Translocations at replication hubs The immunoglobulin heavy chain (IGH) enhancer on chromosome 14 and the MYC

The MYC and IGH loci are

in topological associating domains (TADs) that are close to each other in the nucleus.

locus on chromosome 8 form translocations that are associated with lymphoma. These loci also contain DNA replication origins and the IGH locus undergoes

recombination in B cells during antibody maturation. These processes and the proximity of the two loci increases the opportunity for translocation.

Myc TAD

Phase MYC and IGH separated hub chromatin loops **IGH** Replication loops contain replication Мус enhancer origins. The origin linked concentration of origin replication and recombination Replication and repair proteins repair factors creates a phase separated hub

Replication and recombination creates opportunities for DNA double strand breaks to arise in MYC and IGH, which can be joined to form **oncogenic IGH-MYC translocation**.