

Research Article

Journal of Applied Material Science & Engineering Research

Viscoelastic or Viscoplastic Glucose Theory (VGT #43): A Research of Statistics Methods Influences on Math-Physical Medicine Subjects by using Statistical Correlations as Initial Examination Tool to Explore the Inter-Relationships of Three Selected Biomarkers, CVD Risk %, Averaged Daily Glucose (eAG), and HbA1C % (A1C), within Three Different Selected Time-Windows, 15-Months, 46-Months, and 8.2 Years (98-Months), also Applying VGT and Viscoelastic Perturbation Model to Predict Three Sets of Predicted CVD Risk % Based on the GH-Method: Math-Physical Medicine (No. 624)

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Submitted: 15 Jun 2022; Accepted: 20 Jun 2022; Published: 05 Jul 2022

Citation: Gerald C Hsu. (2022). Viscoelastic or Viscoplastic Glucose Theory (VGT #43): A Research of Statistics Methods Influences on Math-Physical Medicine Subjects by using Statistical Correlations as Initial Examination Tool to Explore the Inter-Relationships of Three Selected Biomarkers, CVD Risk %, Averaged Daily Glucose (eAG), and HbA1C % (A1C), within Three Different Selected Time-Windows, 15-Months, 46-Months, and 8.2 Years (98-Months), also Applying VGT and Viscoelastic Perturbation Model to Predict Three Sets of Predicted CVD Risk % Based on the GH-Method: Math-Physical Medicine (No. 624). J App Mat Sci & Engg Res, 6(2), 01-12

Abstract

Since 2012, the author has been collecting his body weight and finger-piercing glucose values each day. In addition, he accumulates medical conditions data including a combination of data for blood pressure (BP), heart rate (HR), and blood lipids along with lifestyle details (LD). Based on the collected big data, he further organized them into two main groups. The first is the medical conditions group (MC) with 4 categories: weight, glucose, blood pressures, and blood lipids. The second is the LD group with 6 categories: food & diet, exercise, water intake, sleep, stress, and daily routines. At first, he collected his data on a daily basis since Y2012 and then calculated a unique combined daily score for each of the 10 categories within the MC and LD groups. The combined scores of the 2 groups, 10 categories, and 500+ detailed elements constitute an overall "metabolism index (MI) model". This MI model includes the root causes from 6 major lifestyle inputs and symptoms from 4 rudimentary chronic diseases: obesity, diabetes, hypertension, and hyperlipidemia. Therefore, it can serve as the foundation and building block for his additional research work that can expand into various diseases associated with different organs, such as cardiovascular diseases (CVD), and various cancers.

As we know, lifestyle details cause rudimentary chronic diseases which further influence more complicated diseases, such as heart problems (CVD & CHD), chronic kidney disease (CKD), stroke, diabetic retinopathy (DR), neuropathy, hypothyroidism, and others. Some genetic conditions and lifetime unhealthy habits, such as smoking, alcohol consumption, illicit drug use would account for approximately 15% to 25% of the root cause for rudimentary chronic diseases & their complications, including cancers and dementia. In addition to the genetic conditions, lifetime bad habits, and lifestyle details, some external factors, i.e., environmental factors, such as radiation, air and water pollution, food poison and pollution, toxic chemicals, and hormonal therapy, can also contribute to the causes for a variety of cancers. All of the above-mentioned diseases fall into the category of "symptoms" which are the "root-causes" of poor and unhealthy lifestyles.

In articles No. 622 (over 15-month period), No. 623 (over 46-month period), and No. 624 (over 98-month period or 8.2 years), the author applies the viscoelasticity and viscoplasticity theories to conduct his research to discover some hidden behavior or possible relationship among 3 key biomarkers, CVD risk probability %

(CVD risk, a symptom disease), daily average glucose mg/dL (eAG, either CGM sensor or finger-pierced), and its related HbA1C % (A1C, either finger A1C or sensor A1C). The hidden behaviors and possible inter-relationships among the three biomarkers are "time-dependent" which change from time to time. This is why he applies viscoelastic & viscoplastic theories (VGT) from physics and engineering to conduct his medical research work.

The author previously conducted similar analyses for these same datasets of selected biomarkers using a traditional statistical regression method. Generally speaking, statistical methods only deal with numerical characteristics of collected datasets and do not connect with the internal physical characteristics or behaviors of biomarkers in internal organs. In other words, any statistical method has no implicit connection with any internal biophysical behavior or biomedical phenomenon. Incidentally, the accuracy and applicability of results using any statistical method are heavily dependent on internal characteristics of data sample, size of dataset, and the time-window coverage of the chosen data. Therefore, we must be careful in selecting appropriate statistical methods and treat their analysis conclusions cautiously.

For example, in this analysis, the author performed three basic correlation analyses of the same dataset for three biomarkers, CVD risk %, eAG, and A1C %, by choosing three different time-windows, 15-months, 46-months, and 98-months (8.2 years). The following displayed results using "daily data" have shown the vast differences among the three statistical correlation analysis results:

- (1) Correlations using 15 months (10/1/2020 2/28/2022): CVD vs. eAG = -55%; CVD/A1C = -59%, eAG vs. A1C = 97%
- (2) Correlations using 46 months (8/8/2018 3/3/2022): CVD vs. eAG = 70%; CVD/A1C = 70%, eAG vs. A1C = 99%
- (3) Correlations using 418 months in 8.2 years (1/1/2014 3/3/2022): CVD vs. eAG = 80%; CVD/A1C = 79%, eAG vs. A1C = 99%

It is evident that the 15-month window results in negative correlations. The 46-month window provides more moderate correlations, while the 98-month window provides high correlations. This has proven that a longer time-window coverage (not necessarily more data elements) would usually provide additional insights of certain biomarker behaviors; therefore, a complete and more accurate picture could be displayed. This particular statement can also be interpreted that a clear CVD risk's picture can be seen with a longer coverage period of time with meaningful inside information of the biomarker, not just the inclusion of more data elements.

Nevertheless, the correlations between eAG and A1C, regardless of the time-window selection, are always high (97% to 99%). This is due to the fact that, by definition, A1C is determined by the 90-days moving average of eAG.

Therefore, a quick pre-examination by using correlations of the three selected datasets would provide some hints regarding the effectiveness and usefulness for his later analysis results. Obviously, from these studies using 3 different time windows, 15-months or 1.3 years, 46-months or 3.8 years, and 98-month or 8.2 years, a wider time-window coverage of data behaviors, not more data points in the selected time window, usually offers a better understanding of the inner-characteristics for the biomarker datasets which then achieve accurate or useful results.

The following defined equations are used to establish the stress-strain diagram in a space-domain (SD):

```
strain = \varepsilon (CVD risk %)
= individual CVD risk % at present time
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Stress

- = σ (based on change rate of strain, CVD risk, multiplying with a viscosity factor, eAG or A1C)
- $=\eta * (d\varepsilon/dt)$
- $= \eta * (d-strain/d-time)$
- = (viscosity factor η using individual eAG or A1C at present time) * (CVD risk at present time CVD risk at previous time)

Next, he applies the viscoelastic perturbation model to calculate the following predicted CVD risk %.

Perturbed or predicted CVD risk %

= strain value (CVD risk %) at present time + stress value at present time (i.e., CVD risk change rate * eAG or A1C) * amplification factor

For example, in the 8.2 years' case, the selected amplification factor for A1C is 1.0 and for eAG is 0.0625 (or divided by 16) which allows the two stress scales (Y-axis scales) to be on even ground.

To offer a simple explanation to readers who do not have a physics or engineering background, the author includes a brief excerpt from Wikipedia regarding the description of basic concepts for elasticity and plasticity theories, viscoelasticity and viscoplasticity theories from the disciplines of engineering and physics in the Method section.

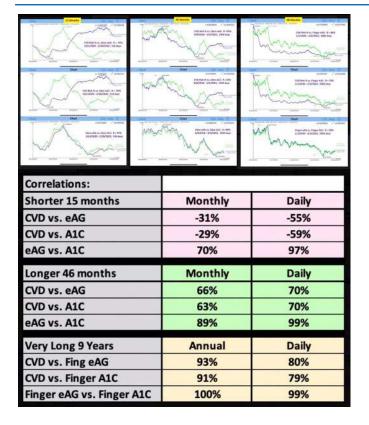
In conclusion, the following three observations outline the findings from this research work of statistical influences on math-physical medical research projects by selecting three different time-windows of CVD risk datasets:

- (1) From the pre-examination using correlation studies in a time-domain, the smallest data points of 9 (8.2 years with 98-months of information) offers the highest correlations among CVD risk, eAG, and A1C. The shortest time frame of 15-months provides the worst correlations among CVD risk, eAG, and A1C, while the 46-months' time window is situated in the middle. These findings prove that the insight information or behaviors of a biomarker is far more important than the sheer number of data points. Usually, a longer time window of a dataset would reveal more insight into the behavior of its associated biomarker, but this statement is still depending upon case by case in medicine. Generally speaking, the higher positive correlations provide a hint that this longer time-window analysis can usually produce meaningful and useful results.
- (2) In the stress-strain diagram in a space-domain (SD), the three curves have different appearances from each other. This is due to the following two reasons. The first is that the strain information (CVD risk) is different in different time-windows. The second is that its associated viscosity factors i.e., eAG or A1C are also different at various time instant. Nevertheless, due to the author's continuous improvement on his eAG (higher in Y2014 and then dropping to its lowest in Y2022), and eAG associated A1C, his strain value (CVD risk) is then moving from the right side of the diagram toward the left side. On the stress-scale, both different strain rates and viscosity factors determine the y-scale values. For an overall viewpoint, the 8.2-years' curve shows a complete view from Y2014 to Y2022 with the biggest CVD risk change rate during the sub-period from Y2014 to Y2017. The 46-months' curve reflects a moderate and confined area for CVD risks and their change rates. The 15-months' curve is the most meaningless case due to its insufficient inclusion of biomarker behaviors. It should be noted that these stress scales have been adjusted by using (A1C) and (eAG / 20 or 16) in order to achieve a better viewing of shape and comparison of the stress-strain curves.
- (3) Using the viscoelastic perturbation model, a waveform comparison study of the metabolism calculated CVD risk % against two predicted CVD risks, using eAG or A1C, can be done. For both time-window cases of 15-months and 46-months, their negative or moderate correlations are not high enough. As a result, the introduction of a "stronger curve vibrations" effect from the perturbation factors i.e., the "stress element", therefore produces less-meaningful or not so useful CVD risk predictions. On the contrary, the 98-month's case only has 9 data points but with very high correlation results. It can produce highly accurate predicted CVD risks using the visco-perturbation model.

In summary, this particular report shows that if using a shorter 15-month dataset, it would result in unsatisfactory results. If using a moderate 46-month dataset, the CVD risk % would have higher correlations with both eAG and A1C, but their associated predicted CVD risks are still not quite useful due to its moderate amount of biomarker information. Only using the 8.2 years (9 data points covering a 98-month period) case can reveal a better picture of inter-relations between CVD risk versus both eAG and A1C.

The author attempts to interpret the above findings using biomedical terminology (not math-physical languages) as follows:

Heart problems, such as CVD, do not occur suddenly, and are usually associated with a patient's lifestyle over a longer period of time. The majority of CVD patients (75% to 80%) also reflect diabetes conditions with hyperglycemia (high eAG and high A1C). Therefore, for patients with both heart problems and diabetes, in order to lower their risks of having CVD, a longer period of continuous efforts on lowering their eGA and A1C is necessary.



Introduction

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Methods

Elasticity, Plasticity, Viscoelasticity and Viscoplasticity

The Difference Between Elastic Materials and Viscoelastic Materials

(from "Soborthans, innovating shock and vibration solutions")

What are Elastic Materials?

Elasticity is the tendency of solid materials to return to their original shape after forces are applied on them. When the forces are removed, the object will return to its initial shape and size if the material is elastic.

What are Viscous Materials?

Viscosity is a measure of a fluid's resistance to flow. A fluid with large viscosity resists motion. A fluid with low viscosity flows. For example, water flows more easily than syrup because it has a lower viscosity. High viscosity materials might include honey, syrups, or gels – generally things that resist flow. Water is a low viscosity material, as it flows readily. Viscous materials are thick or sticky or adhesive. Since heating reduces viscosity, these materials don't flow easily. For example, warm syrup flows more easily than cold.

What is Viscoelastic?

Viscoelasticity is the property of materials that exhibit both viscous and elastic characteristics when undergoing deformation. Synthetic polymers, wood, and human tissue, as well as metals at high temperature, display significant viscoelastic effects. In some applications, even a small viscoelastic response can be significant.

Elastic Behavior Versus Viscoelastic Behavior

The difference between elastic materials and viscoelastic materials is that viscoelastic materials have a viscosity factor and the elastic ones don't. Because viscoelastic materials have the viscosity factor, they have a strain rate dependent on time. Purely elastic materials do not dissipate energy (heat) when a load is applied, then removed; however, a viscoelastic substance does. The following brief introductions are excerpts from Wikipedia:

"Elasticity (Physics)

Physical property when materials or objects return to original shape after deformation

In physics and materials science, elasticity is the ability of a body to resist a distorting influence and to return to its original size and shape when that influence or force is removed. Solid objects will deform when adequate loads are applied to them; if the material is elastic, the object will return to its initial shape and size after removal. This is in contrast to plasticity, in which the object fails to do so and instead remains in its deformed state. The physical reasons for elastic behavior can be quite different for different materials. In metals, the atomic lattice changes size and shape when forces are applied (energy is added to the system). When forces are removed, the lattice goes back to the original lower energy state. For rubbers and other polymers, elasticity is caused by the stretching of polymer chains when forces are applied.

Hooke's law states that the force required to deform elastic objects should be directly proportional to the distance of deformation, regardless of how large that distance becomes. This is

known as perfect elasticity, in which a given object will return to its original shape no matter how strongly it is deformed. This is an ideal concept only; most materials which possess elasticity in practice remain purely elastic only up to very small deformations, after which plastic (permanent) deformation occurs.

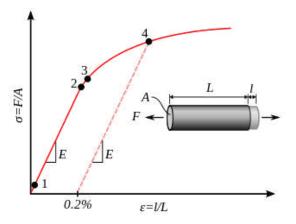
In engineering, the elasticity of a material is quantified by the elastic modulus such as the Young's modulus, bulk modulus or shear modulus which measure the amount of stress needed to achieve a unit of strain; a higher modulus indicates that the material is harder to deform. The material's elastic limit or yield strength is the maximum stress that can arise before the onset of plastic deformation.

Plasticity (Physics)

Deformation of a solid material undergoing non-reversible changes of shape in response to applied forces.

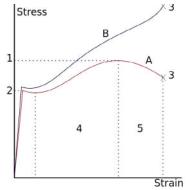
In physics and materials science, plasticity, also known as

is the ability of a solid material to undergo permanent deformation, a non-reversible change of shape in response to applied forces. For example, a solid piece of metal being bent or pounded into a new shape displays plasticity as permanent changes occur within the material itself. In engineering, the transition from elastic behavior to plastic behavior is known as yielding.



Stress-strain curve showing typical yield behavior for nonferrous alloys

- 1. True elastic limit
- 2. Proportionality limit
- 3. Elastic limit
- 4. Offset yield strength



A stress-strain curve typical of structural steel.

- 1: Ultimate strength
- 2: Yield strength (yield point)
- 3: Rupture
- 4: Strain hardening region
- 5: Necking region
- A: Apparent stress (F/A0)
- B: Actual stress (F/A)

Plastic deformation is observed in most materials, particularly metals, soils, rocks, concrete, and foams. However, the physical mechanisms that cause plastic deformation can vary widely. At a crystalline scale, plasticity in metals is usually a consequence of dislocations. Such defects are relatively rare in most crystalline materials, but are numerous in some and part of their crystal structure; in such cases, plastic crystallinity can result. In brittle materials such as rock, concrete and bone, plasticity is caused predominantly by slip at microcracks. In cellular materials such as liquid foams or biological tissues, plasticity is mainly a consequence of bubble or cell rearrangements, notably T1 processes.

For many ductile metals, tensile loading applied to a sample will cause it to behave in an elastic manner. Each increment of load is accompanied by a proportional increment in extension. When the load is removed, the piece returns to its original size. However, once the load exceeds a threshold – the yield strength – the extension increases more rapidly than in the elastic region; now when the load is removed, some degree of extension will remain.

Elastic deformation, however, is an approximation and its quality depends on the time frame considered and loading speed. If, as indicated in the graph opposite, the deformation includes elastic deformation, it is also often referred to as "elasto-plastic deformation" or "elastic-plastic deformation".

Perfect plasticity is a property of materials to undergo irreversible deformation without any increase in stresses or loads. Plastic materials that have been hardened by prior deformation, such as cold forming, may need increasingly higher stresses to deform further. Generally, plastic deformation is also dependent on the deformation speed, i.e. higher stresses usually have to be applied to increase the rate of deformation. Such materials are said to deform visco-plastically."

Viscoelasticity

Property of materials with both viscous and elastic characteristics under deformation

In materials science and continuum mechanics, viscoelasticity is the property of materials that exhibit both viscous and elastic characteristics when undergoing deformation. Viscous materials, like water, resist shear flow and strain linearly with time when a stress is applied. Elastic materials strain when stretched and immediately return to their original state once the stress is removed.

Viscoelastic materials have elements of both of these properties and, as such, exhibit time-dependent strain. Whereas elasticity is usually the result of bond stretching along crystallographic planes in an ordered solid, viscosity is the result of the diffusion of atoms or molecules inside an amorphous material.

In the nineteenth century, physicists such as Maxwell, Boltzmann, and Kelvin researched and experimented with creep and recovery of glasses, metals, and rubbers. Viscoelasticity was further examined in the late twentieth century when synthetic polymers were engineered and used in a variety of applications. Viscoelasticity calculations depend heavily on the viscosity variable, η . The inverse of η is also known as fluidity, φ . The value of either can be derived as a function of temperature or as a given value (i.e. for a dashpot).

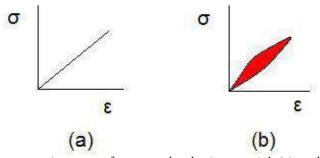
Depending on the change of strain rate versus stress inside a material, the viscosity can be categorized as having a linear, non-linear, or plastic response. When a material exhibits a linear response it is categorized as a Newtonian material. In this case the stress is linearly proportional to the strain rate. If the material exhibits a non-linear response to the strain rate, it is categorized as Non-Newtonian fluid. There is also an interesting case where the viscosity decreases as the shear/strain rate remains constant. A material which exhibits this type of behavior is known as thixotropic. In addition, when the stress is independent of this strain rate, the material exhibits plastic deformation. Many viscoelastic materials exhibit rubber-like behavior explained by the thermodynamic theory of polymer elasticity.

Cracking occurs when the strain is applied quickly and outside of the elastic limit. Ligaments and tendons are viscoelastic, so the extent of the potential damage to them depends both on the rate of the change of their length as well as on the force applied.

A viscoelastic material has the following properties:

- 1. hysteresis is seen in the stress-strain curve
- 2. stress relaxation occurs: step constant strain causes decreasing stress
- 3. creep occurs: step constant stress causes increasing strain
- 4. its stiffness depends on the strain rate or the stress rat

Elastic Versus Viscoelastic Behavior



Stress-strain curves for a purely elastic material (a) and a viscoelastic material (b). The red area is a hysteresis loop and shows the amount of energy lost (as heat) in a loading and unloading cycle. It is equal to

$\oint \sigma d\varepsilon$ where σ is stress and ε is strain.

Unlike purely elastic substances, a viscoelastic substance has an elastic component and a viscous component. The viscosity of a viscoelastic substance gives the substance a strain rate dependence on time. Purely elastic materials do not dissipate energy (heat) when a load is applied, then removed. However, a viscoelastic substance dissipates energy when a load is applied, then removed. Hysteresis is observed in the stress-strain curve, with the area of the loop being equal to the energy lost during the loading cycle. Since viscosity is the resistance to thermally activated plastic deformation, a viscous material will lose energy through a loading cycle. Plastic deformation results in lost energy, which is uncharacteristic of a purely elastic material's reaction to a loading cycle.

Specifically, viscoelasticity is a molecular rearrangement. When a stress is applied to a viscoelastic material such as a polymer, parts of the long polymer chain change positions. This movement or rearrangement is called "creep". Polymers remain a solid material even when these parts of their chains are rearranging in order to accompany the stress, and as this occurs, it creates a back stress in the material. When the back stress is the same magnitude as the applied stress, the material no longer creeps. When the original stress is taken away, the accumulated back stresses will cause the polymer to return to its original form. The material creeps, which gives the prefix visco-, and the material fully recovers, which gives the suffix -elasticity.

Viscoplasticity

Viscoplasticity is a theory in continuum mechanics that describes the rate-dependent inelastic behavior of solids. Rate-dependence in this context means that the deformation of the material depends on the rate at which loads are applied. The inelastic behavior that is the subject of viscoplasticity is plastic deformation which means that the material undergoes unrecoverable deformations when a load level is reached. Rate-dependent plasticity is important for transient plasticity calculations. The main difference between rate-independent plastic and viscoplastic material models is that the latter exhibit not only permanent deformations after the application of loads but continue to undergo a creep flow as a function of time under the influence of the applied load.

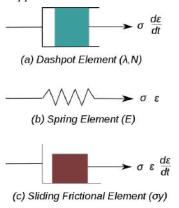


Figure 1. Elements used in one-dimensional models of viscoplastic materials.

The elastic response of viscoplastic materials can be represented in one-dimension by Hookean spring elements. Rate-dependence can be represented by nonlinear dashpot elements in a manner similar to viscoelasticity. Plasticity can be accounted for by adding sliding frictional elements as shown in Figure 1. In the figure E is the modulus of elasticity, λ is the viscosity parameter and N is a power-law type parameter that represents non-linear dashpot $[\sigma(d\varepsilon/dt) = \sigma = \lambda(d\varepsilon/dt)(1/N)]$. The sliding element can have a yield stress (σy) that is strain rate dependent, or even constant, as shown in Figure 1c.

Viscoplasticity is usually modeled in three-dimensions using overstress models of the Perzyna or Duvaut-Lions types. In these models, the stress is allowed to increase beyond the rate-independent yield surface upon application of a load and then allowed to relax back to the yield surface over time. The yield surface is usually assumed not to be rate-dependent in such models. An alternative approach is to add a strain rate dependence to the yield stress and use the techniques of rate independent plasticity to calculate the response of a material

For metals and alloys, viscoplasticity is the macroscopic behavior caused by a mechanism linked to the movement of dislocations in grains, with superposed effects of inter-crystalline gliding. The mechanism usually becomes dominant at temperatures greater than approximately one third of the absolute melting temperature. However, certain alloys exhibit viscoplasticity at room temperature (300K). For polymers, wood, and bitumen, the theory of viscoplasticity is required to describe behavior beyond the limit of elasticity or viscoelasticity.

In general, viscoplasticity theories are useful in areas such as

- the calculation of permanent deformations,
- the prediction of the plastic collapse of structures,
- the investigation of stability,
- crash simulations,
- systems exposed to high temperatures such as turbines in engines, e.g. a power plant,
- dynamic problems and systems exposed to high strain rates.

Phenomenology

For a qualitative analysis, several characteristic tests are performed to describe the phenomenology of viscoplastic materials. Some examples of these tests are

- 1. hardening tests at constant stress or strain rate,
- 2. creep tests at constant force, and
- 3. stress relaxation at constant elongation.

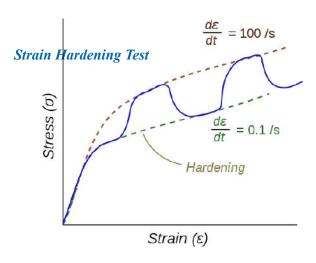


Figure 2. Stress-strain response of a viscoplastic material at different strain rates.

The dotted lines show the response if the strain-rate is held constant. The blue line shows the response when the strain rate is changed suddenly.

One consequence of yielding is that as plastic deformation proceeds, an increase in stress is required to produce additional strain. This phenomenon is known as Strain/Work hardening. For a viscoplastic material the hardening curves are not significantly different from those of rate-independent plastic material. Nevertheless, three essential differences can be observed.

- 1. At the same strain, the higher the rate of strain the higher the stress
- 2. A change in the rate of strain during the test results in an immediate change in the stress–strain curve.
- 3. The concept of a plastic yield limit is no longer strictly applicable.

The hypothesis of partitioning the strains by decoupling the elastic and plastic parts is still applicable where the strains are small i.e.,

$$\varepsilon = \varepsilon e + \varepsilon v p$$

where εe is the elastic strain and εvp is the viscoplastic strain.

To obtain the stress-strain behavior shown in blue in the figure, the material is initially loaded at a strain rate of 0.1/s. The strain rate is then instantaneously raised to 100/s and held constant at that value for some time. At the end of that time period the strain rate is dropped instantaneously back to 0.1/s and the cycle is continued for increasing values of strain. There is clearly a lag between the strain-rate change and the stress response. This lag is modeled quite accurately by overstress models (such as the

Perzyna model) but not by models of rate-independent plasticity that have a rate-dependent yield stress."

Results

Figure 1 displays the data table and calculated results of this study.

JV - 11 - 112				Name of		~ ~		-	_	En more		
11/25/21	Y	X1		11/25/21	Strain	Stress 1	Stress 2			Strain (CVD)		
Period	CVD Risk %	Libre A1C		Period	Strain (CVD)	Stress (A1C)	Stress (eAG			Calculated CVD	Pred. CVD (A1C)	Pred. CVD (eAG)
Y2020M10	48	5.9		/2020M10	48	0.0	0.0	Y2020M1		48	48	48
Y2020M11	48	5.8		/2020M11	48	-1.7	-1.6	Y2020M1		48	46	46
Y2020M12	50	5.9		72020M12	50	13.1	12.8	Y2020M1		50	63	63
Y2021M01	52	6.0		/2021M01	52	10.2	10.1	Y2021M0		52	62	62
Y2021M02	49	6.3		r2021M02	49	-19.1	-18.3	Y2021MC		49	29	30
Y2021M03	50	6.4		/2021M03	50	10.4	9.7	Y2021M0		50	61	60
Y2021M04	52	6.5		/2021M04	52	12.4	11.0	Y2021M0		52	65	63
Y2021M05	53	6.2		72021M05	53	7.7	6.8	Y2021M0		53	61	60
Y2021M06	53	6.1		72021M06	53	-2.2	-2.1	Y2021M0		53	51	51
Y2021M07	53	6.0		72021M07	53	1.6	1.4	Y2021M0		53	55	55
Y2021M08	53	5.8		72021M08	53	-0.7	-0.6	Y2021M0		53	52	53
Y2021M09	54	5.6		/2021M09	54	3.4	3.3	Y2021M0		54	57	57
Y2021M10	54	5.7		/2021M10	54	1.1	1.0	Y2021M1		54	55	55
Y2021M11	54	5.8		72021M11	54	1.6	1.5	Y2021M1	D.	54	56	56
Y2021M12	53	5.7		72021M12	53	-8.3	-7.6	Y2021M1		53	44	45
Y2022M01	53	5.8		72022M01	53	-1.3	-1.2	Y2022M0		53	51	51
Y2022M02	51	5.9	105	/2022M02	51	-9.8	-8.8	Y2022M0	2	51	41	42
Average	52	6.0	110	Average	52	1.1	1.0	Average		51.7	52.7	52.7
Correlation	100%	-29%	-31%					Correlati	on	100%	41%	42%
Correlation			70%					Accuracy			98%	98%
Aller Park						100		-				
		-										
3/4/22	Y	X1	X2	No.	Strain	Stress		Stress 2		Strain (CVD)		
Period	CVD Risk %	Libre A1C	Libre eAG	Period	Strain (CVD)		1C) Street			Calculated CVD	Pred. CVD (A1C)	Pred. CVD (eAG)
Y2018M5	53	6,5	120	Y2018M5	53	0.0			018M5	53	53	53
Y2018M6 Y2018M7	55 55	7.0	134	Y2018M6 Y2018M7	55 55	10.3			018M6 018M7	55	65	65 59
Y2018M8	57	6.9	132	Y2018M8	57	7,9			018M8	57	64	64
Y2018M9	53	7.0	128	Y2018MS	53	-24.8		-22.5 Y2	018M9	53	28	31
Y2018M10	57	7.1	135	Y2018M10	57	28.1			DIBMIO	57	85	84
Y2018M11 Y2018M12	58 54	7.1	130 131	Y2018M11 Y2018M12	58 54	5.2	N.		018M11 018M12	58 54	63	63 26
Y2018M12 Y2019M1	56	7.1	131	Y2018M12 Y2019M1	56	20.3			018M1Z 019M1	56	77	74
Y2019M2	57	6.9	128	Y2019M2	57	1.3			019M2	57	58	58
Y2019M3	59	6.9	132	Y2019M3	59	14.6	P	13.9 Y2	019M3	59	73	73
Y2019M4	60	7,1	132	Y2019M4	60	5.4	- 2		019M4	60	65	65
Y2019M5 Y2019M6	57 60	7.0	126 149	Y2019M5 Y2019M6	57 60	-17.3 17.7			019M5 019M6	57 60	77	41 78
Y2019M7	56	7.4	129	Y2019M7	56	-27.1			019M7	56	29	32
Y2019M8	56	7.1	129	Y2019M8	56	2.3		2.1 Y2	019MB	56	58	58
Y2019M9	55	7.1	133	Y2019M9	55	-7.7	- 3		019M9	55	47	48
Y2019M10	54	7.2	128	Y2019M10	54	-9.7			019M10	54	44	45
Y2019M11 Y2019M12	56 53	7.1	136 130	Y2019M11 Y2019M12	56 53	15.1			019M11 019M12	56 53	71 36	70 37
Y2020M1	54	7.0	131	Y2020M1	54	3.3			020M1	54	57	57
Y2020M2	53	6.9	218	Y2020M2	53	-9.6		-8.1 Y2	020M2	53	43	44
Y2020M3	52	6,7	128	Y2020M3	52	4.5			020143	32	47	47
Y2020M4 Y2020M5	53 53	6.9	129	Y2020M4 Y2020M5	53	4.3			020M4 020M5	53	62 52	51 52
V2020M6	53	6.3	116	Y2020M5	53	1,4			020M6	53	55	54
Y2020M7	51	6.3	113	Y2020M7	51	-10.1	N .	-9.0 Y2	020M7	51	41	42
Y2020MII	52	6,1	111	Y2020M8	52	0.7			020MB	52	52	52
Y2020M9	51	6.1	112	Y2020M9	51	-1.8			020M9	51	50	50
Y2020M10 Y2020M11	48	5.8	107	Y2020M10 Y2020M11	48	20.0 -1.7			020M10 020M11	48	28 46	30 46
Y2020M12	50	5.9	116	Y2020M12	50	13.1			020M12	50	63	63
Y2021M01	52	6.0	119	Y2021M01	52	10.2			021M01	52	62	62
Y2021M02	49	6,3	121	Y2021M02	49	-19.1			021M02	49	29	30
Y2021M03 Y2021M04	50 52	6,4	120	Y2021M03 Y2021M04	50 52	10.4			021M03 021M04	50 52	61 65	60 63
Y2021M04 Y2021M05	53	6.2	109	Y2021M04 Y2021M05	53	7.7			021M04 021M05	53	61	60
Y2021M0€	53	6.1	113	Y2021M06	53	-2.2	- 8		021M06	53	51	51
Y2021M07	53	6.0	107	Y2021M07	53	1.6		1.4 Y2	021M07	53	55	55
Y2021M08	53 54	5.8	101	Y2021M08 Y2021M09	53	-0.7			021M08 021M09	53 54	52	53
Y2021M10	54	5.7	107	Y2021M10	54	1.1			021M09 021M10	54	55	55
Y2021M11	54	5,8	107	Y2021M11	54	1.6	8 0		021M11	54	56	56
Y2021M12	53	5.7	105	Y2021M12	53	-8.3		-7.6 YZ	021M12	53	44	45
Y2022M01 Y2032M02	53 51	5.8	105	Y2022M01 Y2022M02	53	-1.3			022M01 022M02	53	51 41	51 42
						-0.8				51		
Average	54	6,5	121	Average	54	-0.4	_		erage	53,7	53.3	53.5
Correlation	100%	63%	86%			_		Co	rrelation	100%	51%	53%
			92,16					AC			THE R. L.	- CO. C. W.
estreteour.												
agest a causain						rain	Stress	Stress	3/4/22	Y	Y1	YZ
3/4/22	Y	X1	X2	3/4/22	St					_		
3/4/22 Annual Period	Y CVD Risk						tress (A1C)	Stress (eAG/16		Calc. CVD Ri	sk Pert. CVD (Finge	er) Pert. CVD (Lab)
Annual Period Y2014	72.6	% Finger A1C % 7.9	Finger eAG (m	ng/dL) Period Y2014	CVD	Risk S	0.000	0.000	Y2014	Calc. CVD Ri	73%	73%
Annual Period Y2014 Y2015	72.6 61.5	% Finger A1C % 7.9 7.7	Finger eAG (m 135 129	y2014 Y2015	7 6 6	0 Risk 5 3% 2%	0.000 -0.851	0.000 -0.894	Y2014 Y2015	73% 62%	73% 61%	73% 61%
Annual Period Y2014 Y2015 Y2016	72.6 61.5 56.7	% Finger A1C % 7.9 7.7 7.0	Finger eAG (m 135 129 119	y2014 Y2015 Y2016	CVD 77 66	0 Risk S 3% 2% 7%	0.000 -0.851 -0.338	0.000 -0.894 -0.360	Y2014 Y2015 Y2016	73% 62% 57%	73% 61% 56%	73% 61% 56%
Annual Period Y2014 Y2015 Y2016 Y2017	72.6 61.5 56.7 55.3	% Finger AIC% 7.9 7.7 7.0 6.9	Finger eAG (m 135 129 119 117	y2014 Y2015 Y2016 Y2017	CVD 7: 6. 5.	0 Risk S 3% 2% 7% 5%	0.000 -0.851 -0.338 -0.095	0.000 -0.894 -0.360 -0.101	Y2014 Y2015 Y2016 Y2017	73% 62% 57% 55%	73% 61% 56% 55%	73% 61% 56% 55%
Annual Period Y2014 Y2015 Y2016 Y2017 Y2018	72.6 61.5 56.7 55.3 55.4	% Finger A1C % 7.9 7.7 7.0 6.9 6.8	Finger eAG (m 135 129 119 117 116	y2014 Y2014 Y2015 Y2016 Y2017 Y2018	CVD 7: 6. 5. 5. 5. 5. 5. 5. 5. 5. 5. 5. 5. 5. 5.	0 Risk 5 3% 29% 7% 5%	0.000 -0.851 -0.338 -0.095 0.008	0.000 -0.894 -0.360 -0.101 0.008	Y2014 Y2015 Y2016 Y2017 Y2018	73% 62% 57% 55% 55%	73% 61% 56% 55% 55%	73% 61% 56% 55%
Annual Period Y2014 Y2015 Y2016 Y2017 Y2018 Y2019	72.6 61.5 56.7 55.3 55.4 56.5	% Finger A1C % 7.9 7.7 7.0 6.9 6.8 6.7	Finger eAG (m 135 129 119 117 116 114	y2014 Y2014 Y2015 Y2016 Y2017 Y2018 Y2019	CVD 77 66 55 55 55 55 55 55 55 55 55 55 55 55	0 Risk 5 3% 2% 7% 5% 5%	0.000 -0.851 -0.338 -0.095 0.008 0.074	0.000 -0.894 -0.360 -0.101 0.008 0.079	Y2014 Y2015 Y2016 Y2017 Y2018 Y2019	73% 62% 57% 55% 55% 55%	73% 61% 56% 55% 55% 55%	73% 61% 56% 55% 55% 55%
Annual Period Y2014 Y2015 Y2016 Y2017 Y2018	72.6 61.5 56.7 55.3 55.4 56.5 51.6	% Finger A1C % 7.9 7.7 7.0 6.9 6.8	Finger eAG (m 135 129 119 117 116 114 106	y2014 Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020	CVD 77 66 55 55 55 55 55 55 55 55 55 55 55 55	0 Risk 5 3% 29% 7% 5%	0.000 -0.851 -0.338 -0.095 0.008	0.000 -0.894 -0.360 -0.101 0.008 0.079 -0.327	Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020	73% 62% 57% 55% 55%	73% 61% 56% 55% 55% 57% 57%	73% 61% 56% 55%
Annual Period Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020	72.6 61.5 56.7 55.3 55.4 56.5	% Finger A1C % 7.9 7.7 7.0 6.9 6.8 6.7 6.3	Finger eAG (m 135 129 119 117 116 114	y2014 Y2014 Y2015 Y2016 Y2017 Y2018 Y2019	CVD 77: 66: 55: 55: 55: 55: 55: 55: 55: 55: 55	0 Risk 5 3% 22% 7% 55% 57% 22%	0.000 -0.851 -0.338 -0.095 0.008 0.074 -0.310	0.000 -0.894 -0.360 -0.101 0.008 0.079	Y2014 Y2015 Y2016 Y2017 Y2018 Y2019	73% 62% 57% 55% 55% 55% 57% 52%	73% 61% 56% 55% 55% 55%	73% 61% 56% 55% 55% 55% 57%
Annual Period Y2014 Y2015 Y2015 Y2017 Y2018 Y2019 Y2019 Y2020 Y2021 Y2021	72.6 61.5 56.7 55.3 55.4 56.5 51.6 52.4 51.7	% Finger AIC % 7.9 7.7 7.0 6.9 6.8 6.7 6.3 6.1 6.2	Finger eAG (m 135 129 119 117 116 114 106 105	y2014 Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020 Y2021 Y2022	CVD 77 76 66 55 55 55 55 55 55 55 55 55 55 55 55	0 Risk 5 3% 3% 7% 55% 57% 2% 2%	0.000 -0.851 -0.338 -0.095 0.008 0.074 -0.310 0.049 -0.046	0.000 -0.894 -0.360 -0.101 0.008 0.079 -0.327 0.052	Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020 Y2021 Y2021	73% 62% 57% 55% 55% 57% 52% 52% 52%	73% 61% 56% 55% 55% 57% 51% 52% 52%	73% 61% 56% 55% 55% 57% 51% 51% 52%
Annual Period Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020 Y2021 Y2022 Average	72.6 61.5 56.7 55.3 55.4 56.5 51.6 52.4 51.7	% Finger A1C % 7.9 7.7 7.0 6.9 6.8 6.7 6.3 6.1 6.2	Finger eAG (n 135 129 119 117 116 114 106 105 107	y2014 Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020 Y2021	CVD 77 76 66 55 55 55 55 55 55 55 55 55 55 55 55	0 Risk 5 3% 22% 7% 55% 57% 22% 22%	0.000 -0.851 -0.338 -0.095 0.008 0.074 -0.310 0.049	0.000 -0.894 -0.360 -0.101 0.008 0.079 -0.327 0.052	Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020 Y2021 Y2022 Average	73% 62% 57% 55% 55% 57% 52% 52% 52%	73% 61% 56% 55% 55% 57% 51% 52% 52% 52%	73% 61% 56% 55% 55% 57% 51% 52% 52% 56.9%
Annual Period Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2019 Y2020 Y2021 Y2022 Average Correlation	72.6 61.5 56.7 55.3 55.4 56.5 51.6 52.4 51.7	% Finger AIC % 7.9 7.7 7.0 6.9 6.8 6.7 6.3 6.1 6.2 6.9 91%	Finger eAG (m 135 129 119 117 116 114 106 105	y2014 Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020 Y2021 Y2022	CVD 77 76 66 55 55 55 55 55 55 55 55 55 55 55 55	0 Risk 5 3% 3% 7% 55% 57% 2% 2%	0.000 -0.851 -0.338 -0.095 0.008 0.074 -0.310 0.049 -0.046	0.000 -0.894 -0.360 -0.101 0.008 0.079 -0.327 0.052	Y2014 Y2015 Y2016 Y2017 Y2018 Y2019 Y2020 Y2021 Y2021	73% 62% 57% 55% 55% 57% 52% 52% 52%	73% 61% 56% 55% 55% 57% 51% 52% 52%	73% 61% 56% 55% 55% 57% 51% 51% 52%

Figure 1: Data table and calculation results of this study

Figure 2 shows monthly or annually correlations in a time-domain among CVD risk %, eAG, A1C over three time-windows.

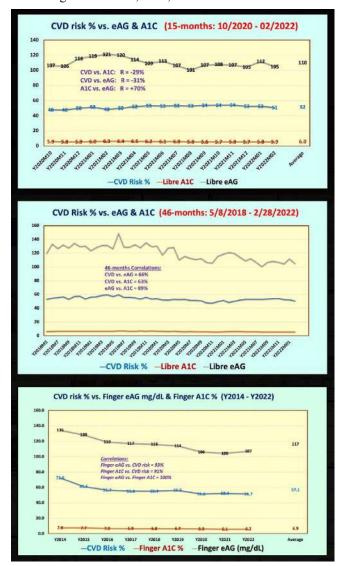


Figure 2: Monthly or annually Correlations among CVD risk %, eAG, and A1C over three time windows

Figure 3 depicts the results of three SD stress-strain diagrams of CVD risk % using A1C and (eAG / 20 or 16) as their viscosity factors (η).

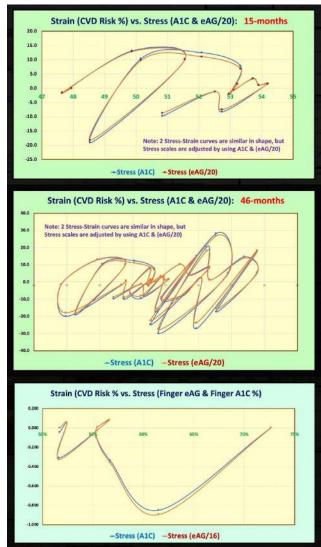


Figure 3: Three SD stress-strain diagrams

Figure 4 reflects a comparison chart between the calculated CVD risk % versus two predicted CVD risks using viscoelastic perturbation model, within 3 time windows.

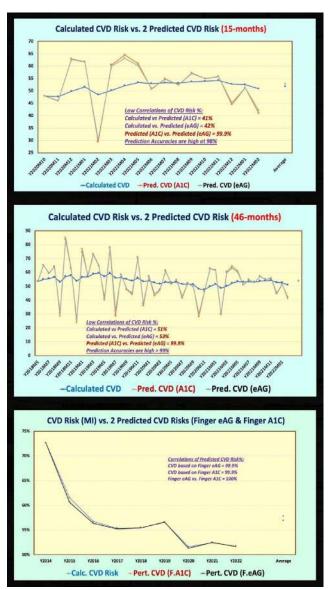


Figure 4: Three daily charts and summarized data table of within 3 time-windows of correlations among CVD risk %, eAG, and A1C %

Figure 5 illustrates three daily charts and a summarized data table within 3 time-windows of correlations among CVD risk %, eAG, and A1C %.

Total Control	CONTROL OF THE PARTY OF T	COMPANY CONTROL OF CON		
Top Color Co	Managing design of the second	Check (F) (Mill) These set in hose left in free left And (Mill) The set in hose left in free left And (Mill) The set in hose left in free left And (Mill) The set in hose left in free left The set in hose left in hose left in hose left The set in hose left in hose left in hose left The set in hose left in		
Correlations:				
Shorter 15 months	Monthly	Daily		
CVD vs. eAG	-31%	-55% -59%		
CVD vs. A1C	-29%			
eAG vs. A1C	70%	97%		
Longer 46 months	Monthly	Daily		
CVD vs. eAG	66%	70%		
CVD vs. A1C	63%	70%		
eAG vs. A1C	89%	99%		
Very Long 9 Years	Annual	Daily		
CVD vs. Fing eAG	93%	80%		
CVD vs. Finger A1C	91%	79%		
are to timber time		99%		

Figure 5: Two Predicted CVD risk % versus calculated CVD risk % using a viscoelastic perturbation model within three time-windows

Conclusion

In conclusion, the following three observations outline the findings from this research work of statistical influences on math-physical medical research projects by selecting three different time-windows of CVD risk datasets:

- (1) From the pre-examination using correlation studies in a time-domain, the smallest data points of 9 (8.2 years with 98-months of information) offers the highest correlations among CVD risk, eAG, and A1C. The shortest time frame of 15-months provides the worst correlations among CVD risk, eAG, and A1C, while the 46-months' time window is situated in the middle. These findings prove that the insight information or behaviors of a biomarker is far more important than the sheer number of data points. Usually, a longer time window of a dataset would reveal more insight into the behavior of its associated biomarker, but this statement is still depending upon case by case in medicine. Generally speaking, the higher positive correlations provide a hint that this longer time-window analysis can usually produce meaningful and useful results.
- (2) In the stress-strain diagram in a space-domain (SD), the three curves have different appearances from each other. This is due to the following two reasons. The first is that the strain information (CVD risk) is different in different time-windows. The second is that its associated viscosity factors i.e., eAG or A1C are also different at various time instant. Nevertheless,

due to the author's continuous improvement on his eAG (higher in Y2014 and then dropping to its lowest in Y2022), and eAG associated A1C, his strain value (CVD risk) is then moving from the right side of the diagram toward the left side. On the stress-scale, both different strain rates and viscosity factors determine the y-scale values. For an overall viewpoint, the 8.2-years' curve shows a complete view from Y2014 to Y2022 with the biggest CVD risk change rate during the sub-period from Y2014 to Y2017. The 46-months' curve reflects a moderate and confined area for CVD risks and their change rates. The 15-months' curve is the most meaningless case due to its insufficient inclusion of biomarker behaviors. It should be noted that these stress scales have been adjusted by using (A1C) and (eAG / 20 or 16) in order to achieve a better viewing of shape and comparison of the stress-strain curves.

(3) Using the viscoelastic perturbation model, a waveform comparison study of the metabolism calculated CVD risk % against two predicted CVD risks, using eAG or A1C, can be done. For both time-window cases of 15-months and 46-months, their negative or moderate correlations are not high enough. As a result, the introduction of a "stronger curve vibrations" effect from the perturbation factors i.e., the "stress element", therefore produces less-meaningful or not so useful CVD risk predictions. On the contrary, the 98-month's case only has 9 data points but with very high correlation results. It can produce highly accurate predicted CVD risks using the visco-perturbation model.

In summary, this particular report shows that if using a shorter 15-month dataset, it would result in unsatisfactory results. If using a moderate 46-month dataset, the CVD risk % would have

higher correlations with both eAG and A1C, but their associated predicted CVD risks are still not quite useful due to its moderate amount of biomarker information. Only using the 8.2 years (9 data points covering a 98-month period) case can reveal a better picture of inter-relations between CVD risk versus both eAG and A1C.

The author attempts to interpret the above findings using biomedical terminology (not math-physical languages) as follows:

Heart problems, such as CVD, do not occur suddenly, and are usually associated with a patient's lifestyle over a longer period of time. The majority of CVD patients (75% to 80%) also reflect diabetes conditions with hyperglycemia (high eAG and high A1C). Therefore, for patients with both heart problems and diabetes, in order to lower their risks of having CVD, a longer period of continuous efforts on lowering their eGA and A1C is necessary.

References

For editing purposes, the majority of the references in this paper, which are self-references, have been removed. Only references from other authors' published sources remain. The bibliography of the author's original self-references can be viewed at www. eclairemd.com.

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