

Figure 3. Features of the Agouti viable yellow allele. (A) The Agouti viable yellow allele (not to scale) has an intracisternal A particle (IAP) insertion (striped box) in pseudoexon 1a (gray box), ~100 kb upstream of the Agouti coding exons (black). IAP long terminal repeats (LTRs) are shown as arrowheads, and transcriptional start sites shown as arrows. (B) An agouti wild-type mouse with an  $A^+$  allele has a brown coat color phenotype with a twotoned hair shaft, in which the base appears black in color, and closer to the tip shows a yellow color (see cartoon in C). This phenotype occurs because the Agouti gene, producing a yellow fur color, is only transiently expressed in the hair follicle microenvironment. The  $A^{vy}$  allele, if expressed ubiquitously from the IAP promoter, produces a phenotype with an entirely yellow hair shaft. There is a spectrum of coat color phenotypes observed, however, because of the  $A^{yy}$  allele ranging from completely yellow when the IAP LTR is active in all cells, to mottled due to patches of active and inactive cells, and finally animals with an agouti colored coat, called pseudoagouti, which are indistinguishable from wild-type agouti animals because of silenced IAP LTR in all cells. (Reproduced from Morgan et al. 1999.) (C) Mice with the Agouti viable yellow allele in a euchromatic state in all cells appear yellow, whereas those with the allele in a heterochromatic state in all cells appear agout in color, termed pseudoagouti. Mice that are mosaic for the heterochromatic and euchromatic allele appear mottled. (D) Summary of the epigenetic marks found at an active or inactive Agouti viable yellow allele. The inactive allele is hypermethylated and enriched for H4K20 trimethylation. The active allele is hypomethylated and enriched for acetylated residues of histone H3 and H4 tails.