Lecture Notes: 17-20 Feb, 2012

Histones

5 primary types of histones in eukaryotes

- H1
- H2A
- H2B
- H3 (highly conserved)
- H4 (highly conserved)
- histone tails are targets for many modifications that alter histone structure

Chromatin folding

- 1. string of nucleosomes (nucleosome filament)
- 2. filament is further folded into 30-nm fiber
- 3. fiber forms loops, centered to the central matrix of chromosomes

Effects of histones on transcription

- core histones repress transcription
- H1 causes further repression
- repression counteracted by transcription factors
- TFs can recruit transferases to modify histones
- nucleosome positioning is important for gene expression

Chromatin modification

We will focus on specific modifications to the tails of H3/H4 histones

- acK (lysine acetylation)
- meR (arginine methylation)
- meK (lysine methylation)
- PS (serine phosphorylation)
- UK (lysine ubiquitination)

Histone code

- one histone modification can influence other
- histone code: a combination of histone modifications that determine gene activities by helping to form euchromatin (active) or heterochromatin (inactive) states
- can be translated by bromodomain
 - interacts with acetylated lysine
 - BRG1 (SWI/SNF):H4 K8
 - TAF250 (TFIID): H3 K9K14
 - bromo-HATs (GCN5, PCAF, TAF250)
- can be translated by chromodomain
 - targets methylation marks
 - HP1 (methylated K9), repression
 - SUV39 HMT (interacts with HP1)
 - chromo-HATs (ESA1), activation
- Other domain
 - PHD (TAF3 or TAF110),
 - Recognizes H3K4me3 activation

Chromatin remodeling

- disrupt the core histones in nucleosomes (ATP-dependent)
- $\bullet\,$ may also move nucleosomes
- create nucleosome-free enhancers with histone acetylases and other proteins

Need to memorize

