# PROTEIN SYNTHESIS AND "FINE TUNING"

Paradigms for the discrete degrees of Divine influx



Stephen H. Smith, MD

"Proteins are subtle and delicate things..."



## Functions of Proteins

Enzymes

Regulatory

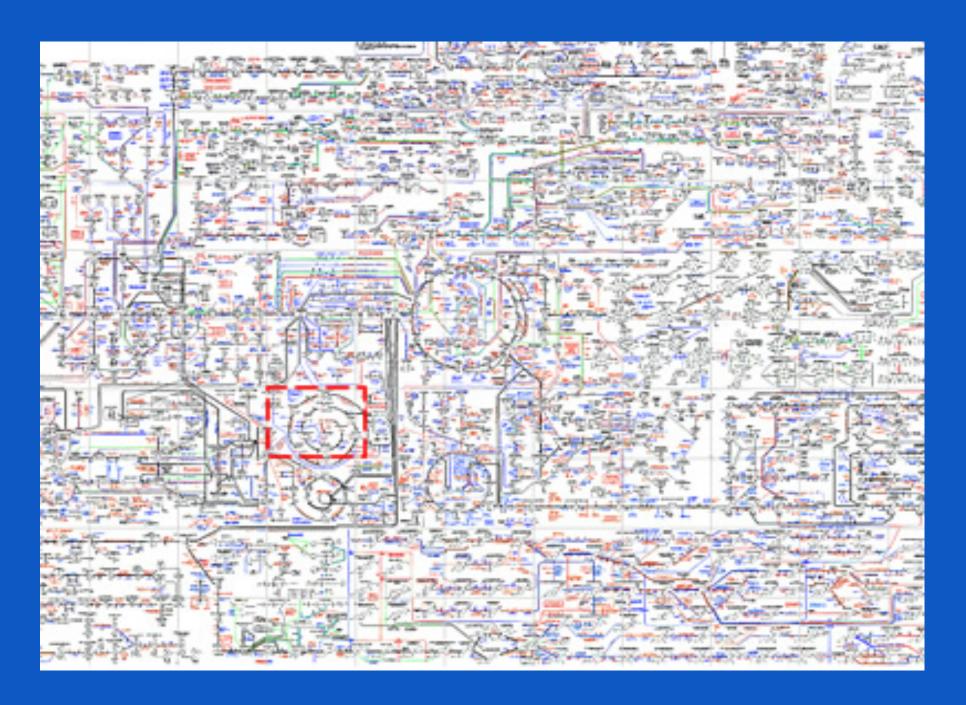
Signaling

Storage

Structural

Immunity

Transport



### Why look at Protein Folding?

- Forces at work operate at the atomic level
- Apparent causes (forces) in nature correspond to spiritual qualities
- Influx "fine tunes" by effects on the electric charge

"Understanding the spiritual cascade of discrete degrees behind the physical degree of life may be a framework that is useful to better understand what is still unknown about protein structure, function and synthesis

Ian Thompson, PhD

## Three Approaches to the Study of Protein Synthesis

- *In Vivo* (within a living cell)
- *In Vitro* (in a test tube))
- In Silico (computational)

## Beginning of Modern Protein Science

- 19<sup>th</sup> century brewing and baking
- Pasteur's discoveries
- Pasteur vs Liebig —— Traube

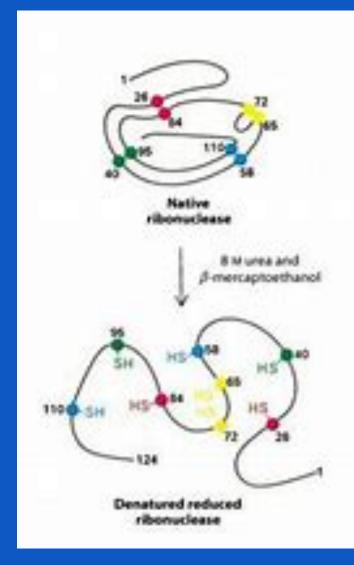
### "Anfinsen's Dogma"

The sequence of amino acids in the primary polypeptide chain determines the final folded tertiary form and, therefore, its function.

## What Determines the Amino Acid Sequence?

 $DNA \longrightarrow mRNA$ 

Ribosome <del>L</del>tRNA



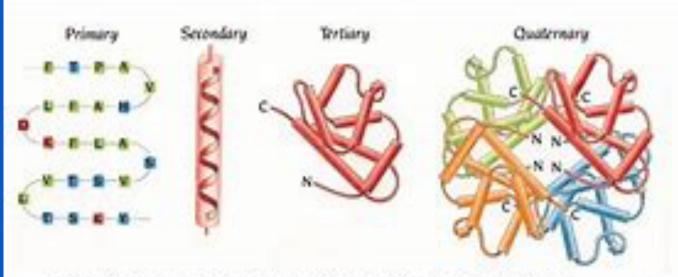
#### Anfinsen Experiment

 Denaturation of ribunuclease A ( 4 disulfide bonds) with 8 M Urea containing βmercaptoethanol to random coil, no activity

## Stages In The Formation Of Proteins

- 1. Primary = polypeptide chain
- 2. Secondary = alpha helix and beta sheets
- 3. Tertiary = "native", active, fully folded form
- 4. Quaternary = combination of two or more discrete polypeptide chains

#### 4 Hierarchies of protein structure

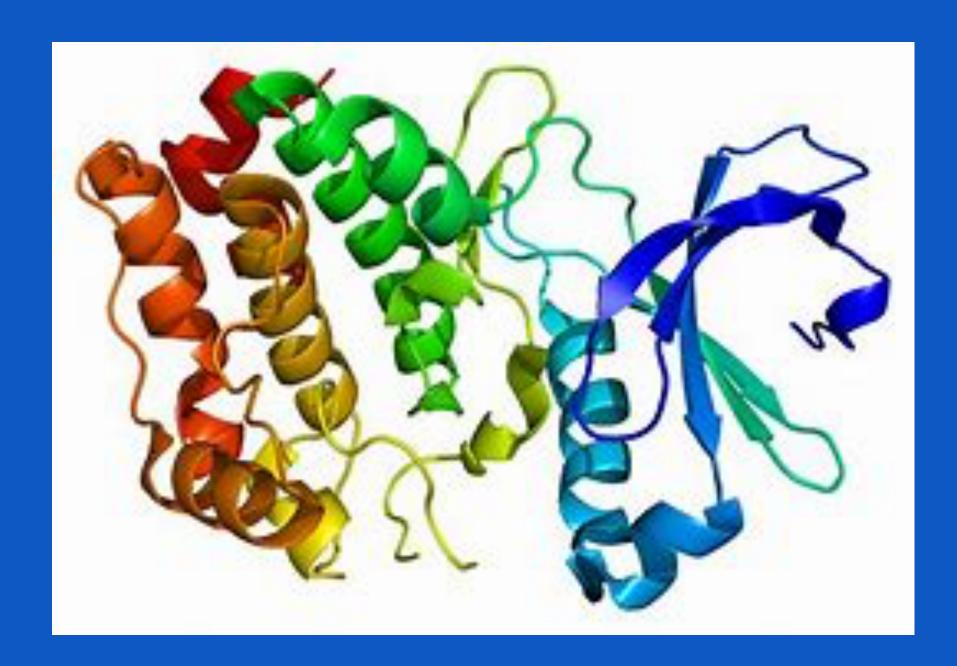


Anfinsen: sequence determines structure



#### Discrete Degrees

The end (goal) is contained within the cause ⇒the cause and the end are contained in the effect.



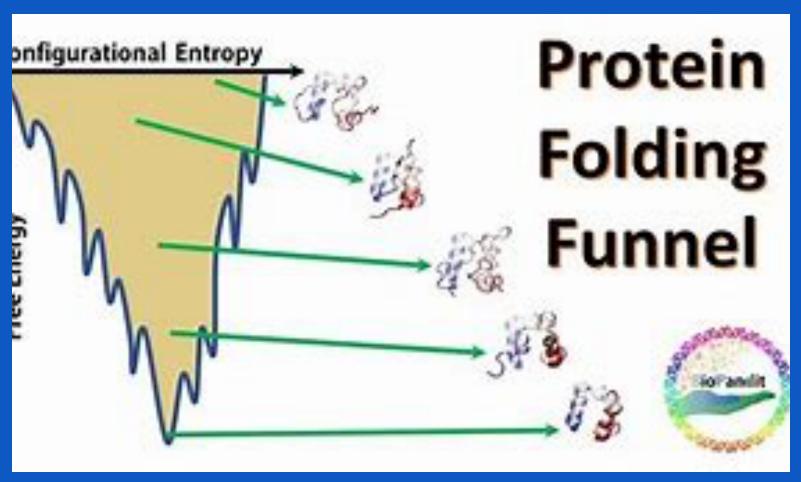
#### "Levinthal's Paradox"

- With only 100 amino acids
- Trial fold once every ten trillionth of a second
- 10<sup>81</sup> years to fold successfully
- Yet protein folding takes only a second!

#### Many Attempts to Solve Levinthal's Paradox

- "Bias" in the pathways? Yet from whence cometh the "bias"?
- "Funnel" energy landscape???

## Landscape "Funnel"



### The Protein Folding Problem

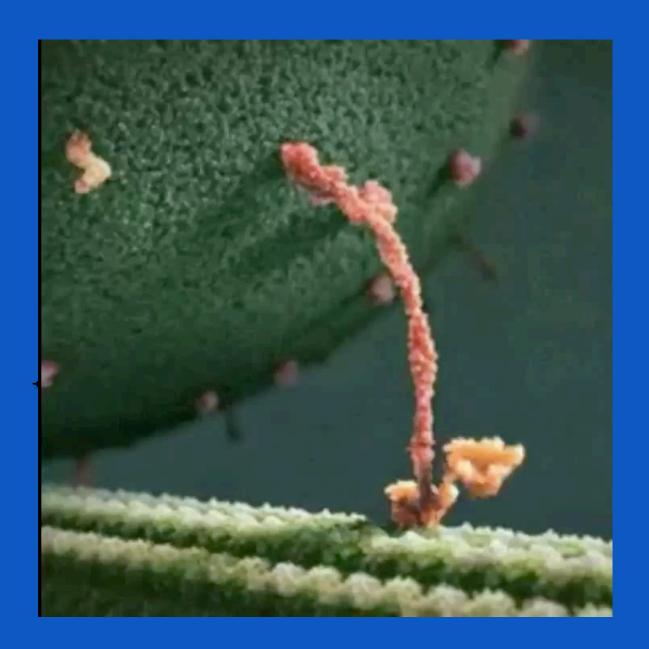
- A new distinct branch of science:
   utilizing chemistry, biology, and physics
   (both classical and quantum physics).
- And now, for the first time theology!

### Key Theological Principle:

Causes are spiritual, they only appear to be from nature!

### Key Theological Principle:

All forms tend toward the human (form).



## Competing Theories of Protein Folding

- Thermodynamics: Gibb's Free Energy
- Place different emphasis on hydrophobic-hydrophilic models
- Research is largely computational and increasingly statistical

#### Protein Folding - from a Theistic Science Perspective

Forces and interactions that fold and stabilize proteins and their possible correspondences!

#### Protein Folding - from a Theistic Science Perspective

Electric charges implicit in the forces and interactions of folding can be fine tuned by immediate Divine influx

# Forces and Correspondences!

Which forces are dominant must be related to use!

#### Hierarchy of Forces

- van der Waals forces
- Hydrogen bonds
- · Ionic bonds (salt bridges)
- Covalent bonds
- Peptide bonds
- Hydrophobic interactions

#### van der Waals interactions

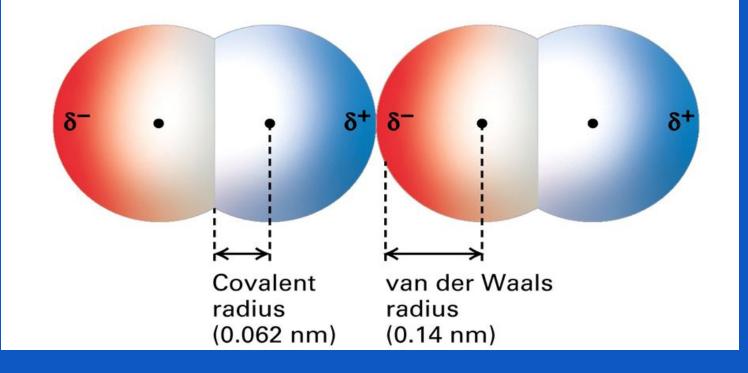
- Atoms moving in close proximity generate a brief, weak attraction
- Almost all the atoms in the interior of the molecule are involved

#### van der Waals interactions

#### van der waals interaction

-caused by transient dipoles, the momentary random fluctuation in the distribution of the electrons of any atoms

- 1/r<sup>6</sup> dependence



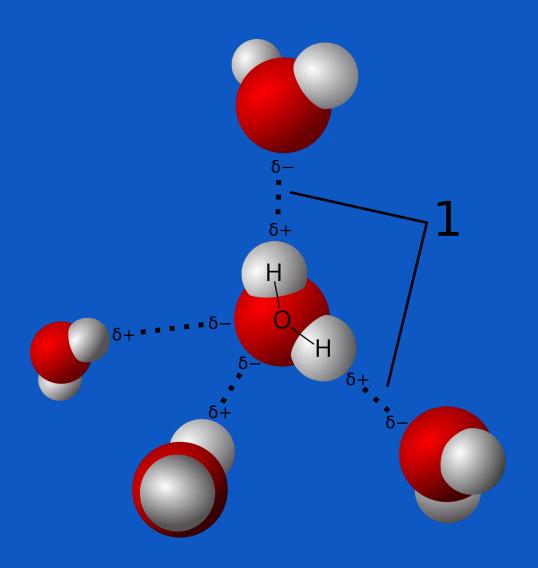
#### van der Waals interactions

- Correspond to "bare facts"?
- Level of "effects" only?
- Corresponds to the sensual/corporeal?

#### Hydrogen Bonds

- Very common in proteins
- 12 of the 20 amino acids form hydrogen bonds
- Asymmetric electron density between closely aligned atoms

### Hydrogen Bonds



### Hydrogen Bonds

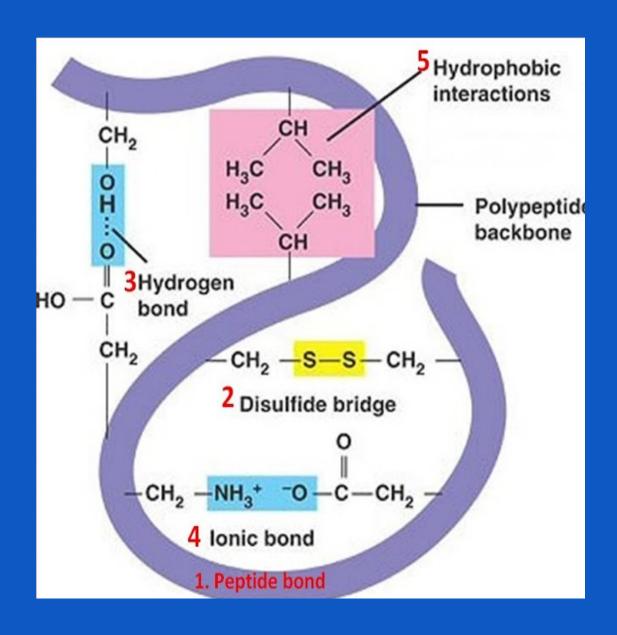
- Corresponds to natural knowledge common to our lives 
   "scientifics"?
- Level of "apparent causes and effects"

#### Salt Bridges (Ionic bonds)

Ion pairing (salt bridging) 

 oppositely charged amino acid
 side-chains are in close spatial
 proximity

#### Ionic Bonds



# Strength of an Ionic Bond:

$$F = q_1 q_2 / d^2 \epsilon^*$$

\* **\varepsilon** "dielectric constant" of the solvent (larger epsilon = smaller attraction!)

# Salt Bridges (Ionic bonds)

 Salt corresponds, in a good sense, to affection for truth

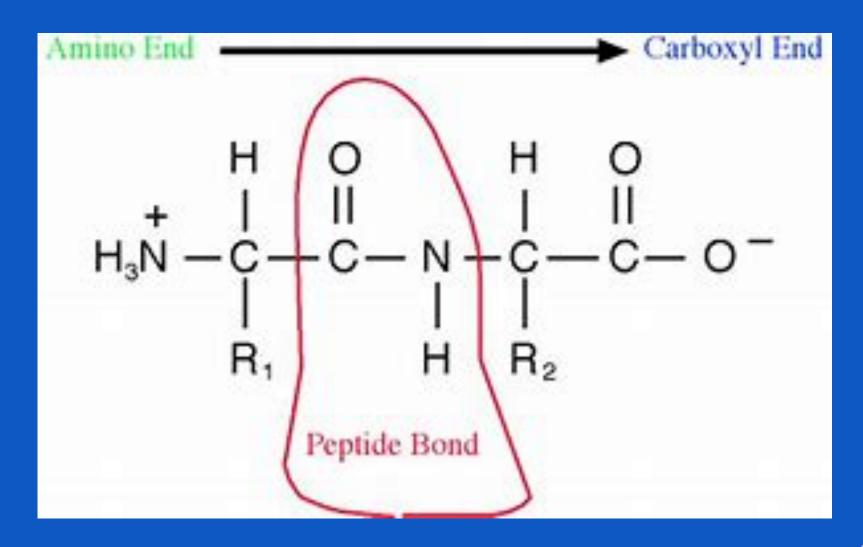
 A salt bridge might therefore correspond to a relationship based on affection for truth.

Now at the level of "cognitions"?

# Peptide Bonds

Form the backbone of protein structure

# Peptide Bonds



#### Peptide Bonds

A tight bond that resists rotation

 Corresponds to the rational; links the natural mind to the spiritual mind in search of truth?

#### Covalent Bonds

Share electron pairs between atoms

#### Covalent Bonds

 Correspondence: a marriage: the bond of good and truth?

# Hydrophobic Interactions

Hydrophobic amino acid residues are found mainly in the core of the functional (native state) protein!

# Hydrophobic Interactions

- Hydrophobic lipid (hydrocarbons)
   corresponds to love (the will)!
- Like the hydrophobic amino acids in the folded state, love is interior!
- Hydrophobic interactions with water are primary drivers in folding!

# Hydrophobic Amino Acids and Water

 Quantum physics of water key to understanding the folding mechanism!

The dielectric constant "ε" of the solvent and the interior environment = an opportunity for "fine tuning"!

#### New Horizons!

- Long range interactions
- Solitons
- The cell as an "electrome"

"Everything in the universe that is in harmony with God's plan relates to goodness and truth."

New Jerusalem and its Heavenly Doctrine 11

#### Arcana Coelestia 3629

"Also, they do not know what correspondence is and what influx is, and that when the spiritual flows into the organic forms belonging to the body, it establishes the living workings such as occur there. Nor do they know that not even the smallest parts of the body can

# Chaperone Proteins

AKA "heat shock" or "stress" proteins

# Chaperone Proteins

Correspondence to celestial angels, spiritual angels and good spirits?

# "Behold I Make All Things New"

An additional role of chaperone protein molecules!



#### Chaperone Proteins

- May preserve a totally new protein for some future use!
- How does it recognize it as potentially useful?

#### Arcana Coelestia 5131

"... To enable the cause to exist the end must act on the level where the cause belongs, calling on assistant means to help it –the endto bring the cause into existence, and to enable the effect to exist the cause likewise must act on the level where the effect belongs, by calling on assistant..."

#### Arcana Coelestia 4523

"...Thus the whole body is an organ composed of the deepest arcana belonging to everything which exists within the natural world, and its formation is determined by hidden forces by which all things act and the wonderful manner in which they flow."

End - Cause - Effect



Transient, small fluctuations in constants of nature

"The existence of life as we know it depends delicately on many seemingly fortuitous features of the laws of physics and the structure of the universe."

Paul Davies, The Mind Of God

"There are three types of change to consider. Tiny infinitesimal changes are possible. If we change the value of the fine structure constant only in the 20<sup>th</sup> decimal place there will be no bad consequences for life...if we change it by a very small amount, say the 2<sup>nd</sup> decimal place, then the changes become more significant. Continued on next slide

Continued from last slide

Properties of atoms are altered and complicated processes like protein folding or DNA replication may be adversely affected."

John Barrow *The Constants of Nature* 

#### Fine Structure Constant

$$lpha = rac{1}{4\piarepsilon_0}rac{e^2}{\hbar c} = rac{\mu_0}{4\pi}rac{e^2c}{\hbar} = rac{k_{
m e}e^2}{\hbar c} = rac{c\mu_0}{2R_{
m K}} = rac{e^2}{4\pi}rac{Z_0}{\hbar}$$

#### where:

- e is the elementary charge (:= 1.602 176 634 × 10<sup>-19</sup> C);
- π is the mathematical constant pi;
- $\hbar = \frac{h}{2\pi}$  is the reduced Planck constant (:=  $\frac{6.626\ 070\ 15 \times 10^{-34}\ J\cdot s}{2\pi}$ );
- c is the speed of light in vacuum (:= 299 792 458 m/s);
- ε<sub>0</sub> is the electric constant or permittivity of free space;
- μ<sub>0</sub> is the magnetic constant or permeability of free space;
- k<sub>e</sub> is the Coulomb constant;

#### Arcana Coelestia 4524

"And because there must be a continuous connection with the spiritual world in order that every single thing may be kept in being, or constantly come into being, it follows that the purer or more interior things within the natural order, and consequently within the human being, spring from that world, and