WHAT IS CLAIMED IS:

1. A mixed carrier, multivalent pneumococcal conjugate composition, comprising 21 different pneumococcal capsular polysaccharide-protein conjugates, wherein each pneumococcal capsular polysaccharide-protein conjugate comprises a protein carrier conjugated to a capsular polysaccharide from a different serotype of *Streptococcus pneumoniae*, wherein the *Streptococcus pneumoniae* serotypes are selected from 1, 3, 4, 5, 6A, 6B, 7F, 8, 9N, 9V, 10A, 11A, 12F, 14, 15B, 18C, 19A, 19F, 22F, 23F, and 33F,

wherein the protein carrier is $CRM_{197}$ or tetanus toxoid, and

wherein four of the capsular polysaccharides are conjugated to tetanus toxoid and the remaining capsular polysaccharides are conjugated to $CRM_{197}$, wherein the four capsular polysaccharides that are conjugated to tetanus toxoid are serotypes 15B, 22F and two serotypes selected from the group consisting of serotypes 1, 3, and 5.

2. The mixed carrier, multivalent pneumococcal conjugate composition of claim 1, wherein the capsular polysaccharides from serotypes 1, 5, 15B and 22F are conjugated to the tetanus toxoid, and the capsular polysaccharides from serotypes 3, 4, 6A, 6B, 7F, 8, 9N, 9V, 10A, 11A, 12F, 14, 18C, 19A, 19F, 23F, and 33F are conjugated to $CRM_{197}$.

3. The mixed carrier, multivalent pneumococcal conjugate composition of claim 1, wherein the capsular polysaccharides from serotypes 1, 3, 15B and 22F are conjugated to the tetanus toxoid, and the capsular polysaccharides from serotypes 4, 5, 6A, 6B, 7F, 8, 9N, 9V, 10A, 11A, 12F, 14, 18C, 19A, 19F, 23F, and 33F are conjugated to $CRM_{197}$.

4. The mixed carrier, multivalent pneumococcal conjugate composition of claim 1, wherein the capsular polysaccharides from serotypes 3, 5, 15B and 22F are conjugated to the tetanus toxoid, and the capsular polysaccharides from serotypes 1, 4, 6A, 6B, 7F, 8, 9N, 9V, 10A, 11A, 12F, 14, 18C, 19A, 19F, 23F, and 33F are conjugated to $CRM_{197}$.

5. The mixed carrier, multivalent pneumococcal conjugate composition of any one of the preceding claims, further comprising an adjuvant.

6. The mixed carrier, multivalent pneumococcal conjugate composition of claim 5, wherein the adjuvant is an aluminum-based adjuvant.

7. The mixed carrier, multivalent pneumococcal conjugate composition of claim 6, wherein the adjuvant is selected from the group consisting of aluminum phosphate, aluminum sulfate, and aluminum hydroxide.

8. The mixed carrier, multivalent pneumococcal conjugate composition of claim 7, wherein the adjuvant is aluminum phosphate.

9. The mixed carrier, multivalent pneumococcal conjugate composition of any one of the preceding claims, wherein:

the capsular polysaccharide from serotype 9N is conjugated to $CRM_{197}$ in a state where the capsular polysaccharide from serotype 9N is activated to have a degree of oxidation of 2-19 or 5-10 and a molecular weight of 200-700 kDa;

the conjugate formed between the capsular polysaccharide from serotype 9N and $CRM_{197}$ has a molecular weight of 500-4,000 kDa;

a ratio of the capsular polysaccharide from serotype 9N to $CRM_{197}$ in the conjugate formed between the capsular polysaccharide from serotype 9N and $CRM_{197}$ is 0.5-2.5 (w/w); and/or

15-60 % of the conjugate formed between the capsular polysaccharide from serotype 9N and $CRM_{197}$ has a $K_d$ of 0.3 or below in a CL-4B column.

10. The use of the mixed carrier, multivalent pneumococcal conjugate composition of any of the preceding claims for prophylaxis against *Streptococcus pneumoniae* infection or disease in a subject.

11. A vaccine comprising the mixed carrier, multivalent pneumococcal conjugate composition of any one of claims 1-9 and a pharmaceutically acceptable excipient.

12. A method for prophylaxis of *Streptococcus pneumoniae* infection or disease in a subject, the method comprising administering a prophylactically effective amount of the mixed carrier, multivalent pneumococcal conjugate composition of any one of claims 1-9 or the vaccine of claim 11 to the subject.

13. The method of claim 12, wherein the subject is a human who is at least 50 years old and the disease is pneumonia or invasive pneumococcal disease (IPD).

14. The method of claim 12, wherein the subject is a human who is at least 6 weeks old and the disease is pneumonia, invasive pneumococcal disease (IPD), or acute otitis media (AOM).

15. The method of claim 14, wherein the subject is 6 weeks to 5 years of age, 2 to 15 months of age, or 6 to 17 years of age.

16. The use of claim 10 or the method of any one of claims 12-15, wherein the subject is a human.

17. The method of any one of claims 12-16, wherein the mixed carrier, multivalent pneumococcal conjugate composition or the vaccine is administered by intramuscular injection.

18. The method of any one of claims 12-17, wherein the mixed carrier, multivalent pneumococcal conjugate composition or the vaccine is administered as part of an immunization series.