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Focusing the immune response on the V3 loop of HIV-I gp I 20 induces cross-clade neutralizing

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Experiments were performed to determine if focusing the immune response on a single neutralizing epitope of gp120 would result in cross-clade neutralizing antibodies (Nabs). Thus, rabbits were immunized with 2-3 priming doses of gp120 DNA (from clades A, B, or C) and with two boosters of fusion proteins carrying the consensus V3 epitope from either clades A, B or C. Immune sera neutralized primary isolates from strains and clades heterologous to those of the prime and boost; ND50 titers exceeded 1:20. ND50 titers >1:800 were achieved against SF162 pseudoviruses carrying the consensus V3 loops of clades A1, AG, B, AE, and F. The neutralizing activity was primarily due to V3-specific Abs as shown by peptide absorption experiments. The neutralizing activity could be significantly broadened as a function of the clade used to construct the gp120 DNA prime and the V3-FP boost.