

Revised Structure of the Capsular Polysaccharide from Streptococcus Pneumoniae Serotype 7C

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The polysaccharide capsule of Streptococcus *pneumoniae* is recognized as the most important virulence factor of pneumococci and the capsular polysaccharides are widely used as the antigens in pneumococcal vaccines. The structure of the polysaccharide from Streptococcus *pneumoniae* serotype 7C was published in 2018 by Christian Kjeldsen et al based on NMR spectroscopy.

A discrepancy in the composition of this polysaccharide was found from both composition and linkage analysis. No galactosamine was detected by HPAEC-PAD from the hydrolysate of 7C polysaccharide, and an extra glucosamine was found instead. The extra glucosamine residue was determined to be in a 3,4-linkage by GC/MS analysis of the partially methylated alditol acetates. The presence of 3, 4-GlcpNAc was further confirmed by NMR analysis. While the 1 H NMR and 2D NMR (COSY, HSQC and HMBC) spectra are comparable with the data published, the large 3 J coupling constant ($^{\sim}9.6$ ppm) between both H-4 and H-3 (J_{3,4}) and H-4 and H-5 (J_{4,5}) of the 3,4-linked HexpNAc from a selective NOE experiment suggested that the H-4 is in an axial position, and the 3,4-HexpNAc should be 3, 4-GlcpNAc instead of 3, 4-GlcpNAc in the published structure. The same conclusion is supported by the strong NOE between the H-4 of 3, 4-GlcpNAc and the H-1 of the glucose link to 3, 4-GlcpNAc through O-4.

In conclusion, the revised structure of the polysaccharide from Streptococcus *pneumoniae* serotype 7C is:

 $[\rightarrow 6)-\alpha\text{-D-Glc}\rho\text{NAc-}(1\rightarrow 2)-\alpha\text{-L-Rha}p\text{-}(1\rightarrow 2)-\beta\text{-L-Rha}p\text{-}(1\rightarrow 4)-\beta\text{-D-Glc}\rho\text{-}(1\rightarrow 4)-\alpha\text{-D-Glc}\rho\text{NAc-}(1\rightarrow O\text{-P}\rightarrow)] \cap \\ \uparrow (1-3) \\ \beta\text{-D-Rib}f\text{-}(1\rightarrow 4)-\alpha\text{-L-Rha}p$

Revised structure of SP 7C

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GSK Vaccines

Bibliographic references:

Christian Kjeldsen, Sofie Slott, Pernille L. Elverdal, Carmen L. Sheppard, Georgia, Kapatai, Norman K. Fry, Ian C. Skovsted, Jens Ø. Duus. Discovery and description of a new serogroup 7 Streptococcus pneumoniae serotype, 7D, and structural analysis of 7C and 7D. Carbonydr. Res. 2018, 463, 24-31. DOI: 10.1016/j.carres.2018.04.011