Fig. 6. Schematic illustration of CCR5 haplotypes. (a) A CCR5 open reading frame (ORF) and polymorphisms in the neighboring DNA sequence that are tightly linked to the CCR5 ORF define different CCR5 haplotypes. Polymorphisms tightly linked to the CCR5 ORF have been found within CCR2 and in the cis-regulatory region of CCR5. In every cell there is a pair of CCR5 haplotypes that differ from each other (i.e., heterozygous) or are identical to each other (homozygous). Different combinations of CCR5 haplotype pairs (e.g., HHG\*2/HHE) may be associated with varied susceptibility to HIV-1 infection and/or altered rates of disease progression in HIV-1-infected individuals. Wt, indicates wild type. To maintain uniformity in the numbering systems, it was proposed at the CCR5--AIDS symposium (held at the NCI-FCRDC, Frederick, MD, April 20, 1999), to designate the first nucleotide of the CCR5 translational start site as +1 and the nucleotide immediately upstream as -1 (3, 4). For reference, we have provided the two commonly used numbering systems along with the new numbering system. (b) The cohorts were genotyped for polymorphisms indicated. Genotyping methods and haplotype classification were as described previously (1, 4). In this classification system, CCR5 haplotypes are grouped into one of seven human haplogroups (HH) -A, -B, -C, -D, -E, -F (F\*1 and F\*2), -G (G\*1 and G\*2). The genotypic characteristics of haplotypes within each haplogroup at the polymorphic positions CCR2-V64I, and CCR5 A-2733G, G-2554T, G-2459A, T-2135C, C-2132T, A-2086G, C-1835T, and  $\pm \Delta 32$  is as shown. Differences from the ancestral CCR5 haplogroup (HHA) are in bold. In this paper, homozygosity refers to two haplotypes of the same haplogroup and heterozygosity indicates two haplotypes from two different haplogroups. [Adapted from Mangano et al. (5).]