

Welcome to Biochemistry! You are embarking on a semester's study of what is, in my humble opinion, the most fascinating material in the intellectual world -- the molecular analysis of life as we know it on earth. This course will touch on many topics that you've visited before, but we are likely to be looking at these familiar issues in new ways, and I hope that you'll be delighted to see how many nuggets of knowledge from earlier coursework fit together in the new analyses you'll be undertaking this term.

I've designed this course, but I strive to create an environment where you will be sufficiently comfortable and sufficiently excited that you will really make this YOUR course, rather than mine. What you are reading is what I call my "syllabook" -- it is a (lengthy) description of the philosophy motivating my design of this course as well as the usual course logistics and scheduling details. It is my sincere hope that it should help you

- get a sense for how I have structured the class
- begin to formulate questions about the course and about Biochemistry
- prepare yourself for what I hope will be an exciting and challenging exploration of the Chemistry of Life.

**What I want to achieve in this course this semester:**

- What we term "biochemistry" includes a huge -- HUGE -- realm of knowledge. I expressly do NOT hope to teach you all of it. Rather, I hope to introduce you to a selection of topics in biochemistry, and through these topics, to a set of analytical approaches for probing the molecular underpinnings of life. It is my intent that by mastering these analytical tools that you will be prepared to master the many domains of biochemical topics that we won't have time to explore this term.
- Specifically, I want to guide you to a deep-seated understanding of the relationships between molecular structure and biological function for the major classes of biological macromolecules, and for proteins in particular.
  - First, you will learn to *explain* how elements of the structure of a macromolecule support its function or properties.
  - By the end of this course, you should have begun to develop the ability to *predict* the change in function that might accompany a particular change in structure of a molecule that you have never seen before. To approach this goal, I intend to introduce you to lots of examples of biologically-important macromolecules, with many of these 'molecular case studies' drawn from the primary research literature.
  - Finally, you should develop the ability to explain how these structure/function relationships form the basis for the real-time control of macromolecular machines in living biological systems.
- Beyond the day-to-day content of this course, I hope to motivate your desire for more sophisticated ways of thinking about biochemical data and biochemical systems. This objective will also lead us into the research literature, and involve the use of 'professional-grade' tools for laboratory analysis, molecular modeling, and data analysis. The goal of stimulating your curiosity and your hunger for more sophisticated knowledge underlies much of the format of this course, and is the greatest goal (in my opinion) that any educator can strive for. I hope that we can achieve it together.

**Who I am and how to get in touch with me:**

Instructor: Peter Kuhlman  
Office: Ebaugh 114  
Phone: 587-6698 (if you let this ring, it will get forwarded to my cell phone -- or voicemail if I'm not immediately available -- if you really need to get in touch with me, don't hesitate to call)  
E-mail: kuhlman@denison.edu  
Office hours: To optimize our opportunities to meet outside of class hours to discuss any questions or difficulties you are having, I don't have regular "office hours", but instead set up individual or group appointments. I will make every effort to ensure that you and I can find a time to meet outside of class hours to discuss any topics that are of concern to you. *I will be available for at least 3 hours of appointments each week*, as long as they are scheduled by the end of the preceding week.

### **Course resources:**

Texts: *Biochemistry, Sixth Edition*, Berg/Tymoczko/Stryer (2006).

The laboratory manual will be handed out on the first day of lab (free of cost to you).

Problem sets: A variety of sample problems will be available to you in order to help you review, deepen your understanding, and prepare for tests. The first installment is online at

<http://personal.denison.edu/~kuhlman/courses/biochem/samples/>

I will sometimes point you to specific problems from these online question banks, but they are always available for you to pursue on your own. As they come to my attention, I will also publish links to other online Biochemistry resources at

<http://personal.denison.edu/~kuhlman/courses/>

I encourage you to take full advantage of them!

### **How we'll achieve the course goals:**

Research into how humans learn suggests that education is most effective if we encounter a new concept or skill in several different ways, followed by the opportunity to practice it and then to apply it to new situations. I have tried to embrace this theory in my design of this course. For most topics, you'll have the opportunity to read about them, to hear me talk about them, to think about them on your own and in class, to answer questions on them, to apply them both in your mind and with your hands, and then to try them out in novel situations. In order to give you multiple ways to explore new concepts, I will structure some of our class meetings differently than a conventional lecture. And because it is my fundamental assumption that I am here as your *partner* in learning, I invite you to join me in taking responsibility for your education. For instance, I expect that you, as an experienced student at Denison, have learned to read textbooks in a sophisticated fashion, and I will therefore typically NOT simply re-cover the material in your text. Rather, I expect you to come to class each day prepared to actively engage the material, and I will intentionally spend much of our class time in ways that I feel *complement* the text rather than reiterate it. I will be disappointed if you have not spent enough time outside of class with your text and notes in order to understand and be able to contribute to the class conversation. To help focus your reading and thinking, I will often provide comments and questions to follow up on the day's class and prepare for the next session.

I will structure our class time in ways intended to help you review, reinforce, and synthesize the material in the text. I will do some lecturing, we will do some (both large and small) group discussions, and I will at times ask probing questions. There will be opportunities over the course of the term for you to practice making observations and reaching conclusions on the basis of those observations. In keeping with my goals for this course, there will be many times when I will ask you to make predictions of the behavior of biochemical systems on the basis of what you have already learned. Throughout, I will be looking more for evidence that you are trying to learn and to think than for evidence that you have the "right" answers. I will ask questions *not* to be cruel or "tough", but rather because I believe it is my job to help you *learn to learn* and *learn to think critically*. That said, please note that although I will endeavor to structure this class to maximize your benefit from it, I recognize that I am fallible (!) and that we don't all learn in the same ways, so I strongly encourage you to share your feelings about the class with me as we go along!

This class format is most effective when every member of the class is engaged. If class members either dominate, distract from, or withdraw from the class, we are all likely to learn less. I will therefore base part of your grade on my evaluation of your contribution to the class discourse. That doesn't mean that you have to say a lot in class, or that every comment or answer you give has to be "right". On the contrary, we often learn much more from our mistaken answers than from our correct ones. Accordingly, I will note only whether your comments and answers indicate (a) that you are coming to class prepared and (b) that you are thinking. Of course, this also means that you must generally be present in class (and you must arrive on time), for if you aren't there, it will be difficult for you to contribute to the class and to show me that you're exploring and thinking.

More specifically, if you *must* be absent from any laboratory or exam day, please see me beforehand if at all possible, and in any case be prepared to rigorously justify your absence (with supporting documentation from, for example, Student Health or the Academic Support office). Absence on a test date will result in a score of zero for that test if you have not *previously* made an arrangement with me to take the test at another time. You're responsible for all class material whether you're present at all class meetings or not, so be sure to make arrangements with me and with your classmates to obtain the information that we covered in class on any days that you are not present. Having said all that, of course, I realize that life has a tendency to bring us surprises at inopportune times; I will try in every situation to be as understanding and creative in

accommodating your unique constraints as I can without compromising my ability to be fair to the class as a whole.

In order to extend your classroom learning experiences, to give you exposure to useful tools and techniques, and to give you a sense for what sorts of questions I am apt to put on the tests before you sit down to take them, I will occasionally recommend problems from the textbook or give you challenge questions. In addition, I have compiled a long list of questions to stimulate your studying and reflecting. These are available online at the URL listed above under "Course Resources".

I will strive to keep the format and content of this class flexible so that I can respond to your needs and interests as we go through the semester. Consequently, you should view the course calendar and even the grading scheme as approximations of the final outcomes at the end of the course. I agree with past students that this flexibility enhances my ability to make this Biochemistry course *your* Biochemistry course. I also realize, however, that this very flexibility can be unsettling, and I will therefore also strive to make it very clear what my view of the class structure and priorities are. Furthermore, in order to make changes to the course on the fly, we need to be in ready communication, and so I ask that you make it a habit to *check your e-mail daily* in case we need to be in touch about class-related matters. In return, I will make every effort to respond to *your* email within a day of receiving it.

Several of the assignments for this course involve writing. In science, as in nearly all aspects of professional and personal life, clear and effective communication skills are a tremendous asset. I encourage each of you to take advantage of the campus Writing Center to improve your written communication skills. Although there is a widespread perception that writing in the sciences is somehow "different" from writing in other disciplines, I regard that as a myth. Styles certainly differ between fields, but the fundamentals of good communication are nearly universal. The Center is a free resource available to all Denison students. Student writing consultants from many majors help writers one-on-one in all phases of the writing process, from deciphering the assignment, to discussing ideas, to developing an argument, to finalizing a draft. Because proofreading is a last step in that process, writers should leave plenty of time for getting their ideas right before expecting proofreading help. Consultants also can help writers with personal documents, like job and internship applications. The Center is located on the fourth floor of Barney-Davis Hall; a satellite location is in the Reference section of the Library (the Entry level). Go to the Writing Center website at <http://www.denison.edu/writingctr/> to see their current hours or make an appointment to meet with a tutor.

Finally, I think that it is important for you to learn how to decide which of the many, many bits of information that you encounter in this class are more important and which are less important. Similarly, I want to encourage you to reflect on which aspects of the material *I* think are important and am apt to stress on exams. To stimulate these kinds of higher-order learning, I'll give you one extra point for every exam question that you predict I will ask. Of course, I don't expect you to come up with the exact questions, but with some work I think that you'll be able to predict the *types* of questions I'll ask. I will accept up to four of these sample questions for each exam. In order to receive credit for your submissions, you must send me *sample questions and correct answers by email before the test is administered*. Please ask me about this in class if you have any questions.

### **Building on your previous knowledge**

This course has an uncommonly long list of prerequisite courses: 2 semesters of General Chemistry, 2 semesters of Organic Chemistry, one semester of Introductory Biology, and one semester of Cellular and Molecular Biology. And we really will build explicitly and implicitly on material from all of those courses. To underscore the importance of those prerequisites and to ensure that you learned material from those courses that is fundamental to this course, you will take a Fundamentals Test as an element of this course. Although your score on the Fundamentals Test will not be reflected in your final course grade, *failure to achieve a score of at least 70% on the Fundamentals Test will result in a grade penalty of 18% of the possible points for this class. This penalty will prevent you from receiving a grade above a B- for this course.* Because I place such a high value on your competency with this material, the test will be offered weekly for eight weeks so that you have ample opportunity to solidify your knowledge and multiple chances to demonstrate to me that you have mastered the fundamental concepts you will need to build on in this course. Note, however, that while I'll give you lots of opportunities to re-take the test, it will be up to you to take the initiative to do so. The second-to-last page of this Syllabook provides a list of the concept areas you will be responsible for.

### **Laboratory participation:**

The laboratory component of this course is an integral part of it. This is reflected in its large numerical weight in the overall grade scheme of the course. To pass this course, you **MUST** complete all assigned laboratory work. Period.

A schedule of the lab exercises appears later in this syllabook. As noted above, you'll receive the laboratory manual on the first day of lab.

### **Learning from your mistakes**

I believe very firmly that learning is an iterative process; very few of us get things exactly right the first time through, and there is often a great opportunity for learning in repeating an assignment after receiving feedback on it. This philosophy underlies much of my approach to grading and point values for assignments in this class. I want you to feel that you can learn and benefit from your mistakes, that you will be rewarded for getting it right the second time around. Accordingly, if you ever want to revise an assignment and submit it for reconsideration, you are welcome to come and discuss that with me. And for *any* assignment on which you receive a grade of 50% or less, you are strongly encouraged to revise and submit the assignment for regrading.

In the same vein, if you receive any assignment back on which any part of one of your answers has been marked in orange highlighter, you should interpret this as an invitation to revise your answer and submit it for regrading. Typically this will indicate that I had trouble understanding the logic behind your answer, or felt that you answered a different question from the one I asked. In any event, this indicates that I think you have a better grasp on the material than your answer shows, and I'd like to give you another chance to demonstrate your mastery.

In any resubmission situation, please

- ensure that you include the original graded version of the assignment so that I can compare my original comments with your revised version. Resubmissions that are not accompanied by the original version will not be regraded. I strongly encourage you to make a copy of your original assignment before giving it back to me, so that you can study from it. *Resubmissions get the lowest priority on my grading to-do list, and I won't guarantee getting them back to you before the end of finals week.*
- ensure that you carefully address the deficiencies of the original. I try to put care and time into the comments on your assignments when I hand them back, with the intent that my comments will help you think more carefully about your work. Revised versions that come back to me with the original problems un-addressed are likely to receive a less-than-generous review. If you have any questions about my comments, by all means come to talk with me about them *before* making your revisions.
- be aware that I grade *at least as stringently* on revisions as on the originals -- on the revision, I assume that you understand my expectations, so I am less likely to give you the benefit of the doubt when your intent or procedure is not clear from your work.
- include with your resubmitted assignment a brief description of what you will do differently for the next test or assignment. That is to say, beyond the errors that led to a lower-than-desired score, what can you change about the way you study or the way that you are thinking about this course that may improve your performance the next time around? Resubmissions that are not accompanied by a description of how you improve your learning habits will not be regraded.
- understand that your final grade for the assignment will be the average of your original and revised grades. With a substantial improvement over the original assignment, this can make a very meaningful difference in your overall grade.

Finally, it would be naive to think that simply copying down answers off of someone else's paper, or off of the answer key, constitutes learning from your mistakes. While I won't be able to judge whether a correctly revised answer shows that you've *really learned* the subject matter, *you* should be able to make that assessment. I strongly encourage you to ask yourself that question honestly every time that you hand in revised answers. If you are not honest with yourself about this, you could get a rude surprise on the final exam, which is worth more points than all the other tests put together, and on which you only get one opportunity to show me what you truly have learned.

### **Due dates and deadlines**

Let's face it. We're all busy, and we all find ourselves in the nasty situation from time to time where we have more things to do than we have time to get them done. It's a very real part of the modern lifestyle. With that in mind, and in an effort to avoid holding you to a standard that's higher than the one to which I hold myself, here's my policy on handing things in on time: if you hand any assignment in after the deadline but before I have time to grade it, there will be no cost for your tardiness. If you hand in an assignment after the due date, and up to one week *after I have graded* that assignment for your classmates, there will be an automatic 12% deduction in your score (that is, the max you can get for a late submission is 88% instead of 100%). I hope that you never find yourself turning things in later than that, but if you do, your score will drop an additional 12% for every additional week *after I grade* the assignment.

### **Special Needs**

Different students come to this class with different training, different backgrounds, and different abilities. If you feel that because of personal factors you would benefit from some modification of course procedures, such as special test-taking arrangements, I ask that you contact me privately *at your earliest convenience*. Also, please note that reading is the pre-eminent way of taking in information in science. If you have difficulty reading, please make an appointment to see me at your earliest opportunity.

I will work with you and with the Academic Support and Enrichment Center (Doane Hall, room 102) to optimize your learning experience. Certain accommodations will require verification of disability based on documentation on file in the Academic Support office.

### **Academic and personal honesty**

My fundamental assumption about this class is that you are here to learn. All feedback that I provide you, all learning opportunities that I set up, are predicated on that assumption. For you to effectively learn, and for me to effectively advise you on your learning, you must be honest with me and with yourself about what you do and do not know. I will attach no stigma to your admission that you don't understand something. Indeed, that admission is the first step down the road to mastery of this course.

In 2008, the DCGA and the Denison University faculty signed off on a set of policies envisioned and largely crafted by Denison students. This set of policies, known as the Code of Academic Integrity, "provides a means of assuring that values essential to learning—trust, responsibility, and ethics—are promoted and maintained by all members of the Denison community." (this quote is taken from the description of the Code at <http://www.denison.edu/about/integrity.html>) I encourage you to become well acquainted with the Code; it not only governs how Denison deals with academic dishonesty (more on that in a minute), but it also provides a call to take greater personal responsibility for creating the learning environment that you want to experience at Denison.

If the atmosphere of trust and learning that I try to create in this class is compromised by individuals behaving in an academically dishonest manner -- for instance, passing off someone else's work as their own - I will be deeply disappointed and quite upset. Academic dishonesty is tantamount to intellectual theft. This standard applies to all work ranging from simple lab assignments to the final exam. I recommend that you carefully read the Denison University Code of Academic Integrity (at the URL listed above), the Academic Dishonesty Policy as printed in the student handbook, and the section in the Bedford Handbook entitled "Citing sources" on pages 592-608. Neither ignorance nor carelessness is an acceptable defense in cases of plagiarism.

The grade penalty associated with a confirmed case of dishonesty will ordinarily be a score of zero for that assignment. Furthermore, by Denison policy, I must refer every act of academic dishonesty to the Associate Provost, and violations may result in failure of the course, suspension, or expulsion.

Don't be foolish. Engaging in dishonest behavior in order to bring your grade up from a B to an A, or to save yourself some time, is simply not worth the cost. It saddens me every time that I catch a student cutting corners like this, because the cost is so disproportionate to the potential gain. And I'm sorry to say that I've had to turn in at least one student for academic dishonesty every semester for the past several years. Don't join their ranks. Be proud of what you've learned, not what you've gotten away with. Be honest.

**How will your learning be assessed?**

As I currently envision the course (and subject to feedback from you), there will be three types of opportunities (in addition to class discussions) for you to demonstrate to me your mastery of biochemical concepts -- tests, the laboratory exercises, and a final exam:

Laboratory	225 points	(see explanation later in this syllabook)
Subjective evaluation of intellectual engagement and contribution to the class	50 points	
Hour-long tests	4 x 50 pts =	200 points
Final exam (including a take-home portion)	250 points	
<b>TOTAL</b>	<b>775 points</b>	

**Please note:**

- The four hour exams only account for 28% of your grade in this course, and the final exam for 34%. I've done this for three reasons. First, it makes each in-class exam significantly shorter; I want to test how well you understand the material, not how quickly you can write. Second, it allows you to learn from your mistakes -- you can do poorly on a test, but if you learn what you did wrong and you address your deficiencies, then you will have the chance to score well on the final and come out of the class with a grade that reflects *what you learned in the end*. Third, this scoring system reflects my belief that what you know at the end of the semester is really important -- I don't want to reward you for cramming for one test and then immediately forgetting the material. Rather, I want to reward you for truly learning the material and for integrating each new topic into your mental picture of how the world works.
- As noted above, I feel it is very important that you have retained key elements of content knowledge from the courses that stand as prerequisites to this course. Consequently, if you cannot correctly answer at least 70% of the questions on the Fundamentals Test after the maximum number of allowed attempts, then I will assess a grade penalty of 18% of the total possible points in the course. This penalty will make it impossible to earn a grade above B- for this course if you haven't demonstrated competency in these key elements of those previous courses.
- Grades will be assigned *roughly* according to the scale below. Note that I may change the scale so that it more accurately reflects what I feel to be the performance of members of the class. For instance, if everyone does poorly on the tests and if I conclude that it is because I did something wrong, then I will likely adjust all grades upward from the following distribution. On the other hand, if I think that the tests are sufficiently challenging but everyone does very well and is earning "A" marks, then I will be pleased as punch and am *unlikely* to change the scale. For the most part, you may consider these to be the most demanding standards that I am apt to apply.

90% and up	flavors of A
80 to 89%	flavors of B
70 to 79%	flavors of C
60 to 66%	flavors of D
below 60%	F

- Finally, I wish to make clear my interpretation of letter grades. First, let me say I recognize that the way we assign grades in American higher education is not perfect, or even wonderful. It is, however, a necessity of our education system, and a way to allow communication between student, faculty, and outside constituents -- employers, med school admissions committees, etc -- about how well you met the goals of this class. I will try very hard to ensure that all grades that I give in this class are based on careful, principled evaluation. I also try to ensure that I apply fair standards to all students in my class. Again, however, I recognize that there are many desirable learning attributes -- persistence, creativity, and positive outlook, to name only a few -- that aren't well captured in a simple letter grade. If you ever ask me for a narrative evaluation or letter of recommendation, I'll be sure to comment at length on these other facets of learning.

Second, and in the same vein, I want to acknowledge up front that I don't give grades for effort, per se. Hard work, diligence, and motivation are indispensable for good learning. But they don't, in themselves, *constitute* good learning or high achievement. And those are the things that are primarily reflected in my grading. Let me give a few examples that may illustrate what I mean. Not everyone, for instance, can play

in the (W)NBA when they grow up, regardless of whether they've wanted to be a professional basketball player all their life or not. Nor can everyone be a doctor, or a stand-up comic, or an opera singer. You must have aptitude AND attitude AND a whole lot of effort. Other things will have to be sacrificed along the way. Many of us decide that those sacrifices are not worth the goal. And that is OK; choosing not to devote yourself solely to Biochemistry this fall will not make you a failure in my eyes, and it shouldn't make you view yourself as a failure either.

I view an "F" as a strong indication that the level of preparation and/or commitment brought to the class by a student are incompatible with the goals of this course. I hope not to give your class any "F"s.

I view a "D" as an indication that a student is passing the class, but performing well below my standards and failing to achieve a substantial portion of the course goals. Usually, this means that the student is performing well below her or his true abilities. I hope not to give your class any "D"s.

I view a "C" as notice that a student is doing only a fair job. Frequently, this means that the student is present but not fully motivated or engaged. A "C" student is doing adequately but probably came to the class with insufficient preparation and/or has not committed the personal resources to learn most effectively.

I view a "B" as a very respectable grade. The student to whom I give a "B" may be trying very hard but still struggling with mastery of the material, or may be working less hard and stopping short of achieving excellence.

I view a "B+" as an indication that a student is doing a truly good job. This grade indicates to me that the student is expending significant care and effort to ensure that s/he is learning the material.

I view an "A-" as a very good grade. I do not give this grade lightly or without evidence that a student is nearly approaching mastery of the ideas and techniques of this class; this grade indicates that a student has met my expectations for the course.

I view an "A" as an indication of true excellence. In order to achieve an "A" in my course, a student must demonstrate to me that s/he has not only committed the necessary resources to master the material, but also that s/he is *aggressively engaging the questions that we explore*. This is a grade to be proud of, a grade to be earned by serious work and mental sharpness.

The single biggest difference that I have found between "A" and "B" students over the years is that the former group takes a much more active role in the course, that they demonstrate a personal commitment to excellence in learning.

## LAB SCHEDULE

Date	Lab activities	Assignment due this week
<b><u>Experiment #1</u></b>		
Sep. 2	“The Big Picture”, make buffers	
Sep. 09	protein standard curve pour gel filtration column	<i>Fundamentals Test will be given at the start of lab today</i>
Sep. 16	lysate prep, cation exchange protein assay, abs @410	<i>std curve, sample calculations (10 pts)</i>
Sep. 23	dialysis, purification table clinic	<i>Ten Questions (self-graded) lab notebook interviews</i>
Sep. 30	sample concentration, std. gel filtration column	<i>dry run purification table (5 pts)</i>
Oct. 07	gel filtration of sample, plan SDS-PAGE samples	<i>graph of gel filtration data (ungraded)</i>
Oct. 14	SDS-PAGE	<i>purification table (10 pts; due Friday)</i>
Oct. 21	<b><i>Lab Test (50 pts)</i></b>	
Oct. 28	Back to the Big Picture: what does it mean?	<i>gel-filtration and SDS-PAGE analysis (25 pts; due Friday) Hand in lab notes from first experiment (10 pts; due Friday)</i>
<b><u>Experiment #2</u></b>		
Nov. 04	Big Picture #2 pH profile	
Nov. 11	K <sub>M</sub> and inhibitors	<i>graph of pH/activity profile (10 pts)</i>
Nov. 18	independent projects	<i>L-B analysis on inhibitors (5 pts)</i>
Dec. 02	independent projects	<i>by Friday, hand in poster abstract and data tables for ungraded feedback</i>
Dec. 09	check out	<i>last chance to hand in lab report revisions for regrading</i>
Dec. 10, 6 PM	Combined class poster session	<i>Poster Presentation (50 pts) hand in lab notes for Experiment #2 (20 pts)</i>



## OVERALL COURSE CALENDAR

Topical Outline	When we'll be there	Recommended chapters in B/T/S 6e*	Important dates and Assignments
<b>I. The Basics</b>			
A. Introduction to the context of biochemistry	week 1	1	
B. Introduction to the molecules	week 2	(4,11,12) 2.1, 2.2	
C. Introduction to protein analysis			<i>test #1, 21 Sep</i>
i. proteins in their native habitat			
ii. protein architecture	week 3	2.3-2.5	
iii. how proteins interact with themselves and other molecules	week 4	2.6	
iv. protein purification and characterization	weeks 5/6	3	
v. testing hypotheses in protein biochemistry via the central dogma of molecular biology	week 6	5,6	<i>test #2, 12 Oct</i>
<b>II. Towards a predictive understanding of protein architecture, function, and regulation</b>			
A. Case studies in protein S/F			
i. hemoglobin	weeks 7/8	7	<i>lab test, 21 Oct</i>
ii. receptor proteins	week 9		
B. Thermodynamic and kinetic tools in protein analysis	weeks 9,10	8, 9	<i>test #3, 6 Nov</i>
C. Control of protein function by modification of protein structure			
i. allosteric inhibition/activation	week 11	9, 10.1, 10.2	
ii. selective proteolysis	week 11	10.4	
iii. covalent modification	weeks 11/12	10.3, 21.2-3	
iv. further case studies in enzyme action and control	week 12		<i>test #4, 20 Nov</i>
<b>III. Proteins in the wild:</b>			
Function and regulation of proteins in complex biological systems	weeks 13/14	to be determined with class input	<i>course evals 4 Dec</i> <i>lab poster presentations, 10 Dec, 6 p.m.</i>  <i>FINAL EXAM</i> <i>In-class portion Monday 21 Dec, 9 a.m.</i>  <i>take-home portion due 18 Dec at 9 am</i>

\* Specific sections of chapters will be recommended in class, depending on the pace of our coverage and the abilities and interests of the members of the class.

**"Base Topics" for Biochemistry 302:**  
**concepts and techniques that we are *assuming* you are conversant with as you enter this class**

General and organic chemical principles

- **fluency** with metric units of measurement and prefixes (pico-, nano-, micro-, milli-, centi-, kilo-)
- **fluency** with scientific notation
- stoichiometry (gram  $\leftrightarrow$  mole relationships)
- solution properties and solution stoichiometry (concentration, dilution, ionic strength)
- periodic trends in bonding and reactivity of non-metals
- types and magnitudes of intermolecular interactions/forces
- the structure of small molecules and the impact of structure on reactivity, intermolecular forces, etc.
- dynamic chemical equilibrium (Le Chatelier's principle,  $K_{eq}$ )
- basic aqueous equilibrium calculations and pH ( $K_a$ ,  $K_b$ ,  $K_w$ , the Henderson-Hasselbalch equation, buffers)
- the interaction of light with matter (i.e., spectroscopy; especially Beer's Law and standard curves)
- kinetics (reaction order, Arrhenius equation, catalysis, collision geometry)
- thermodynamics (reaction coordinate diagrams; S, H, G; relationship of kinetic properties to thermodynamic properties; relationship of thermodynamic properties to equilibrium constant and behavior)
- structure and reactivity of common organic functional groups
- organic reaction mechanisms and "electron pushing"
- identification of nucleophiles/electrophiles and rationalization of their chemical behavior
- stereochemistry

Molecular and cellular biology

- cellular organization of prokaryotes and eukaryotes
- intracellular transport
- cellular information organization (the "central dogma": DNA  $\rightarrow$  RNA  $\rightarrow$  protein; basic molecular players in these processes)
- basic elements of gene structure and mRNA structure
- basic aspects of molecular cloning
- general understanding of proteins (what are enzymes?, the link between structure and function)

Lab Techniques/Skills

- pH meter use
- stir/heat plate use
- balance use (top-load and analytical and *when to use each*)
- column chromatography -- basic principles
- electrophoresis (protein and DNA)
- micropipet use (if you have any questions about proper use of micropipetters, *please* ask in lab!)
- **basic solution preparation** (e.g., how would you make 1.00 L of a 0.3 M solution of NaCl?)
- spectrophotometer principles and use
- **dealing with uncertainty in measurements (averages, standard deviations, confidence intervals for collections of numbers *and for lines*)**

Computer/Technology Skills

- basic use of Microsoft Word, Excel (including linear regression), and PowerPoint
- Web searching of data bases
- Web/online searching of literature sources
- E-mail use
- Network file sharing and information exchange

In addition, you will be expected to gain full familiarity (immediate recall) with the structures, names, one-letter-codes, and physicochemical properties of the 20 amino acids most commonly found in proteins. This information can be found in detail in chapter 2 of your text; it is summarized on the next page. This is an ideal project to begin on your first weekend of the semester.

Name	one-letter ID	notable physicochemical properties
All of them	X	$pK_A$ of the amino group is about 8 (an oversimplification, but...) $pK_A$ of the carboxyl group is about 3
Alanine	A, as in <b>A</b> lanine	small; in many ways this is the generic amino acid is often interchangeable, biochemically and evolutionarily, with G, S, or T
Glycine	G, as in <b>G</b> lycine	very small; can fit in tight spots is often interchangeable, biochemically and evolutionarily, with A (and less often w/ S, T)
Proline	P, as in <b>P</b> roline	unusual and constrained cyclic backbone geometry
Serine	S, as in <b>S</b> erine	small and modestly hydrophilic -OH group is a target for some covalent modifications as we'll discuss later is often interchangeable, biochemically and evolutionarily, with A and T
Threonine	T, as in <b>T</b> hreonine	-OH group is a target for some covalent modifications is often interchangeable, biochemically and evolutionarily, with A and S
Cysteine	C, as in <b>C</b> ysteine	contains sulfur can form a (covalent) disulfide bond with another C is sometimes interchangeable, biochemically and evolutionarily, with S; less often with A, T
Valine	V, as in <b>V</b> aline	bulky and hydrophobic is often interchangeable, biochemically and evolutionarily, w/ I, L, M; less often w/ F, Y (&W)
Isoleucine	I, as in <b>I</b> soleucine	bulky and hydrophobic is often interchangeable, biochemically and evolutionarily, w/ V, L, M; less often w/ F, Y (&W)
Leucine	L, as in <b>L</b> eucine	bulky and hydrophobic is often interchangeable, biochemically and evolutionarily, w/ V, I, M; less often w/ F, Y (&W)
Methionine	M, as in <b>M</b> ethionine	contains sulfur bulky and hydrophobic is often interchangeable, biochemically and evolutionarily, w/ V, I, L; less often w/ F, Y (&W)
Phenylalanine	F, as in <b>F</b> enylalanine	bulky and hydrophobic weakly absorbs UV light around 280 nm is often interchangeable, biochemically and evolutionarily, with Y; less often with I, V, L, M, W
Tyrosine	Y, as in <b>tY</b> rosine	bulky and hydrophobic absorbs UV light around 280 nm -OH group is a target for some covalent modifications is often interchangeable, biochemically and evolutionarily, with F; less often with I, V, L, M, W
Tryptophan	W, as in <b>tW</b> yptophan (just say it with a lisp!)	bulky and hydrophobic absorbs UV light around 280 nm occurs only rarely, but is generally interchangeable, biochemically and evolutionarily, with F and Y
Asparagine	N, as in asparagi <b>N</b> e	hydrophilic releases ammonia by amide hydrolysis when treated with strong acid is often interchangeable, biochemically and evolutionarily, w/ Q; sometimes with D, E
Glutamine	Q, as in <b>Q</b> tamine	hydrophilic releases ammonia by amide hydrolysis when treated with strong acid is often interchangeable, biochemically and evolutionarily, w/ N; sometimes with D, E
Aspartic acid	D, as in aspar <b>D</b> ic acid	acidic $pK_A$ , about 4 negative charge at pH above $pK_A$ is often interchangeable, biochemically and evolutionarily, w/ E; sometimes with N, Q
Glutamic acid	E, as in glu <b>E</b> -tamic acid	acidic $pK_A$ , about 4 negative charge at pH above $pK_A$ is often interchangeable, biochemically and evolutionarily, w/ D; sometimes with N, Q
Histidine	H, as in <b>H</b> istidine	acidic $pK_A$ , about 6 neutral charge at pH above $pK_A$ , positive charge at pH < $pK_A$
Lysine	K, as in the letter before L for lysine	basic $pK_A$ , about 11 positive charge at pH < $pK_A$ is often interchangeable, biochemically and evolutionarily, w/ R
Arginine	R, as in <b>R</b> ginine	basic $pK_A$ , about 12.5 positive charge at pH < $pK_A$ is often interchangeable, biochemically and evolutionarily, w/ K