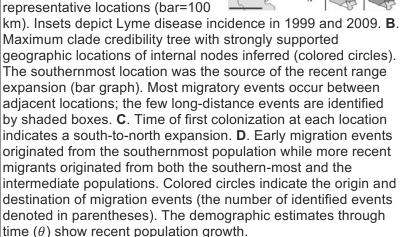
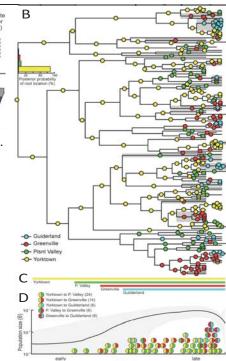
Fig 3. Nearly all recent migration occurred in a south-to-north direction among adjacent locations resulting in a gradual *I. scapularis* range expansion A. Arrows indicate strongly supported migration paths among 4





than *Taq* polymerase)^{77–80}. Uniform priming with random hexamers results in uniform amplification of all DNA within a sample. In contrast, SWGA takes advantage of the fact that every species has different mutational biases that create over- and under-represented sequence motifs within their genomes⁴⁹. Motifs that are over-represented in the target microbial genome, but under-represented in the tick genome, are used to create primer sets (2-20 primers each 6-12bp in length) for \$\phi29\$ reactions that amplify only the target microbial genome⁵⁰. We have used these technologies to develop successful SWGA protocols to sequence a variety of microbial species without prior culturing including: *Wolbachia pipientis* (fruit flies)⁴⁹, multiple *Plasmodium* species (primate blood)⁵¹, *Trypanosoma cruzi* (triatomine bug), *Mycobacterium tuberculosis* (human) and *B. burgdorferi* (ticks). Here we describe three representative cases. *W. pipientis* - a maternally-inherited bacterium in fruit flies - accounts for ~2% of MiSeq reads from genomic extracts of infected fruit flies⁴⁹. Thus, 98% of reads are wasted on non-target DNA if *W. pipientis* is sequenced directly from fly genomic extracts. However, sequencing *W. pipientis* from the same fly extract after SWG amplification results in ~70% of sequencing reads mapping to *W*.

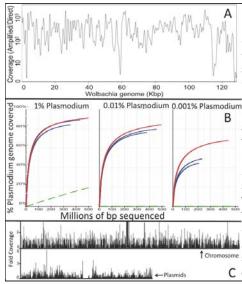


Fig 4. Selective whole genome amplification dramatically improves sequencing coverage of the target genome. A. Sequencing coverage (normalized by sequencing effort) was ~100X greater in **SWG amplified** samples than directly sequenced samples across the Wolbachia genome. Coverage of amplified samples at each bp was divided by coverage from directly sequenced samples. Only 4 locations in the Wolbachia genome had equivalent coverage in the amplified and non-amplified samples (no reads mapped to those areas for either sample). B. SWGA performed on primate genomic DNA containing known quantities of Plasmodium DNA (1% to 0.001%). The sequencing effort is shown in relation to the percent of the *Plasmodium* genome with sequencing coverage. Red lines show the combination of two independent amplifications (blue lines). Coverage without SWGA is shown in green. **C**. Although no reads map to *B*. burgdorferi from tick extracts prior to SWGA, post-SWGA †chromosome samples with equivalent sequencing effort have relatively C even coverage.