (16.1) Measuring Toxicity (18.0) The Toxicity of Benzene (18.1) Glucose/Dextrose (20.9) Water Softeners: Examples of Cation-Exchange Chromatography (21.5) Curing a Hangover with Vitamin B_1 (23.3)

Contents

PART An Introduction to the Study of Organic Chemistry 1 ONE

Remembering General Chemistry: Electronic Structure and Bonding 2

CHEMICAL CONNECTION: Natural versus Synthetic Organic Compounds 3

- 1.1 The Structure of an Atom 4
- 1.2 How the Electrons in an Atom are Distributed 5
- 1.3 Covalent Bonds 7
- 1.4 How the Structure of a Compound is Represented 13 PROBLEM-SOLVING STRATEGY 15
- 1.5 Atomic Orbitals 19
- 1.6 An Introduction to Molecular Orbital Theory 21
- 1.7 How Single Bonds are Formed in Organic Compounds 25
- How a Double Bond is Formed: The Bonds in Ethene 29
 CHEMICAL CONNECTION: Diamond, Graphite, Graphene, and Fullerenes: Substances that Contain Only Carbon Atoms 31
- **1.9** How a Triple Bond is Formed: The Bonds in Ethyne **31**
- 1.10 The Bonds in the Methyl Cation, the Methyl Radical, and the Methyl Anion 33
- 1.11 The Bonds in Ammonia and in the Ammonium Ion 35
- 1.12 The Bonds in Water 36
 - CHEMICAL CONNECTION: Water—A Unique Compound 37
- **1.13** The Bond in a Hydrogen Halide **38**
- 1.14 Hybridization and Molecular Geometry 39 PROBLEM-SOLVING STRATEGY 39
- 1.15 Summary: Hybridization, Bond Lengths, Bond Strengths, and Bond Angles 40 PROBLEM-SOLVING STRATEGY 44
- 1.16 Dipole Moments of Molecules 44
- ESSENTIAL CONCEPTS 46 PROBLEMS 47

Acids and Bases: Central to Understanding Organic Chemistry 50

- 2.1 An Introduction to Acids and Bases 50
- 2.2 pK_a and pH 52 PROBLEM-SOLVING STRATEGY 54 CHEMICAL CONNECTION: Acid Rain 54
- 2.3 Organic Acids and Bases 55 BIOLOGICAL CONNECTION: Poisonous Amines 56 PROBLEM-SOLVING STRATEGY 58
- 2.4 How to Predict the Outcome of an Acid-Base Reaction 58
- 2.5 How to Determine the Position of Equilibrium 59
- **2.6** How the Structure of an Acid Affects its pK_a Value **60**
- 2.7 How Substituents Affect the Strength of an Acid 64 PROBLEM-SOLVING STRATEGY 64
- An Introduction to Delocalized Electrons 66
 MEDICAL CONNECTION: Fosamax Prevents Bones from Being Nibbled Away 67
 PROBLEM-SOLVING STRATEGY 68
- 2.9 A Summary of the Factors that Determine Acid Strength 69
- How pH Affects the Structure of an Organic Compound 70
 PROBLEM-SOLVING STRATEGY 71
 CHEMICAL CONNECTION: Derivation of the Henderson-Hasselbalch Equation 72
 MEDICAL CONNECTION: Aspirin Must Be in its Basic Form to be Physiologically Active 74
- 2.11 Buffer Solutions 74 MEDICAL CONNECTION: Blood: A Buffered Solution 75
 2.12 Lewis Acids and Bases 76
 - ESSENTIAL CONCEPTS 77 PROBLEMS 77

MasteringChemistry[®] for Organic Chemistry

MasteringChemistry tutorials guide you through the toughest topics in chemistry with self-paced tutorials that provide individualized coaching. These assignable, in-depth tutorials are designed to coach you with hints and feedback specific to your individual misconceptions. For additional practice on Acids and Bases, go to MasteringChemistry, where the following tutorials are available:

- Acids and Bases: Definitions
- Acids and Bases: Factors That Influence Acid Strength
- Acids and Bases: Base Strength and the Effect of pH on Structure
- Acids and Bases: Predicting the Position of Equilibrium

TUTORIAL Acids and Bases 80

3	An Introduction to Organic Compounds: Nomenclature, Physical Properties, and Structure 88
3.1	
3.1	Alkyl Groups 92 The Nomenclature of Alkanes 95
3.2	INDUSTRIAL CONNECTION: How is the Octane Number of Gasoline Determined? 98
3.3	The Nomenclature of Cycloalkanes 99
3.3	PROBLEM-SOLVING STRATEGY 101
2.4	
3.4	The Nomenclature of Alkyl Halides 101
3.5	The Nomenclature of Ethers 103
3.6	The Nomenclature of Alcohols 104
3.7	The Nomenclature of Amines 106
3.8	CHEMICAL CONNECTION: Bad-Smelling Compounds 109 The Structures of Alkyl Halides, Alcohols, Ethers, and Amines 109
3.0 3.9	Noncovalent Interactions 110
3.5	PROBLEM-SOLVING STRATEGY 114
3.10	MEDICAL CONNECTION: Drugs Bind to Their Receptors 114 The Solubility of Organic Compounds 116
3.10	BIOLOGICAL CONNECTION: Cell Membranes 118
3.11	Rotation Occurs about Carbon–Carbon Single Bonds 118
3.12	Some Cycloalkanes Have Angle Strain 122
J.12	CHEMICAL CONNECTION: Von Baeyer, Barbituric Acid, and Blue Jeans 123
	PROBLEM-SOLVING STRATEGY 123
3.13	
3.13	Conformers of Cyclohexane 124 Conformers of Monosubstituted Cyclohexanes 127
J.14	CHEMICAL CONNECTION: Starch and Cellulose—Axial and Equatorial 128
3.15	Conformers of Disubstituted Cyclohexanes 129
0.10	PROBLEM-SOLVING STRATEGY 130
2 10	PROBLEM-SOLVING STRATEGY 132
3.16	Fused Cyclohexane Rings 134
	MEDICAL CONNECTION: Cholesterol and Heart Disease 134
	MEDICAL CONNECTION: How High Cholesterol is Treated Clinically 135

PART Electrophilic Addition Reactions, Stereochemistry, and Electron Delocalization 141

TUTORIAL Using Molecular Models 142

4. Isomers: The Arrangement of Atoms in Space 143
4.1 Cis–Trans Isomers Result from Restricted Rotation 145

ESSENTIAL CONCEPTS 135 PROBLEMS 136

4.2 CHEMICAL CONNECTION: Cis-Trans Interconversion in Vision 1474.2 Using the *E,Z* System to Distinguish Isomers 147

PROBLEM-SOLVING STRATEGY 150

- **4.3** A Chiral Object Has a Nonsuperimposable Mirror Image **150**
- 4.4 An Asymmetric Center is a Cause of Chirality in a Molecule 151
- 4.5 Isomers with One Asymmetric Center 152
- **4.6** Asymmetric Centers and Stereocenters **153**
- 4.7 How to Draw Enantiomers 153
- 4.8 Naming Enantiomers by the *R*,*S* System 154 PROBLEM-SOLVING STRATEGY 157

PROBLEM-SOLVING STRATEGY 158

- 4.9 Chiral Compounds Are Optically Active 159
- 4.10 How Specific Rotation Is Measured 161
- 4.11 Enantiomeric Excess 163
- 4.12 Compounds with More than One Asymmetric Center 164
- 4.13 Stereoisomers of Cyclic Compounds 166 PROBLEM-SOLVING STRATEGY 168
- 4.14 Meso Compounds Have Asymmetric Centers but Are Optically Inactive 169 PROBLEM-SOLVING STRATEGY 171

MasteringChemistry*

for Organic Chemistry

Mastering Chemistry tutorials guide you through the toughest topics in chemistry with self-paced tutorials that provide individualized coaching. These assignable, in-depth tutorials are designed to coach you with hints and feedback specific to your individual misconceptions. For additional practice on Molecular Models, go to MasteringChemistry where the following tutorials are available:

- Basics of Model Building
- Building and Recognizing Chiral Molecules
- Recognizing Chirality in Cyclic Molecules

Using the *E*,*Z* system to name alkenes was moved to Chapter 4, so now it appears immediately after using cis and trans to distinguish alkene stereoisomers.

MasteringChemistry*

for Organic Chemistry

MasteringChemistry tutorials guide you through the toughest topics in chemistry with self-paced tutorials that provide individualized coaching. These assignable, in-depth tutorials are designed to coach you with hints and feedback specific to your individual misconceptions. For additional practice on Interconverting Structural Representations, go to MasteringChemistry where the following tutorials are available:

- Interconverting Fischer Projections and Perspective Formulas
- Interconverting Perspective Formulas, Fischer Projections, and Skeletal Structures
- Interconverting Perspective Formulas, Fischer Projections, and Newman Projections

Catalytic hydrogenation and relative stabilities of alkenes were moved from Chapter 6 to Chapter 5 (thermodynamics), so they can be used to illustrate how ΔH° values can be used to determine relative stabilities.

MasteringChemistry[®]

for Organic Chemistry

MasteringChemistry tutorials guide you through the toughest topics in chemistry with self-paced tutorials that provide individualized coaching. These assignable, in-depth tutorials are designed to coach you with hints and feedback specific to your individual misconceptions. For additional practice on Drawing Curved Arrows: Pushing Electrons, go to MasteringChemistry where the following tutorials are available:

- An Exercise in Drawing Curved Arrows: Pushing Electrons
- An Exercise in Drawing Curved Arrows: Predicting Electron Movement
- An Exercise in Drawing Curved Arrows: Interpreting Electron Movement

All the reactions in Chapter 6 follow the same mechanism the first step is always addition of the electrophile to the sp^2 carbon bonded to the most hydrogens.

- 4.15 How to Name Isomers with More than One Asymmetric Center 172 PROBLEM-SOLVING STRATEGY 175
- 4.16 Nitrogen and Phosphorus Atoms Can Be Asymmetric Centers 177
- 4.17 Receptors 178
- MEDICAL CONNECTION: The Enantiomers of Thalidomide 1794.18 How Enantiomers Can Be Separated 179
 - PHARMACEUTICAL CONNECTION: Chiral Drugs 180

ESSENTIAL CONCEPTS 181 PROBLEMS 181

TUTORIAL Interconverting Structural Representations 187

5 Alkenes: Structure, Nomenclature, and an Introduction to Reactivity • Thermodynamics and Kinetics 190

ENVIRONMENTAL CONNECTION: Pheromones 191

- 5.1 Molecular Formulas and the Degree of Unsaturation 191
- 5.2 The Nomenclature of Alkenes 192
- 5.3 The Structure of Alkenes 195

PROBLEM-SOLVING STRATEGY 196

- 5.4 How An Organic Compound Reacts Depends on Its Functional Group 197
- 5.5 How Alkenes React Curved Arrows Show the Flow of Electrons 198
- **GENERAL CONNECTION:** A Few Words About Curved Arrows **200 5.6** Thermodynamics: How Much Product is Formed? **202**
- 5.7 Increasing the Amount of Product Formed in a Reaction 205
- **5.8** Calculating ΔH° Values **206**
- **5.9** Using ΔH° Values to Determine the Relative Stabilities of Alkenes **207 PROBLEM-SOLVING STRATEGY 208**

NUTRITIONAL CONNECTION: Trans Fats 211

- 5.10 Kinetics: How Fast is the Product Formed? 211
- **5.11** The Rate of a Chemical Reaction **213 CHEMICAL CONNECTION:** The Difference between ΔG^{\ddagger} and E_a **215**
- 5.12 A Reaction Coordinate Diagram Describes the Energy Changes That Take Place During a Reaction 215
- **5.13** Catalysis **218**
- 5.14 Catalysis by Enzymes 219
 - ESSENTIAL CONCEPTS 220 PROBLEMS 221 CHEMICAL CONNECTION: Calculating Kinetic Parameters 224

TUTORIAL Drawing Curved Arrows 225

- 6 The Reactions of Alkenes The Stereochemistry of Addition Reactions
- 6.1 The Addition of a Hydrogen Halide to an Alkene 236
- 6.2 Carbocation Stability Depends on the Number of Alkyl Groups Attached to the Positively Charged Carbon 237

235

- 6.3 What Does the Structure of the Transition State Look Like? 239
- 6.4 Electrophilic Addition Reactions Are Regioselective 241 PROBLEM-SOLVING STRATEGY 243
- 6.5 The Addition of Water to an Alkene 245
- **6.6** The Addition of an Alcohol to an Alkene **246**
- 6.7 A Carbocation Will Rearrange if It Can Form a More Stable Carbocation 248
- 6.8 The Addition of Borane to an Alkene: Hydroboration–Oxidation 250
- **CHEMICAL CONNECTION:** Borane and Diborane **251 6.9** The Addition of a Halogen to an Alkene **254**
- PROBLEM-SOLVING STRATEGY 257
- 6.10 The Addition of a Peroxyacid to an Alkene 257
- 6.11 The Addition of Ozone to an Alkene: Ozonolysis 259 PROBLEM-SOLVING STRATEGY 261
- 6.12 Regioselective, Stereoselective, And Stereospecific Reactions 263
- 6.13 The Stereochemistry of Electrophilic Addition Reactions 264 CHEMICAL CONNECTION: Cyclic Alkenes 269 PROBLEM-SOLVING STRATEGY 274
- 6.14 The Stereochemistry of Enzyme-Catalyzed Reactions 276

6.15	Enantiomers Can Be Distinguished by Biological Molecules 277 CHEMICAL CONNECTION: Chiral Catalysts 278
6.16	Reactions and Synthesis 278 ENVIRONMENTAL CONNECTION: Which are More Harmful: Natural Pesticides or Synthetic Pesticides? 280
	ESSENTIAL CONCEPTS 280 SUMMARY OF REACTIONS 281 PROBLEMS 282
7	The Reactions of Alkynes • An Introduction to Multistep Synthesis 288
	MEDICAL CONNECTION: Synthetic Alkynes Are Used to Treat Parkinson's Disease 289
7.1	PHARMACEUTICAL CONNECTION: Why Are Drugs so Expensive?290The Nomenclature of Alkynes290MEDICAL CONNECTION: Synthetic Alkynes Are Used for Birth Control291
7.2 7.3	How to Name a Compound That Has More than One Functional Group 292 The Structure of Alkynes 293
7.4	BIOLOGICAL CONNECTION: How a Banana Slug Knows What to Eat 293 The Physical Properties of Unsaturated Hydrocarbons 294
7.5 7.6	The Reactivity of Alkynes 295 The Addition of Hydrogen Halides and the Addition of Halogens to an Alkyne 296
7.7 7.8	The Addition of Water to an Alkyne 299 The Addition of Borane to an Alkyne: Hydroboration–Oxidation 301
7.9 7.10	The Addition of Hydrogen to an Alkyne 302 A Hydrogen Bonded to an <i>sp</i> Carbon Is "Acidic" 304
7.10	CHEMICAL CONNECTION: Sodium Amide and Sodium in Ammonia 305 PROBLEM-SOLVING STRATEGY 305
7.11 7.12	Synthesis Using Acetylide Ions 306 DESIGNING A SYNTHESIS I: An Introduction to Multistep Synthesis 307
7.12	ENVIRONMENTAL CONNECTION: Green Chemistry: Aiming for Sustainability 312 ESSENTIAL CONCEPTS 312 SUMMARY OF REACTIONS 313 PROBLEMS 314
•	
8	Delocalized Electrons: Their Effect on Stability, pK_a , and the Products of a Reaction • Aromaticity and Electronic Effects: An Introduction to the
	Reactions of Benzene 318
8.1	Delocalized Electrons Explain Benzene's Structure 319 HISTORICAL CONNECTION: Kekule's Dream 321
8.2 8.3	The Bonding in Benzene 321 Resonance Contributors and the Resonance Hybrid 322
8.4	How to Draw Resonance Contributors 323
	BIOLOGICAL CONNECTION: Electron Delocalization Affects the Three-Dimensional Shape of Proteins 326
8.5	The Predicted Stabilities of Resonance Contributors 326 PROBLEM-SOLVING STRATEGY 328
8.6	Delocalization Energy is the Additional Stability Delocalized Electrons Give to a Compound 329
8.7	Delocalized Electrons Increase Stability 330
8.8	INDUSTRIAL CONNECTION: Organic Compounds That Conduct Electricity 333 A Molecular Orbital Description of Stability 335
8.9	Delocalized Electrons Affect pK_a Values 339 PROBLEM-SOLVING STRATEGY 342
8.10 8.11	Electronic Effects 342 Delocalized Electrons Can Affect the Product of a Reaction 346
8.12	Reactions of Dienes 347
8.13 8.14	Thermodynamic Versus Kinetic Control 350 The Diels–Alder Reaction is a 1,4-Addition Reaction 355
8.15 8.16	Retrosynthetic Analysis of the Diels–Alder Reaction 361 Benzene is an Aromatic Compound 362
8.17 8.18	The Two Criteria for Aromaticity 363 Applying the Criteria for Aromaticity 364
0.10	CHEMICAL CONNECTION: Buckyballs 365
8.19	PROBLEM-SOLVING STRATEGY 366 A Molecular Orbital Description of Aromaticity 367
-	

Chapter 8 starts by discussing the structure of benzene because it is the ideal compound to use to explain delocalized electrons. This chapter also includes a discussion of aromaticity, so a short introduction to electrophilic aromatic substitution reactions is now included. This allows students to see how aromaticity causes benzene to undergo electrophilic substitution rather than electrophilic addition the reactions they have just finished studying.

Traditionally, electronic effects are taught so students can understand the directing effects of substituents on benzene rings. Now that most of the chemistry of benzene follows carbonyl chemistry, students need to know about electronic effects before they get to benzene chemistry (so they are better prepared for spectroscopy and carbonyl chemistry). Therefore, electronic effects are now discussed in Chapter 8 and used to teach students how substituents affect the pK_a values of phenols, benzoic acids, and anilinium ions. Electronic effects are then reviewed in the chapter on benzene.

MasteringChemistry[®]

for Organic Chemistry

MasteringChemistry tutorials guide you through the toughest topics in chemistry with self-paced tutorials that provide individualized coaching. These assignable, in-depth tutorials are designed to coach you with hints and feedback specific to your individual misconceptions. For additional practice on Drawing Resonance Contributors, go to MasteringChemistry where the following tutorials are available:

- Drawing Resonance Contributors: Moving π Electrons
- Drawing Resonance Contributors: Predicting
 Aromaticity
- Drawing Resonance Contributors: Substituted Benzene Rings

U.ZU AIUIIIalic Heleiucychic Cumpulius 3	8.20	pounds 36	Aromatic Heterocyclic Compou	ınds 30
--	------	-----------	------------------------------	---------

8.21 How Benzene Reacts 370

8.22 Organizing What We Know About the Reactions of Organic Compounds (Group I) 372

ESSENTIAL CONCEPTS 373 SUMMARY OF REACTIONS 374 PROBLEMS 375

TUTORIAL Drawing Resonance Contributors 382

PART Substitution and Elimination Reactions 390 THREE

9 Substitution and Elimination Reactions of Alkyl Halides 391

ENVIRONMENTAL CONNECTION: The Birth of the Environmental Movement 392

- **9.1** The S_N2 Reaction **393**
- **9.2** Factors That Affect S_N2 Reactions **398**
- CHEMICAL CONNECTION: Why Are Living Organisms Composed of Carbon Instead of Silicon? 405
- **9.3** The S_N1 Reaction **406**
- **9.4** Factors That Affect S_N1 Reactions **409**
- 9.5 Competition Between S_N2 and S_N1 Reactions 410 PROBLEM-SOLVING STRATEGY 411
 - BIOLOGICAL CONNECTION: Naturally Occurring Alkyl Halides That Defend Against Predators 412
- 9.6 Elimination Reactions of Alkyl Halides 412
- 9.7 The E2 Reaction 413
- 9.8 The E1 Reaction 419

PROBLEM-SOLVING STRATEGY 421

- 9.9 Competition Between E2 and E1 Reactions 422
- **9.10** E2 and E1 Reactions are Stereoselective **423**
 - PROBLEM-SOLVING STRATEGY 425
- **9.11** Elimination from Substituted Cyclohexanes **427**
- **9.12** Predicting the Products of the Reaction of an Alkyl Halide with a Nucleophile/Base **429**
- **9.13** Benzylic Halides, Allylic Halides, Vinylic Halides, and Aryl Halides **433 PROBLEM-SOLVING STRATEGY 434**
 - PROBLEM-SOLVING STRATEGY 437
- 9.14 Solvent Effects 438 CHEMICAL CONNECTION: Solvation Effects 438 ENVIRONMENTAL CONNECTION: Environmental Adaptation 441
- **9.15** Substitution and Elimination Reactions in Synthesis **442**
- 9.16 Intermolecular Versus Intramolecular Reactions 444 PROBLEM-SOLVING STRATEGY 446
- 9.17 DESIGNING A SYNTHESIS II: Approaching the Problem 446 ESSENTIAL CONCEPTS 449 SUMMARY OF REACTIONS 450 PROBLEMS 451
- 10 Reactions of Alcohols, Ethers, Epoxides, Amines, and Sulfur-Containing Compounds 458
- 10.1 Nucleophilic Substitution Reactions of Alcohols: Forming Alkyl Halides 459 CHEMICAL CONNECTION: The Lucas Test 461 GENERAL CONNECTION: Grain Alcohol and Wood Alcohol 462
- 10.2 Other Methods Used to Convert Alcohols into Alkyl Halides 463
- 10.3 Converting an Alcohol Into a Sulfonate Ester 465 MEDICAL CONNECTION: The Inability to Perform an S_N2 Reaction Causes a Severe Clinical Disorder 467
- 10.4 Elimination Reactions of Alcohols: Dehydration 468 PROBLEM-SOLVING STRATEGY 471
 - BIOLOGICAL CONNECTION: Biological Dehydrations 473
- 10.5 Oxidation of Alcohols 474
 GENERAL CONNECTION: Blood Alcohol Concentration 476
 MEDICAL CONNECTION: Treating Alcoholism with Antabuse 476
 MEDICAL CONNECTION: Methanol Poisoning 477

The two chapters in the previous edition on substitution and elimination reactions of alkenes have been combined into one chapter. The recent compelling evidence showing that secondary alkyl halides do not undergo S_N1 solvolysis reactions has allowed this material to be greatly simplified, so now it fits nicely into one chapter.

10.7 10.8 10.9 10.10	Nucleophilic Substitution Reactions of Ethers 477 MEDICAL CONNECTION: Anesthetics 478 Nucleophilic Substitution Reactions of Epoxides 480 CHEMICAL CONNECTION: Crown Ethers—Another Example of Molecular Recognition 484 CHEMICAL CONNECTION: Crown Ethers Can be Used to Catalyze SN ₂ Reactions 485 Arene Oxides 485 ENVIRONMENTAL CONNECTION: Benzo[<i>a</i>]pyrene and Cancer 488 ENVIRONMENTAL CONNECTION: Chimney Sweeps and Cancer 489 Amines Do Not Undergo Substitution or Elimination Reactions 490 BIOLOGICAL CONNECTION: Alkaloids 491 PHARMACEUTICAL CONNECTION: Lead Compounds for the Development of Drugs 491 Quaternary Ammonium Hydroxides Undergo Elimination Reactions 492		
	Thiols, Sulfides, and Sulfonium Ions 494 HISTORICAL CONNECTION: Mustard Gas–A Chemical Warfare Agent 495 MEDICAL CONNECTION: Alkylating Agents as Cancer Drugs 496		
10.12	Methylating Agents Used by Chemists versus Those Used by Cells 496 CHEMICAL CONNECTION: Eradicating Termites 497 MEDICAL CONNECTION: S-Adenosylmethionine: A Natural Antidepressant 498		
10.13	Organizing What We Know About the Reactions of Organic Compounds (Group II) 499		
	ESSENTIAL CONCEPTS 500 SUMMARY OF REACTIONS 501 PROBLEMS 503		
11	Organometallic Compounds 508		The discussion of palladium- catalyzed coupling reactions has
11.1	Organolithium and Organomagnesium Compounds 509		been expanded, and the cyclic
11.2	Transmetallation 511		catalytic mechanisms are shown.
11.3	Organocuprates 512		
11.4	Palladium-Catalyzed Coupling Reactions 515		
11.5	PROBLEM-SOLVING STRATEGY 521		
11.5	Alkene Metathesis 522 HISTORICAL CONNECTION: Grubbs, Schrock, Suzuki, and Heck Receive the Nobel Prize 526 HISTORICAL CONNECTION: The Nobel Prize 526 ESSENTIAL CONCEPTS 527 - SUMMARY OF REACTIONS 527 - PROBLEMS 528		
12	Radicals 532		
12.1	Alkanes are Unreactive Compounds 532		
	GENERAL CONNECTION: Natural Gas and Petroleum 533		
	GENERAL CONNECTION: Fossil Fuels: A Problematic Energy Source 533		
12.2	The Chlorination and Bromination of Alkanes 534 HISTORICAL CONNECTION: Why Radicals No Longer Have to Be Called Free Radicals 536		
12.3	Radical Stability Depends on the Number of Alkyl Groups Attached to the Carbon with the Unpaired Electron 536		
12.4	The Distribution of Products Depends on Probability and Reactivity 537	Masto	ringChemistry*
12.5	The Reactivity–Selectivity Principle 539		- /
	PROBLEM-SOLVING STRATEGY 541		anic Chemistry
12.6 12.7			Chemistry tutorials guide you through est topics in chemistry with self-paced
12.7	The Starson hamistry of Dadian Substitution and Dadian Addition Departience FAC		nat provide individualized coaching. These
12.9			e, in-depth tutorials are designed to 1 with hints and feedback specific to your
	CHEMICAL CONNECTION: Cyclopropane 550	, ndividual	misconceptions. For additional practice on
	beoranna A or articlo in. More r ractice with Multistep Oynthesis 550		Curved Arrows in Radical Systems, go to Chemistry where the following tutorials
12.11	Radical Reactions in Biological Systems 552	re availat	ble:
	NUTRITIONAL CONNECTION: Decaffeinated Coffee and the Cancer Scare 553 NUTRITIONAL CONNECTION: Food Preservatives 555		Arrows in Radical Systems: Interpreting
	NUTRITIONAL CONNECTION to Changelate a Lingth Food?		l Arrows I Arrows in Radical Systems: Drawing
12.12	Radicals and Stratospheric Ozone 556		Arrows
	MEDICAL CONNECTION: Artificial Blood 558		Arrows in Radical Systems: Drawing
	ESSENTIAL CONCEPTS 558 SUMMARY OF REACTIONS 559 PROBLEMS 559	nesona	ance Contributors

TUTORIAL Drawing Curved Arrows in Radical Systems 563

In addition to the more than 170 spectroscopy problems in Chapters 13 and 14, there are 60 additional spectroscopy problems in the *Study Guide and Solutions Manual*.



PART Identification of Organic Compounds 566 FOUR

Mass Spectrometry; Infrared Spectroscopy; UV/Vis Spectroscopy 567

- 13.1 Mass Spectrometry 569
- 13.2 The Mass Spectrum Fragmentation 570
- **13.3** Using The *m*/*z* Value of the Molecular Ion to Calculate the Molecular Formula **572 PROBLEM-SOLVING STRATEGY 573**
- **13.4** Isotopes in Mass Spectrometry **574**
- 13.5 High-Resolution Mass Spectrometry Can Reveal Molecular Formulas 575
- 13.6 The Fragmentation Patterns of Functional Groups 575
- 13.7 Other Ionization Methods 583
- 13.8 Gas Chromatography–Mass Spectrometry 583 GENERAL CONNECTION: Mass Spectrometry in Forensics 583
- 13.9 Spectroscopy and the Electromagnetic Spectrum 583
- 13.10 Infrared Spectroscopy 585
- 13.11 Characteristic Infrared Absorption Bands 588
- 13.12 The Intensity of Absorption Bands 589
- 13.13 The Position of Absorption Bands 590 GENERAL CONNECTION: The Originator of Hooke's Law 590
- **13.14** The Position and Shape of an Absorption Band is Affected by Electron Delocalization and Hydrogen Bonding **591**
 - PROBLEM-SOLVING STRATEGY 593
- 13.15 C—H Absorption Bands 595
- **13.16** The Absence of Absorption Bands **598**
- 13.17 Some Vibrations are Infrared Inactive 599
- 13.18 How to Interpret an Infrared Spectrum 600
- 13.19 Ultraviolet and Visible Spectroscopy 602 GENERAL CONNECTION: Ultraviolet Light and Sunscreens 603
- **13.20** The Beer–Lambert Law **604**
- **13.21** The Effect of Conjugation on λ_{max} 605
- 13.22 The Visible Spectrum and Color 606 CHEMICAL CONNECTION: What Makes Blueberries Blue and Strawberries Red? 607
- 13.23 Some Uses of UV/Vis Spectroscopy 608

ESSENTIAL CONCEPTS 610 PROBLEMS 611

MR Spectroscopy 620

- 14.1 An Introduction to NMR Spectroscopy 620 HISTORICAL CONNECTION: Nikola Tesla (1856–1943) 622
- **14.2** Fourier Transform NMR **623**
- 14.3 Shielding Causes Different Nuclei to Show Signals at Different Frequencies 623
- 14.4 The Number of Signals in an ¹H NMR Spectrum 624 PROBLEM-SOLVING STRATEGY 625
- 14.5 The Chemical Shift Tells How Far the Signal Is from the Reference Signal 626
- **14.6** The Relative Positions of ¹H NMR Signals **628**
- 14.7 The Characteristic Values of Chemical Shifts 629
- 14.8 Diamagnetic Anisotropy 631
- **14.9** The Integration of NMR Signals Reveals the Relative Number of Protons Causing Each Signal **632**
- **14.10** The Splitting of Signals Is Described by the N + 1 Rule **634**
- 14.11 What Causes Splitting? 637
- 14.12 More Examples of ¹H NMR Spectra 639
- 14.13 Coupling Constants Identify Coupled Protons 644 PROBLEM-SOLVING STRATEGY 646
- 14.14 Splitting Diagrams Explain the Multiplicity of a Signal 647
- **14.15** Enantiotopic and Diastereotopic Hydrogens **650**
- 14.16 The Time Dependence of NMR Spectroscopy 652

- 14.17 Protons Bonded to Oxygen and Nitrogen 652
- 14.18 The Use of Deuterium in ¹H NMR Spectroscopy 654
- **14.19** The Resolution of ¹H NMR Spectra **655**
- 14.20 ¹³C NMR Spectroscopy 657 PROBLEM-SOLVING STRATEGY 660
- 14.21 Dept ¹³C NMR Spectra 662
- 14.22 Two-Dimensional NMR Spectroscopy 662
- 14.23 NMR Used in Medicine is Called Magnetic Resonance Imaging 665
- 14.24 X-Ray Crystallography 666 GENERAL CONNECTION: Structural Databases 667 ESSENTIAL CONCEPTS 668 PROBLEMS 669

PART Carbonyl Compounds 685 FIVE

Reactions of Carboxylic Acids and Carboxylic Acid Derivatives 686 The Nomenclature of Carboxylic Acids and Carboxylic Acid Derivatives 688 MEDICAL CONNECTION: Nature's Sleeping Pill 691

- **15.2** The Structures of Carboxylic Acids and Carboxylic Acid Derivatives **692**
- **15.3** The Physical Properties of Carbonyl Compounds **693**
- 15.4 How Carboxylic Acids and Carboxylic Acid Derivatives React 694
- PROBLEM-SOLVING STRATEGY 696 15.5 The Relative Reactivities of Carboxylic Acids and Carboxylic Acid Derivatives 696
- **15.6** Reactions of Acyl Chlorides **698**
- **15.7** Reactions of Esters **701**
- **15.8** Acid-Catalyzed Ester Hydrolysis and Transesterification **702**
- 15.9 Hydroxide-Ion-Promoted Ester Hydrolysis 706 PHARMACEUTICAL CONNECTION: Aspirin, NSAIDs, and COX-2 Inhibitors 707
- 15.10 Reactions of Carboxylic Acids 709 PROBLEM-SOLVING STRATEGY 710
- 15.11 Reactions of Amides 711 BIOLOGICAL CONNECTION: Dalmatians: Do Not Fool with Mother Nature 711
- 15.12 Acid-Catalyzed Amide Hydrolysis and Alcoholysis 712
 HISTORICAL CONNECTION: The Discovery of Penicillin 713
 MEDICAL CONNECTION: Penicillin and Drug Resistance 713
 PHARMACEUTICAL CONNECTION: Penicillins in Clinical Use 714
 BIOLOGICAL CONNECTION: A Semisynthetic Penicillin 714
- 15.13 Hydroxide-Ion-Promoted Hydrolysis of Amides 715 INDUSTRIAL CONNECTION: Synthetic Polymers 715 MEDICAL CONNECTION: Dissolving Sutures 716
- 15.14 Hydrolysis of an Imide: a Way to Synthesize a Primary Amine 716
- 15.15 Nitriles 717
- 15.16 Acid Anhydrides 719
 - GENERAL CONNECTION: What Drug-Enforcement Dogs Are Really Detecting 721
- 15.17 Dicarboxylic Acids 721
- 15.18 How Chemists Activate Carboxylic Acids 723
 15.19 How Cells Activate Carboxylic Acids 724
 CHEMICAL CONNECTION: Nerve Impulses, Paralysis, and Insecticides 727
 ESSENTIAL CONCEPTS 728 SUMMARY OF REACTIONS 729 PROBLEMS 731

16 Reactions of Aldehydes and Ketones • More Reactions of Carboxylic Acid Derivatives 739

- 16.1 The Nomenclature of Aldehydes and Ketones 740 GENERAL CONNECTION: Butanedione: An Unpleasant Compound 742
- **16.2** The Relative Reactivities of Carbonyl Compounds **743**
- **16.3** How Aldehydes and Ketones React **744**

The focus of the first chapter on carbonyl chemistry is all about how a tetrahedral intermediate partitions. If students understand this, then carbonyl chemistry becomes pretty straightforward. I found that the lipid materil that had been put into this chapter in the last edition detracted from the main message of the chapter. Therefore, the lipid material was removed and put into a new chapter exclusively about lipids.

- 16.5 Reactions of Carbonyl Compounds with Hydride Ion 752
- **16.6** More About Reduction Reactions **757**
- 16.7 Chemoselective Reactions 759
- 16.8 Reactions of Aldehydes and Ketones with Nitrogen Nucleophiles
 760 PHARMACEUTICAL CONNECTION: Serendipity in Drug Development
 765
- 16.9 Reactions of Aldehydes and Ketones with Oxygen Nucleophiles 766
 BIOLOGICAL CONNECTION: Preserving Biological Specimens 768
 CHEMICAL CONNECTION: Carbohydrates 770
 PROBLEM-SOLVING STRATEGY 771
- **16.10** Protecting Groups **772**
- 16.11 Reactions of Aldehydes and Ketones with Sulfur Nucleophiles 774
- 16.12 Reactions of Aldehydes and Ketones with a Peroxyacid 774
- 16.13 The Wittig Reaction Forms an Alkene 776 CHEMICAL CONNECTION: β-Carotene 777
- 16.14 DESIGNING A SYNTHESIS IV: Disconnections, Synthons, and Synthetic Equivalents 779
 CHEMICAL CONNECTION: Synthesizing Organic Compounds 781
 PHARMACEUTICAL CONNECTION: Semisynthetic Drugs 781
- **16.15** Nucleophilic Addition to α,β -Unsaturated Aldehydes and Ketones **781**
- 16.16Nucleophilic Addition to α,β-Unsaturated Carboxylic Acid Derivatives785CHEMICAL CONNECTION: Enzyme-Catalyzed Cis-Trans Interconversion785
- 16.17 Conjugate Addition Reactions in Biological Systems 786 MEDICAL CONNECTION: Cancer Chemotherapy 786

ESSENTIAL CONCEPTS 787 SUMMARY OF REACTIONS 788 PROBLEMS 791

Reactions at the \alpha-Carbon 801

- 17.1 The Acidity of an α-Hydrogen 802 PROBLEM-SOLVING STRATEGY 804
- 17.2 Keto–Enol Tautomers 805
- 17.3 Keto-Enol Interconversion 806
- **17.4** Halogenation of the α -Carbon of Aldehydes and Ketones **807**
- **17.5** Halogenation of the α -Carbon of Carboxylic Acids **809**
- 17.6 Forming an Enolate Ion 810
- 17.7 Alkylating the α-Carbon 811
 INDUSTRIAL CONNECTION: The Synthesis of Aspirin 813
 PROBLEM-SOLVING STRATEGY 813
- **17.8** Alkylating and Acylating the α -Carbon Via an Enamine Intermediate **814**
- **17.9** Alkylating the β -Carbon **815**
- **17.10** An Aldol Addition Forms a β -Hydroxyaldehyde or a β -Hydroxyketone **817**
- **17.11** The Dehydration of Aldol Addition Products Forms α,β -Unsaturated Aldehydes and Ketones **819**
- 17.12 A Crossed Aldol Addition 821 MEDICAL CONNECTION: Breast Cancer and Aromatase Inhibitors 823
- **17.13** A Claisen Condensation Forms a β -Keto Ester **824**
- 17.14 Other Crossed Condensations 827
- 17.15 Intramolecular Condensations and Intramolecular Aldol Additions 827
- 17.16 The Robinson Annulation 830

PROBLEM-SOLVING STRATEGY 830

- 17.17 CO₂ Can be Removed from a Carboxylic Acid that has a Carbonyl Group at the 3-Position 831
- 17.18 The Malonic Ester Synthesis: A Way to Synthesize a Carboxylic Acid 833
- 17.19 The Acetoacetic Ester Synthesis: A Way to Synthesize a Methyl Ketone 834
- 17.20 DESIGNING A SYNTHESIS V: Making New Carbon–Carbon Bonds 836
- **17.21** Reactions at the α -Carbon in Living Systems 838
- 17.22 Organizing What We Know About the Reactions of Organic Compounds (Group III) 841

ESSENTIAL CONCEPTS 843 SUMMARY OF REACTIONS 844 PROBLEMS 846

TUTORIAL Synthesis and Retrosynthetic Analysis 854

This chapter was reorganized and rewritten for ease of understanding.

PART Aromatic Compounds 867 SIX

18	Reactions of Benzene and Substituted Benzenes 868
18.1 18.2 18.3	GENERAL CONNECTION: Measuring Toxicity 869 The Nomenclature of Monosubstituted Benzenes 870 GENERAL CONNECTION: The Toxicity of Benzene 871 The General Mechanism for Electrophilic Aromatic Substitution Reactions 871 Halogenation of Benzene 872
18.4 18.5 18.6 18.7	MEDICAL CONNECTION: Thyroxine874Nitration of Benzene874Sulfonation of Benzene875Friedel–Crafts Acylation of Benzene876Friedel–Crafts Alkylation of Benzene877CHEMICAL CONNECTION: Incipient Primary Carbocations879
18.10 18.11 18.12 18.13	BIOLOGICAL CONNECTION: A Biological Friedel-Crafts Alkylation879Alkylation of Benzene by Acylation-Reduction880Using Coupling Reactions to Alkylate Benzene881How Some Substituents on a Benzene Ring Can Be Chemically Changed882The Nomenclature of Disubstituted and Polysubstituted Benzenes884The Effect of Substituents on Reactivity886The Effect of Substituents on Orientation890The Ortho-Para Ratio894
18.15 18.16 18.17 18.18	Additional Considerations Regarding Substituent Effects 894 DESIGNING A SYNTHESIS VI: The Synthesis of Monosubstituted and Disubstituted Benzenes 896 The Synthesis of Trisubstituted Benzenes 898 Synthesizing Substituted Benzenes Using Arenediazonium Salts 900 Azobenzenes 903
18.20	HISTORICAL CONNECTION: Discovery of the First Antibiotic 904 PHARMACEUTICAL CONNECTION: Drug Safety 904 The Mechanism for the Formation of a Diazonium Ion 905 MEDICAL CONNECTION: A New Cancer-Fighting Drug 905 NUTRITIONAL CONNECTION: Nitrosamines and Cancer 906
	Nucleophilic Aromatic Substitution 907 DESIGNING A SYNTHESIS VII: The Synthesis of Cyclic Compounds 909 ESSENTIAL CONCEPTS 910 SUMMARY OF REACTIONS 911 PROBLEMS 913
19	More About Amines • Reactions of Heterocyclic Compounds 924
19.1 19.2	More About Nomenclature 925 More About the Acid–Base Properties of Amines 926 MEDICAL CONNECTION: Atropine 927
19.3 19.4 19.5	Amines React as Bases and as Nucleophiles 927 Synthesis of Amines 929 Aromatic Five-Membered-Ring Heterocycles 929 PROBLEM-SOLVING STRATEGY 931
19.6 19.7	Aromatic Six-Membered-Ring Heterocycles 934 Some Heterocyclic Amines Have Important Roles in Nature 939 PHARMACEUTICAL CONNECTION: Searching for Drugs: An Antihistamine, a Nonsedating Antihistamine, and a Drug for Ulcers 940 MEDICAL CONNECTION: Porphyrin, Bilirubin, and Jaundice 943
19.8	Organizing What We Know About the Reactions of Organic Compounds (Group IV) 943
	ESSENTIAL CONCEPTS 944 SUMMARY OF REACTIONS 945 PROBLEMS 946

MasteringChemistry[®]

for Organic Chemistry

MasteringChemistry tutorials guide you through the toughest topics in chemistry with self-paced tutorials that provide individualized coaching. These assignable, in-depth tutorials are designed to coach you with hints and feedback specific to your individual misconceptions. For additional practice on Synthesis and Retrosynthetic Analysis, go to MasteringChemistry where the following tutorials are available:

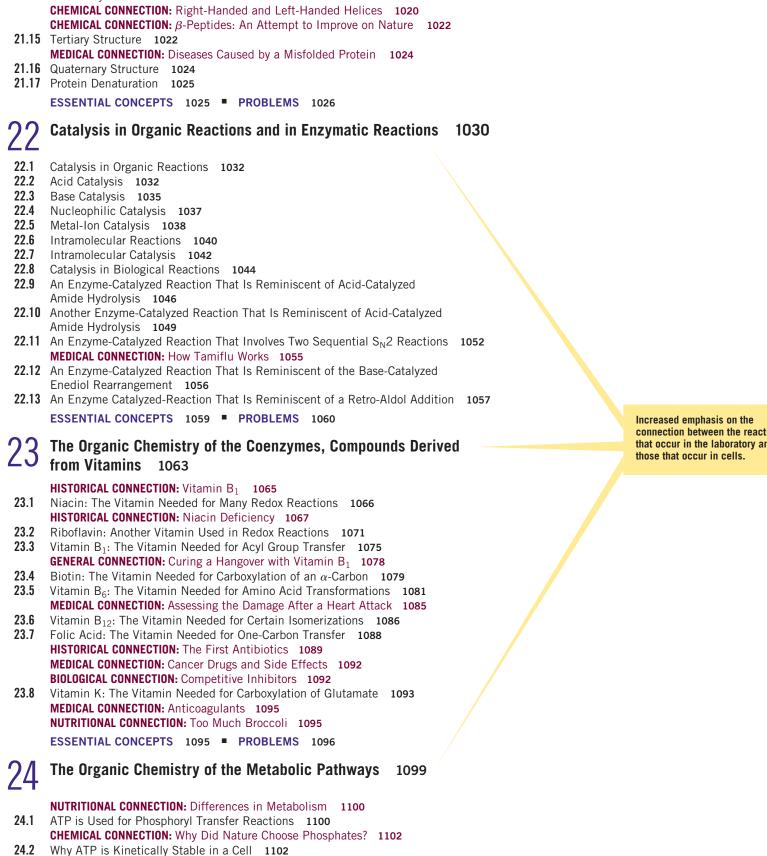
- Synthesis and Retrosynthetic Analysis: Changing the Functional Group
- Synthesis and Retrosynthetic Analysis: Disconnections
- Synthesis and Retrosynthetic Analysis: Synthesis of Carbonyl Compounds

PART Bioorganic Compounds 949 SEVEN

20 The Organic Chemistry of Carbohydrates 950

- 20.1 Classifying Carbohydrates 951
- **20.2** The D and L Notation **952**
- 20.3 The Configurations of Aldoses 953
- 20.4 The Configurations of Ketoses 955
- 20.5 The Reactions of Monosaccharides in Basic Solutions 956
- 20.6 Oxidation–Reduction Reactions of Monosaccharides 957
- 20.7 Lengthening the Chain: The Kiliani–Fischer Synthesis 958
- 20.8 Shortening the Chain: The Wohl Degradation 959
- MEDICAL CONNECTION: Measuring the Blood Glucose Levels in Diabetes 960
- **20.9** The Stereochemistry of Glucose: The Fischer Proof **960**
- GENERAL CONNECTION: Glucose/Dextrose 962
- 20.10 Monosaccharides Form Cyclic Hemiacetals 962
- 20.11 Glucose is the Most Stable Aldohexose 965
- 20.12 Formation of Glycosides 967
- 20.13 The Anomeric Effect 968
- 20.14 Reducing and Nonreducing Sugars 969
- 20.15 Disaccharides 969 NUTRITIONAL CONNECTION: Lactose Intolerance 971 MEDICAL CONNECTION: Galactosemia 971 BIOLOGICAL CONNECTION: A Toxic Disaccharide 972
- 20.16 Polysaccharides 973 MEDICAL CONNECTION: Why the Dentist is Right 974 BIOLOGICAL CONNECTION: Controlling Fleas 975
- 20.17 Some Naturally Occurring Compounds Derived from Carbohydrates 976
 MEDICAL CONNECTION: Resistance to Antibiotics 976
 MEDICAL CONNECTION: Heparin–A Natural Anticoagulant 977
 HISTORICAL CONNECTION: Vitamin C 978
- 20.18 Carbohydrates on Cell Surfaces 978
- 20.19 Artificial Sweeteners 979 NUTRITIONAL CONNECTION: Acceptable Daily Intake 981 ESSENTIAL CONCEPTS 981
 SUMMARY OF REACTIONS 982
 PROBLEMS 983
 - Amino Acids, Peptides, and Proteins 986
- 21.1 The Nomenclature of Amino Acids 987
 - NUTRITIONAL CONNECTION: Proteins and Nutrition 99121.2 The Configuration of Amino Acids 991
 - MEDICAL CONNECTION: Amino Acids and Disease 992 PHARMACEUTICAL CONNECTION: A Peptide Antibiotic 992
 - 21.3 Acid–Base Properties of Amino Acids 993
 - 21.4 The Isoelectric Point 995
 - 21.5 Separating Amino Acids 996 GENERAL CONNECTION: Water Softeners: Examples of Cation-Exchange Chromatography 1000
 - **21.6** Synthesis of Amino Acids 1000
 - 21.7 Resolution of Racemic Mixtures of Amino Acids 1002
 - Peptide Bonds and Disulfide Bonds 1003 MEDICAL CONNECTION: Diabetes 1006 CHEMICAL CONNECTION: Hair: Straight or Curly? 1006
 - **21.9** Some Interesting Peptides 1006
 - 21.10 The Strategy of Peptide Bond Synthesis: N-Protection and C-Activation 1007
 - **21.11** Automated Peptide Synthesis **1010**
 - **21.12** An Introduction to Protein Structure **1013**
 - BIOLOGICAL CONNECTION: Primary Structure and Taxonomic Relationship 1013
 - 21.13 How to Determine the Primary Structure of a Polypeptide or a Protein 1013 PROBLEM-SOLVING STRATEGY 1015

New art adds clarity.



- 24.3 The "High-Energy" Character of Phosphoanhydride Bonds 1102
- 24.4 The Four Stages of Catabolism 1104

21.14 Secondary Structure 1019

- 24.5 The Catabolism of Fats: Stages 1 and 2 1105
- The Catabolism of Carbohydrates: Stages 1 and 2 1108 24.6 **PROBLEM-SOLVING STRATEGY 1111**

connection between the reactions that occur in the laboratory and

NUTRITIONAL CONNECTION: Fats Versus Carbohydrates as a Source of Energy 1112

- 24.7 The Fate of Pyruvate 1112
- **24.8** The Catabolism of Proteins: Stages 1 and 2 1113 **MEDICAL CONNECTION:** Phenylketonuria (PKU): An Inborn Error of Metabolism 1114 **MEDICAL CONNECTION:** Alcaptonuria 1115
- 24.9 The Citric Acid Cycle: Stage 3 1115
- 24.10 Oxidative Phosphorylation: Stage 4 1118 NUTRITIONAL CONNECTION: Basal Metabolic Rate 1119
- 24.11 Anabolism 1119
- 24.12 Gluconeogenesis 1120
- 24.13 Regulating Metabolic Pathways 1122

24.14 Amino Acid Biosynthesis 1123

ESSENTIAL CONCEPTS 1124
PROBLEMS 1125

25	The Organic Chemistry of Lipids	1127
LJ		

The lipid material previously in
the chapter on carboxylic acids
and their derivatives has been
moved into this new chapter. The
discussion of terpenes from the
metabolism chapter has also been
moved into this chapter, along with
some new material.

25.1	Fatty Acids Are Long-Chain Carboxylic Acids	1128
	NUTRITIONAL CONNECTION: Omega Fatty Acids	1129

- 25.2 Waxes Are High-Molecular-Weight Esters 1130
 25.3 Fats and Oils Are Triglycerides 1130
 NUTRITIONAL CONNECTION: Olestra: Nonfat with Flavor 1132
 BIOLOGICAL CONNECTION: Whales and Echolocation 1132
- 25.4 Soaps and Micelles 1132
- 25.5 Phospholipids Are Components of Cell Membranes 1134 BIOLOGICAL CONNECTION: Snake Venom 1136 MEDICAL CONNECTION: Multiple Sclerosis and the Myelin Sheath 1137
- **25.6** Prostaglandins Regulate Physiological Responses **1137**
- **25.7** Terpenes Contain Carbon Atoms in Multiples of Five **1139**
- 25.8 How Terpenes Are Biosynthesized 1141 MEDICAL CONNECTION: How Statins Lower Cholesterol Levels 1142 PROBLEM-SOLVING STRATEGY 1144 CHEMICAL CONNECTION: Protein Prenylation 1146
- **25.9** How Nature Synthesizes Cholesterol 1147
- 23.5 How Nature Synthesizes Cholesteron 1147 25.10 Steroids 1148 MEDICAL CONNECTION: One Drug—Two Effects 1149
- 25.11 Synthetic Steroids 1150
 - ESSENTIAL CONCEPTS 1151

 PROBLEMS 1152
- 26 The Chemistry of the Nucleic Acids 1155
- 26.1 Nucleosides and Nucleotides 1155 HISTORICAL CONNECTION: The Structure of DNA: Watson, Crick, Franklin, and Wilkins 1158 BIOLOGICAL CONNECTION: Cyclic AMP 1159
- 26.2 Nucleic Acids Are Composed of Nucleotide Subunits 1159
- 26.3 The Secondary Structure of DNA 1161
- 26.4 Why DNA Does Not Have A 2'-OH Group 1163
- 26.5 The Biosynthesis of DNA Is Called Replication 1163
- 26.6 DNA and Heredity 1164
- PHARMACEUTICAL CONNECTION: Natural Products That Modify DNA 1165
- **26.7** The Biosynthesis of RNA Is Called Transcription **1165**
- BIOLOGICAL CONNECTION: There Are More Than Four Bases in DNA 1166
- **26.8** The RNAs Used for Protein Biosynthesis **1167**
- 26.9 The Biosynthesis of Proteins Is Called Translation 1169
 MEDICAL CONNECTION: Sickle Cell Anemia 1171
 MEDICAL CONNECTION: Antibiotics That Act by Inhibiting Translation 1172
- 26.10 Why DNA Contains Thymine Instead of Uracil 1173
- MEDICAL CONNECTION: Antibiotics Act by a Common Mechanism117426.11Antiviral Drugs1174
 - HISTORICAL CONNECTION: Influenza Pandemics 1175
- 26.12 How the Base Sequence of DNA Is Determined 1175
- 26.13 Genetic Engineering 1177

PART Special Topics in Organic Chemistry 1181 EIGHT

27 Syntl

Synthetic Polymers 1182

- 27.1 There Are Two Major Classes of Synthetic Polymers 1183
- 27.2 An Introduction To Chain-Growth Polymers 1184
 27.3 Radical Polymerization 1184
 INDUSTRIAL CONNECTION: Teflon: An Accidental Discovery 1187
 ENVIRONMENTAL CONNECTION: Recycling Symbols 1189
- 27.4 Cationic Polymerization 1189
- 27.5 Anionic Polymerization 1192
- 27.6 Ring-Opening Polymerizations 1193
- 27.7 Stereochemistry of Polymerization Ziegler–Natta Catalysts 1195
- **27.8** Polymerization of Dienes **1196**
- 27.9 Copolymers 1198

PHARMACEUTICAL CONNECTION: Nanocontainers 1198

- 27.10 An Introduction to Step-Growth Polymers 1199
 27.11 Classes of Step-Growth Polymers 1200
 MEDICAL CONNECTION: Health Concerns: Bisphenol A and Phthalates 1202
 INDUSTRIAL CONNECTION: Designing a Polymer 1203
- 27.12 Physical Properties of Polymers 1204 NUTRITIONAL CONNECTION: Melamine Poisoning 1205
- **27.13** Recycling Polymers **1206**
- 27.14 Biodegradable Polymers 1207

ESSENTIAL CONCEPTS 1208 PROBLEMS 1208

Pericyclic Reactions 1212

- 28.1 There Are Three Kinds of Pericyclic Reactions 1213
- 28.2 Molecular Orbitals and Orbital Symmetry 1215
- 28.3 Electrocyclic Reactions 1218
- 28.4 Cycloaddition Reactions 1224
- 28.5 Sigmatropic Rearrangements 1227
- 28.6 Pericyclic Reactions in Biological Systems 1232
 CHEMICAL CONNECTION: Bioluminescence 1233
 NUTRITIONAL CONNECTION: The Sunshine Vitamin 1234
 NUTRITIONAL CONNECTION: Animals, Birds, Fish—And Vitamin D 1235
- **28.7** Summary of the Selection Rules for Pericyclic Reactions **1235**

ESSENTIAL CONCEPTS 1236 PROBLEMS 1236

Appendices A-1

- PKA VALUES A-1
- I KINETICS A-3
- III SUMMARY OF METHODS USED TO SYNTHESIZE A PARTICULAR FUNCTIONAL GROUP A-8
- IV SUMMARY OF METHODS EMPLOYED TO FORM CARBON-CARBON BONDS A-11
- V SPECTROSCOPY TABLES A-12
- VI PHYSICAL PROPERTIES OF ORGANIC COMPOUNDS A-18

ANSWERS TO SELECTED PROBLEMS ANS-1

GLOSSARY G-1

CREDITS C-1