

Streptococcus

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Streptococcus is a genus of coccus (spherical) Gram-positive bacteria belonging to the phylum Firmicutes^[3] and the Lactobacillales (lactic acid bacteria) order. Cellular division occurs along a single axis in these bacteria, and thus they grow in chains or pairs, hence the name—from Greek στρεπτός *streptos*, meaning easily bent or twisted, like a chain (twisted chain).

Contrast this with *Staphylococci*, which divide along multiple axes and generate grape-like clusters of cells. Most *Streptococci* are oxidase- and catalase-negative, and many are facultative anaerobes.

In 1984, many organisms formerly considered *Streptococcus* were separated out into the genera *Enterococcus* and *Lactococcus*.^[4] Currently, over 50 species are recognised in this genus.

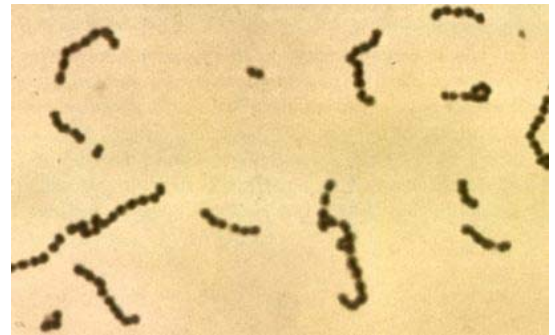
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Pathogenesis and classification

In addition to streptococcal pharyngitis (strep throat), certain *Streptococcus* species are responsible for many cases of pink eye,^[5] meningitis, bacterial pneumonia, endocarditis, erysipelas, and necrotizing fasciitis (the 'flesh-eating' bacterial infections). However, many streptococcal species are not

Streptococcus



Scientific classification

Kingdom:	Bacteria
Phylum:	Firmicutes
Class:	Bacilli ^[1]
Order:	Lactobacillales
Family:	Streptococcaceae
Genus:	<i>Streptococcus</i>
	Rosenbach, 1884

Species^[2]

- *Streptococcus agalactiae*
- *Streptococcus anginosus*
- *Streptococcus bovis*
- *Streptococcus canis*
- *Streptococcus constellatus*
- *Streptococcus dysgalactiae*
- *Streptococcus equinus*
- *Streptococcus iniae*
- *Streptococcus intermedius*
- *Streptococcus milleri*
- *Streptococcus mitis*
- *Streptococcus mutans*
- *Streptococcus oralis*
- *Streptococcus parasanguinis*
- *Streptococcus peroris*
- *Streptococcus pneumoniae*
- *Streptococcus pseudopneumoniae*
- *Streptococcus pyogenes*
- *Streptococcus rattii*

pathogenic, and form part of the commensal human microbiota of the mouth, skin, intestine, and upper respiratory tract. Furthermore, streptococci are a necessary ingredient in producing Emmentaler ("Swiss") cheese.

Species of *Streptococcus* are classified based on their hemolytic properties.^[6] Alpha-hemolytic species cause oxidation of iron in hemoglobin molecules within red blood cells, giving it a greenish color on blood agar. Beta-hemolytic species cause complete rupture of red blood cells. On blood agar, this appears as wide areas clear of blood cells surrounding bacterial colonies. Gamma-hemolytic species cause no hemolysis.

Beta-hemolytic streptococci are further classified by Lancefield grouping, a serotype classification (that is, describing specific carbohydrates present on the bacterial cell wall).^[7] The 20 described serotypes are named Lancefield groups A to V (excluding I and J).

In the medical setting, the most important groups are the alpha-hemolytic streptococci *S. pneumoniae* and *Streptococcus viridans* group, and the beta-hemolytic streptococci of Lancefield groups A and B (also known as "group A strep" and "group B strep").

Alpha-hemolytic

When alpha hemolysis (α -hemolysis) is present, the agar under the colony is dark and greenish. *Streptococcus pneumoniae* and a group of oral streptococci (*Streptococcus viridans* or viridans streptococci) display alpha hemolysis. This is sometimes called green hemolysis because of the color change in the agar. Other synonymous terms are incomplete hemolysis and partial hemolysis. Alpha hemolysis is caused by hydrogen peroxide produced by the bacterium, oxidizing hemoglobin to green methemoglobin.

Pneumococci

- *S. pneumoniae* (sometimes called pneumococcus), is a leading cause of bacterial pneumonia and occasional etiology of otitis media, sinusitis, meningitis, and peritonitis. Inflammation is thought to be the major cause of how pneumococci cause disease, hence the tendency of diagnoses associated with them to involve inflammation.

The viridans group: alpha-hemolytic

- The viridans streptococci are a large group of commensal bacteria, that are either α -hemolytic, producing a green coloration on blood agar plates (hence the name "viridans", from Latin *vīridis*, green), or nonhemolytic. They possess no Lancefield antigens.^[3]

Table: Medically relevant streptococci^[6]

- *Streptococcus salivarius*
- *Streptococcus tigurinus*
- *Streptococcus thermophilus*
- *Streptococcus sanguinis*
- *Streptococcus sobrinus*
- *Streptococcus suis*
- *Streptococcus uberis*
- *Streptococcus vestibularis*
- *Streptococcus viridans*
- *Streptococcus zooepidemicus*
- etc.

Species	Host	Disease
<i>S. pyogenes</i>	human	pharyngitis
<i>S. agalactiae</i>	human, cattle	neonatal meningitis and sepsis
<i>S. equisimilis</i>	human, animals	endocarditis, bacteremia, pneumonia, meningitis, respiratory infections
<i>S. bovis</i>	human, animals	biliary or urinary tract infections, endocarditis
<i>S. anginosus</i>	human, animals	subcutaneous/organ abscesses, meningitis, respiratory infections
<i>S. sanguinis</i>	human	endocarditis, dental caries
<i>S. suis</i>	swine	meningitis
<i>S. mitis</i>	human	endocarditis
<i>S. mutans</i>	human	dental caries
<i>S. pneumoniae</i>	human	pneumonia

Beta-hemolytic

Beta hemolysis (β -hemolysis), sometimes called complete hemolysis, is a complete lysis of red cells in the media around and under the colonies: the area appears lightened (yellow) and transparent. Streptolysin, an exotoxin, is the enzyme produced by the bacteria which causes the complete lysis of red blood cells. There are two types of streptolysin: Streptolysin O (SLO) and streptolysin S (SLS). Streptolysin O is an oxygen-sensitive cytotoxin, secreted by most Group A streptococcus (GAS), and interacts with cholesterol in the membrane of eukaryotic cells (mainly red and white blood cells, macrophages, and platelets), and usually results in β -hemolysis under the surface of blood agar. Streptolysin S is an oxygen-stable cytotoxin also produced by most GAS strains which results in clearing on the surface of blood agar. SLS affects immune cells, including polymorphonuclear leukocytes and lymphocytes, and is thought to prevent the host immune system from clearing infection. *Streptococcus pyogenes*, or Group A beta-hemolytic Strep (GAS), displays beta hemolysis.

Some weakly beta-hemolytic species cause intense beta hemolysis when grown together with a strain of *Staphylococcus*. This is called the CAMP test.[1] *Streptococcus agalactiae* displays this property. *Clostridium perfringens* can be identified presumptively with this test. *Listeria monocytogenes* is also positive on sheep's blood agar.

Group A

S. pyogenes, also known as group A *Streptococcus* (**GAS**), is the causative agent in a wide range of group A streptococcal infections. These infections may be noninvasive or invasive. The noninvasive infections tend to be more common and less severe. The most common of these infections include streptococcal pharyngitis (strep throat) and impetigo.[8] Scarlet fever is also a noninvasive infection, but has not been as common in recent years.

The invasive infections caused by group A β -hemolytic streptococci tend to be more severe and less common. This occurs when the bacterium is able to infect areas where it is not usually found, such as the blood and the organs.[9] The diseases that may be caused include streptococcal toxic shock syndrome, necrotizing fasciitis, pneumonia, and bacteremia.[8]

Additional complications may be caused by GAS, namely acute rheumatic fever and acute glomerulonephritis. Rheumatic fever, a disease that affects the joints, kidneys, and heart valves, is a



Alpha-hemolytic *S. viridans* (right) and beta-hemolytic *S. pyogenes* (left) streptococci growing on blood agar

consequence of untreated strep A infection caused not by the bacterium itself. Rheumatic fever is caused by the antibodies created by the immune system to fight off the infection cross-reacting with other proteins in the body. This "cross-reaction" causes the body to essentially attack itself and leads to the damage above. Globally, GAS has been estimated to cause more than 500,000 deaths every year, making it one of the world's leading pathogens.^[8] Group A *Streptococcus* infection is generally diagnosed with a rapid strep test or by culture.

Group B

S. agalactiae, or group B *Streptococcus*, **GBS**, causes pneumonia and meningitis in neonates and the elderly, with occasional systemic bacteremia. They can also colonize the intestines and the female reproductive tract, increasing the risk for premature rupture of membranes during pregnancy, and transmission of the organism to the infant. The American Congress of Obstetricians and Gynecologists (formerly the American College of Obstetricians and Gynecologists), American Academy of Pediatrics, and the Centers for Disease Control recommend all pregnant women between 35 and 37 weeks gestation to be tested for GBS. Women who test positive should be given prophylactic antibiotics during labor, which will usually prevent transmission to the infant.^[10]

The United Kingdom has chosen to adopt a risk factor-based protocol, rather than the culture-based protocol followed in the US. Current guidelines state that if one or more of the following risk factors are present, then women should be treated with intrapartum antibiotics:

- Preterm labour (<37 weeks)
- Prolonged rupture of membranes (>18 hours)
- Intrapartum fever (>38C)
- Prior GBS affected infant
- GBS bacteriuria during this pregnancy

This protocol results in treatment of 15–20% of pregnant women and prevention of 65–70% of cases of early onset GBS sepsis.^[11]

Group C

This group includes *S. equi*, which causes strangles in horses,^[12] and *S. zooepidemicus*—*S. equi* is a clonal descendent or biovar of the ancestral *S. zooepidemicus*—which causes infections in several species of mammals, including cattle and horses. *S. dysgalactiae* is also a member of group C, β-haemolytic streptococci that can cause pharyngitis and other pyogenic infections similar to group A streptococci.

Group D (enterococci)

Many former group D streptococci have been reclassified and placed in the genus *Enterococcus* (including *E. faecalis*, *E. faecium*, *E. durans*, and *E. avium*).^[13] For example, *Streptococcus faecalis* is now *Enterococcus faecalis*. *E. faecalis* is sometimes alpha hemolytic and *E. faecium* is sometimes beta hemolytic.^[14]

The remaining nonenterococcal group D strains include *Streptococcus bovis* and *Streptococcus equinus*.

Nonhemolytic streptococci rarely cause illness. However, weakly hemolytic group D beta-hemolytic streptococci and *Listeria monocytogenes* (which is actually a Gram-positive bacillus) should not be confused with nonhemolytic streptococci.

Group F streptococci

Group F streptococci were first described in 1934 by Long and Bliss amongst the "minute haemolytic

streptococci".^[15] They are also known as *Streptococcus anginosus* (according to the Lancefield classification system) or as members of the *S. milleri* group (according to the European system).

Group G streptococci

These streptococci are usually, but not exclusively, beta-hemolytic. *S. canis* is an example of a GGS which is typically found on animals, but can cause infection in humans.

Group H streptococci

Group H streptococci cause infections in medium-sized canines. Group H streptococci rarely cause illness unless a human has direct contact with the mouth of a canine. One of the most common ways this can be spread is human-to-canine, mouth-to-mouth contact. However, the canine may lick the human's hand and infection can be spread, as well.^[16]

Molecular taxonomy and phylogenetics

Streptococci have been divided into six groups on the basis of their 16S rDNA sequences: *S. anginosus*, *S. bovis*, *S. mitis*, *S. mutans*, *S. pyogenes* and *S. salivarius*.^[18] The 16S groups have been confirmed by whole genome sequencing (see figure). The important pathogens *S. pneumoniae* and *S. pyogenes* belong to the *S. mitis* and *S. pyogenes* groups, respectively, while the causative agent of dental caries, *Streptococcus mutans*, is basal to the *Streptococcus* group.

Genomics

The genomes of hundreds of species have been sequenced.^[20] Most *Streptococcus* genomes are 1.8 to 2.3 Mb in size and encode 1,700 to 2,300 proteins. Some important genomes are listed in the table.^[21] The four species shown in the table (*S. pyogenes*, *S. agalactiae*, *S. pneumoniae*, and *S. mutans*) have an average pairwise protein sequence identity of about 70%.^[21]

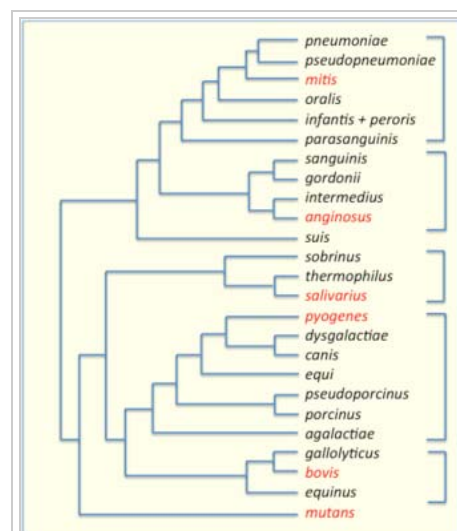
feature	<i>S. pyogenes</i>	<i>S. agalactiae</i>	<i>S. pneumoniae</i>	<i>S. mutans</i>
base pairs	1,852,442	2,211,488	2,160,837	2,030,921
ORFs	1792	2118	2236	1963
prophages	yes	no	no	no

Treatment

See also

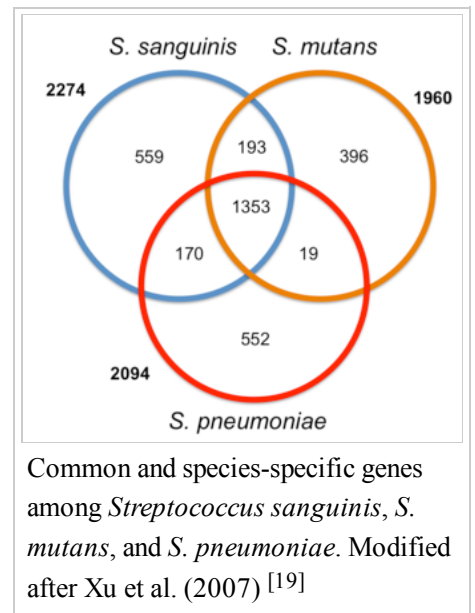
- Quellung reaction
- Streptococcal infection in poultry
- Streptokinase

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Phylogenetic tree of *Streptococcus* species, based on data from PATRIC.^[17] 16S groups are indicated by brackets and their key members are highlighted in red.

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External links

- *Streptococcus* (http://patricbrc.org/portal/portal/patric/Taxon?cType=taxon&cId=1301) genomes and related information at PATRIC (http://patricbrc.org/), a Bioinformatics Resource Center funded by NIAID (http://www.niaid.nih.gov/)
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- The UK (http://www.gbss.org.uk) Group B Strep Support charity

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