



dnasmile

Oral and systemic health

Welcome

Example

to your DNA Smile report

Date of birth:

Date reported: 18 Nov 2021

Sample number: 12345678

Referring practitioner:

DNA Smile is a genetic test designed to give you insight into the state of your dental and oral health and risk for developing periodontal disease and tooth caries.



Oral and dental health



Heart disease



Risk of cancer and chronic disease



Cognitive function



Type 2 diabetes

Genetics and personalised medicine

Genes are segments of DNA that contain the instructions your body needs to make each of the many thousands of proteins required for life. Each gene is comprised of thousands of combinations of “letters” (called bases) which make up your genetic code. The code gives the instructions to make the proteins required for proper development and function.

Genetic variations can affect the expression of a gene, thereby affecting metabolic processes that are important for maintaining cellular health and how we respond to environmental interventions such as diet, lifestyle, supplements, and medication.

Knowledge of these genetic variations offers unparalleled insight into your biological systems, allowing your healthcare practitioner to recommend precise interventions aimed at helping you reach your goals and achieve optimal health.



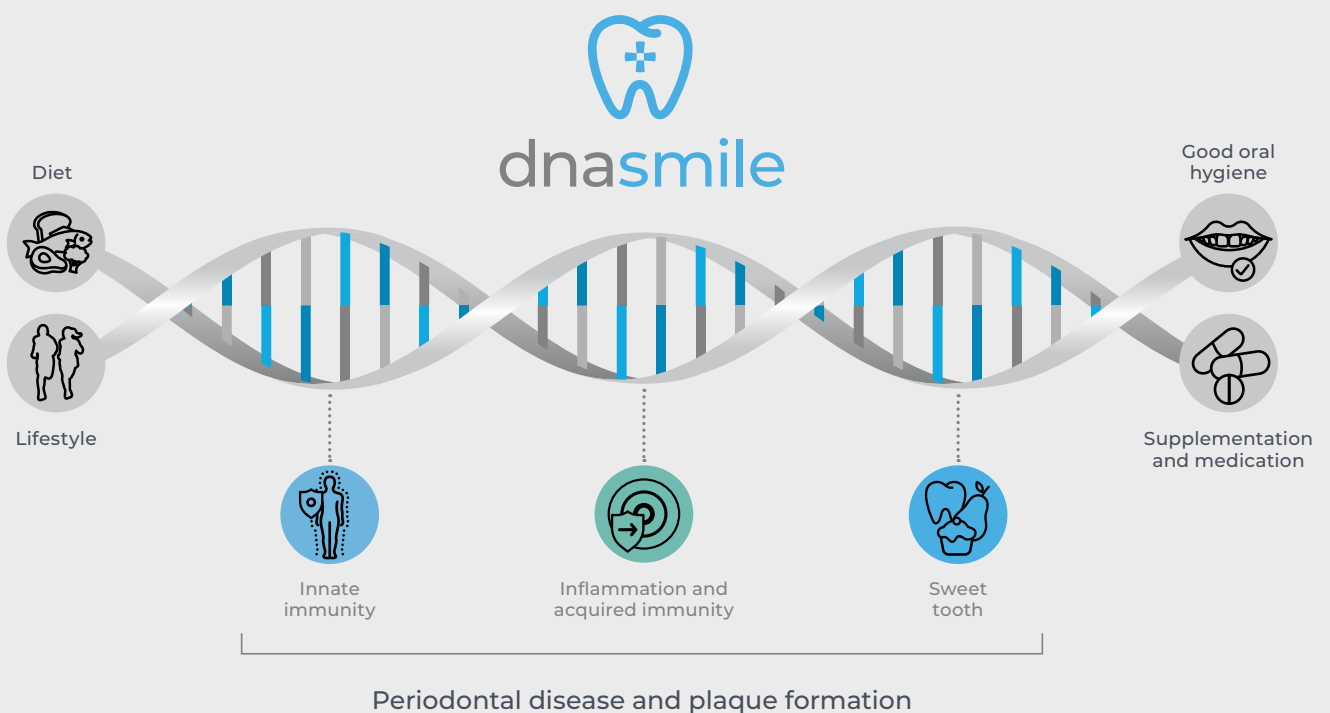
NORMAL GENE
Genotype resulting in baseline potential for disease risk



VARIANT GENE
Genotype resulting in increased potential for disease risk and need for personalised intervention

Personalised medicine and dental health

Your oral health is strongly related to risk for development of chronic diseases, from cognitive decline to type 2 diabetes and heart disease. Early, targeted interventions using a holistic, systems-biology approach is essential to improve overall health outcomes. This report provides you with actionable information to improve oral health and prevent disease.



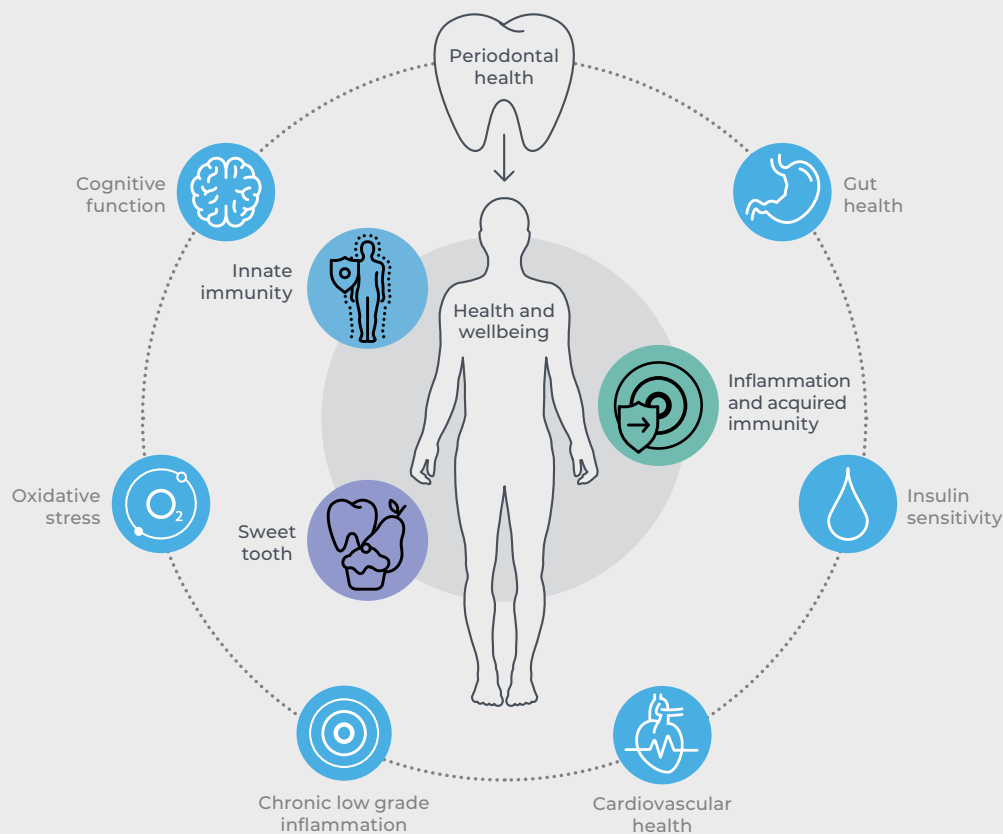
Understanding periodontal disease

Periodontal disease refers to both gingivitis (gum disease) and periodontitis. Periodontitis is a chronic inflammatory disease of the mouth that begins when bacteria, found in dental plaque, causes a major negative immune response, which leads to the damage of the connective tissue and the bones surrounding your teeth. If left untreated, the damage is irreversible and causes tooth loss.

Your risk for developing periodontal disease is based on a number of factors, including oral hygiene practices, dietary intake (a high sugar intake), the presence of other inflammatory illnesses, exposure to certain pathogenic bacteria and an altered oral and gut microbiome, and, importantly, genetics.

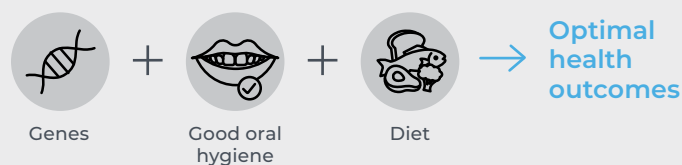
Besides the importance of tending to your oral health to prevent tooth loss, is the relevance that your oral health has on the rest of your body. Periodontal disease has been strongly linked to an increased risk for a number of chronic inflammatory diseases, including cognitive decline and late-onset Alzheimer's disease, as well as heart disease, type 2 diabetes and certain cancers (oesophageal and oral cancer). Therefore, by understanding your risk for periodontal disease and ensuring good oral health with personalised interventions, you are also able to control and decrease your risk for other inflammatory diseases.

DNA SMILE CAN HELP IDENTIFY HEALTH RISKS:






To improve your health outcomes and reduce the risk of disease:

understand your risk and ensure good oral health with early targeted interventions






Result summary

BIOLOGICAL AREA	PRIORITY
 Innate immunity	
 Inflammation and acquired immunity	
 Sweet tooth	

Genotype results

No Impact
 Low Impact
 Moderate Impact
 High Impact
 Beneficial Impact

BIOLOGICAL AREA	GENE NAME	GENE VARIATION	RESULT	GENE IMPACT
 Innate immunity	DEFB	3'UTR c*5G>A	AA	
		-44 C>G	GG	
	TLR4	896 A>G	GG	
		1196 C>T	TT	
 Inflammation and acquired immunity	IL-1A	-889 C>T	CC	
		4845 G>T	GG	
	IL1B	3954 C>T	CC	
		-511 T>C	AA	
	IL1RN	2018 T>C	TT	
	IL6	-174 C>G	GG	
	IL17A	-197 G>A	GA	
	MMP3	Lys45Glu (A>G)	GG	
TNFA	G>A	GG		
 Sweet tooth	SLC2A2	Thr110Ile (T>C)	TT	
	TAS1R2	Ile191Val (G>A)	AA	

Gene results per biological area with personalised recommendations



Innate immunity

The innate immune system comprises physical defensive barriers such as the skin; defensive mechanisms such as saliva; and the general immune response, which includes immune cells and proteins. The innate immune system is activated by the presence of antigens and their chemical properties and depends on pattern recognition receptors (PRRs) to detect conserved structures of pathogenic microorganisms (like bacteria, virus, fungus and protozoa), which are called pathogen-associated molecular patterns (PAMPs).



Innate immunity results

Genotype result table:

GENE NAME	GENE VARIATION	RESULT	GENE IMPACT
DEFB	3'UTR c*5G>A	AA	
	-44 C>G	GG	
TLR4	896 A>G	GG	
	1196 C>T	TT	

Gene and genotype specific information



DEFB

The DEFB1 gene encodes the beta-defensin 1 protein. The defensin family is a group of microbicidal and cytotoxic peptides that are essential in the innate immune response. Beta-defensin 1 is an antimicrobial peptide implicated in the resistance of epithelial surfaces to microbial colonization. The SNP has been found to influence susceptibility to and severity of periodontal diseases.

Result: AA

The AA genotype is associated with an increased risk for both chronic and aggressive periodontitis. Do not smoke and keep blood glucose levels well-controlled. Ensure good dental hygiene and consult with your dentist regarding risk management. Follow an anti-inflammatory diet, avoid refined carbohydrates and sugars.

Result: GG

The GG genotype is associated with normal protein function.



TLR4

The TLR4 gene encodes the toll like receptor 4, which is found on the surface of immune competent cells and plays an important role in autoimmunity, occupying a significant position as a receptor recognizing pathogen-associated molecular patterns (PAMPs).

TLR4 has been shown to contribute to the host's susceptibility for periodontitis. TLR4 cytokine expression is significantly increased in both macrophages and gingival fibroblasts located in inflamed gingival tissues indicating its importance in the inflammatory process. The binding of bacteria lipopolysaccharides (LPS) to TLR4 on human gingival fibroblasts (HGFs) can activate various second messenger systems, such as the nuclear factor κ B (NF- κ B) pathway. These processes are closely associated with the destruction of periodontal tissue. The two TLR 4 variants, 896 A>G and 1196 C>T, reported here, are frequently inherited together.

Result: GG

Individuals with TLR4 GG genotype have an increased risk for chronic periodontitis and are more susceptible to general infection with gram-negative bacteria.

Result: TT

Individuals with TLR4 TT genotype have an increased risk for chronic periodontitis.



Inflammation and acquired immunity

Acquired immunity, also referred to as adaptive immunity, develops over your lifetime as you are exposed to a vaccine, an infection or disease, or another person's antibodies. The acquired immune system is composed of specialised, systemic cells and processes that eliminates pathogens by preventing their growth.

Inflammation, and the inflammatory response, are important factors to consider with acquired immunity. Cytokines are molecules that are used for cell signalling. A major function of cytokines is for them to be able to communicate with neighbouring or distant cells in order to initiate an immune response. Cytokines are also used to trigger cell trafficking to a specific area of the body. Increased levels of pro-inflammatory cytokines have been associated with chronic, low-grade inflammation, which is an important contributing factor to many chronic diseases including periodontitis. Chronic, low-grade inflammation, is also associated with an exaggerated response to an infection, increasing risk for adverse outcomes.



Inflammation and acquired immunity results

Genotype result table:

GENE NAME	GENE VARIATION	RESULT	GENE IMPACT
IL1A	889 C>T	CC	
	4845 G>T	GG	
IL1B	3954 C>T	CC	
	-511 T>C	AA	
IL1RN	2018 T>C	TT	
IL6	-174 C>G	GG	
IL17A	-197 G>A	GA	
MMP3	Lys45Glu (A>G)	GG	
TNFA	G>A	GG	

Inflammation and acquired immunity (continued)



Gene and genotype specific information



IL-1

IL-1 has been increasingly implicated as an important leverage point in the inflammatory cascade, and IL-1 expression is therefore key in the pathogenesis of several chronic diseases. The biological activity of IL-1 involves the two agonists – IL-1alpha (IL-1A) and IL-1beta (IL-1B), specific IL-1 receptors, and an IL-1 receptor antagonist (IL-1RN), which is a negative regulator of the pro-inflammatory response.

IL-1 is a potent regulator of the extracellular matrix, where increased expression leads to activation of osteoclasts and thus cellular degradation.

Certain genetic variations in IL-1A, IL-1B and IL-1RN lead to a more active inflammatory response, and have been associated with increased risk for a number of chronic diseases including periodontitis.

Result:

Individuals with variations in IL-1A, IL-1B, or IL-1RN are associated with having increased IL-1 plasma concentrations and have been linked with a number of pro-inflammatory chronic diseases, including periodontitis, coronary artery disease, certain autoimmune diseases, and cancers. Increase intake of nutrients known to inhibit the secretion of pro-inflammatory markers.



IL6

Interleukin 6 is a pro-inflammatory cytokine that plays a crucial role in inflammation and regulates expression of CRP. This gene has been implicated in the development of periodontal disease and other chronic inflammatory diseases of lifestyle including low-grade chronic obesity and, insulin resistance, dyslipidaemia and increased risk for cardiovascular disease.

Result: GG

The GG genotype leads to normal expression of this gene and thus is not associated with an increased risk for chronic, low-grade inflammation.



Gene and genotype specific information (continued)



IL17A

IL-17A encodes the proinflammatory cytokine, Interleukin-17A, which is the most studied member of the IL-17 family, and is secreted by T helper-17 (Th-17) cells. It is a powerful activator of neutrophils and is contributing factor in the pathogenesis of various autoimmune and inflammatory diseases. IL-17 stimulates the production and expression of TNF-alpha and IL-1 beta by human macrophages and regulates antimicrobial activity of molecules such as calgranulins, β -defensins, and mucin. Increased levels IL-17 have been documented in the saliva of patients with periodontal disease.

Result: GA

The GA genotype leads to increased expression of IL-17A and higher serum levels of the proinflammatory cytokine have been found in IL-17A GA genotype carriers. The risk variant is associated with chronic periodontitis and is linked to worse clinical and inflammatory periodontal parameters. The GA genotype has also been shown to be a major contributor to the development of a number of inflammatory diseases including coronary artery disease and osteoarthritis. Obesity, a modern Western diet, and high levels of oral bacteria stimulate the production of IL-17A.



MMP3

Matrix metalloproteinases (MMPs) are involved in the breakdown of the extracellular matrix in normal physiological processes, such as embryonic development and tissue remodelling, as well as in disease processes, such as arthritis and metastasis. MMPs have been implicated as a significant mediator of periodontitis-associated tissue breakdown, due to their role in the pathological destruction of extracellular matrix and immune responses related to periodontal inflammation. MMP3 (stromelysin-1) can degrade collagen in the basal membrane and can induce the synthesis of other MMPs, such as MMP1, MMP8 and MMP9.

Result: GG

Individuals with the GG genotype are at greater risk of developing periodontitis. The SNP is also related to a greater number of periodontal pockets as well as increased expression of MMP8 in the saliva. The MMP3 Lys45Glu variant alters the transcription levels of this matrix-degrading enzyme, affecting its ability to degrade connective tissue and thus may lead to the progression of periodontitis. Lifestyle factors, such as smoking, diet, and psychosocial conditions as well as comorbidities, such as diabetes and exposure to certain harmful bacteria in the mouth all contribute toward the development of periodontal disease.



TNFA

Tumour necrosis factor- α (TNF α), a proinflammatory cytokine secreted by both macrophages and adipocytes has been shown to alter whole body glucose homeostasis, and has been implicated in the development of several chronic diseases of lifestyle.

Result: GG

The GG genotype is associated with normal expression of this proinflammatory cytokine and thus no increased risk for chronic inflammatory diseases of lifestyle.



Sweet tooth

Higher sugar consumption has been strongly linked to the risk of obesity as well as developing tooth caries. Genetics contributes to the interindividual variability in sugar consumption, where sweet taste receptors and glucose sensors play an important role in influencing an individuals sweet craving behaviour and sugar intake and thus risk for tooth caries.



Sweet tooth results

Genotype result table:

GENE NAME	GENE VARIATION	RESULT	GENE IMPACT
SLC2A2	Thr110Ile (T>C)	TT	
TAS1R2	Ile191Val (G>A)	AA	

Gene and genotype specific information



SLC2A2

GLUT2, coded by the SLC2A2 gene, is a member of the facilitative glucose transport protein (GLUT) family and is expressed in the pancreas, liver, small intestine, kidney, and brain. GLUT2 facilitates the first step in glucose induced insulin secretion, with the entry of glucose into the pancreatic β cell. Because of its low affinity for glucose, it has been suggested as a glucose sensor, is considered to be important in the postprandial state, and is involved in food intake and regulation.

Result: TT

Individuals with the TT genotype have been found to consume significantly higher amounts of sugar than usual and have an increased risk for developing tooth caries. They have also been associated with having an elevated risk of type 2 diabetes.



TAS1R2

TAS1R2 encodes the taste receptor type 1 member 2, which is strongly involved in sensing sweet taste in foods, both from natural sugars as well as artificial sweeteners.

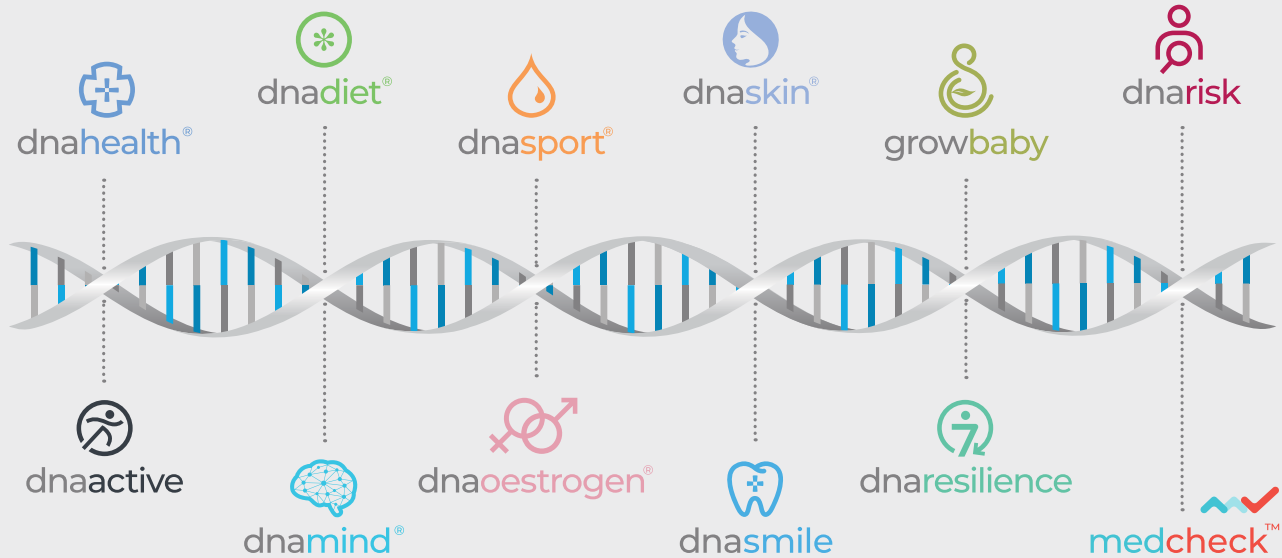
Result: AA

The AA genotype is associated with altered receptor function and individuals with this genotype have been reported to consume more high-sugar foods and sweet beverages, increasing susceptibility to developing tooth caries.



A lifetime of optimal health awaits you

Your genes do not change, which means our laboratories will only ever need one sample* from you. Throughout your life, as your health goals and priorities change, we can continue to provide valuable health insights from this single sample* to support your unique health journey.



*Requires finger prick blood spot sample collection

Our Commitment

DNALYSIS Biotechnology is continuously developing new tests with the highest standards of scientific rigour. Our commitment to ensuring the ethical and appropriate use of genetic tests in practice means that gene variants are only included in panels once there is sound motivation for their clinical utility and their impact on health outcomes.

ADVANCED | **ACTIONABLE** | **APPROPRIATE**
technology | interventions | use in practice

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Risks and Limitations:

DNALYSIS Biotechnology has a laboratory with standard and effective procedures in place for handling samples and effective protocols in place to protect against technical and operational problems. However as with all laboratories, laboratory error can occur; examples include, but are not limited to, sample or DNA mislabelling or contamination, failure to obtain an interpretable report, or other operational laboratory errors. Occasionally due to circumstances beyond DNALYSIS Biotechnology's control it may not be possible to obtain SNP specific results.

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