### Moles, DHEA, Etc.

Everyone knows what a mole is, but no one understands what a mole really is. Moles are interesting because they are simultaneously normal and a little abnormal. But it seems that no one has taken them seriously enough to study them very scientifically. At least, I don't know of any publications on "the epidemiology of moles," or "the experimental production of moles."

Advertisements from the cancer society fundraisers have taught people that "any change" in a mole could be a sign of cancer. The idea seems reasonable enough, but I think it has no clear scientific basis The fear of cancer has caused many people to have things excised as soon as they are noticed, but during this generation in which people have learned to have things cut out promptly, the incidence of melanoma has risen sharply. Melanoma first appears in normal- looking skin about as often as it does in pigmented moles, so the "preventive" removal of all moles is no guarantee that you won't get melanoma.

At least some pigment cells are "amoeboid," meaning that they can creep around. At body temperature, white blood cells under the microscope can walk about two inches per day.(1) These facts indicate that it could be seriously misleading to think of a bit of tissue, such as a mole, as a "strictly localized" thing, the removal of which has no meaning for the rest of the body.

I will just briefly mention some of the observations that have made me think of moles as "systemic phenomena."

A friend had many protuberant moles that had been present for most of her life (if not since birth), and one summer they withered, dried up, and fell off. She was taking a little more thyroid and triiodothyronine that summer. One hot summer, I had a four ounce glass of Modelo beer, discovered I liked it, and for the rest of the summer I would often drink a similar small amount during he hot part of the day, while at my friend's house to water the plants. Two moles appeared on my forehead, and grew rapidly, until the bigger one was the size of a small pea. When the beer in the refrigerator was gone, the summer was nearly over, and I had thought increasingly about the estrogen content of beer, so I didn't drink any more. Within about a week, the moles were shrinking. The smaller one disappeared while the other one was still about the size of match-head. After about three weeks, the remaining mole was about the size of a dried pea, and one morning it rolled free when I touched it. It had become firm and dry.

# **DHEA Changes Moles**

In 1980, a familiar old mole on my belly had begun growing and darkening. A couple of times when I had my shirt off on a hot day physician friends looked worried when they saw it and told me I'd better have it removed. It became thoroughly black, took on a sharply triangular shape, and had a firm leathery texture. By 1981, it was about 15mm. long, but still fairly flat. I had been experimenting with oral doses of DHEA in oil for two or three weeks, and had noticed that it made me feel like an adolescent, but it was causing me to get pimples.

One night as I was going to bed, I looked down and was shocked to see what looked like a maraschino cherry on my belly. I spontaneously flicked at it with my finger, thinking it might be an engorged tick, though the color and gloss weren't appropriate. Just before the second flick brushed it, I too late saw something black on the top of the globe, and realized that it was my mole. The black material brushed away, leaving a small drop of blood, but the dome remained glossy and red. By the next day, it was smaller and was no longer shiny from internal pressure. That afternoon I photographed it, and there was a red flare surrounding it, about 2 inches in diameter. By the third day, I could see that the deflating thing was going to be a brown mole, of roughly the same size as the original one that had been there a few years earlier. The only difference from its earlier state is that its margin might be more diffuse now.

In his mid-50s, my brother mentioned that he was getting a molelike spot on his forehead. When I told him about my experience, and that I thought oral DHEA might have been responsible for the mole's disappearance, he applied a concentrated solution of DHEA in vitamin E directly onto the mole. A day or two later, he noticed a spot of blood on his pillow, and had his doctor cut the thing off, because of fear that bleeding signalled a malignant change.

After I found that a copper solution quickly caused my white eyebrows to return to their normal color, and repeated the experiment several times over about 3 years, I tried the same solution on my sideburns, and saw that some of those hairs had darkened after one application. It wasn't convenient to observe those hairs, so I decided to apply the solution to some white whiskers on my chin. After two or three applications, I carelessly used a solution that was too strong. Within less than a minute I felt it stinging, and washed it off. A stinging sensation was noticeable during

the next 2 or 3 hours, and I washed the spot thoroughly several times. About 4 hours later it was still a little uncomfortable, and when I looked at it in a mirror I was horrified to see what appeared to be a large, coffee-colored mole on my chin, in the area where I had applied the copper, and where the burning sensation persisted. The edges were slightly raised about the surrounding skin, and it looked li ke a normal light brown mole, roughly the size of a nickel. It was so hard for me to believe that such an anatomical change could have occurred within 4 hours that I wondered if I had just failed to observe something that had been developing gradually. But after about a week it had disappeared, and no trace of it has returned, so I had to conclude that the high concentration of copper had acted with, or in place of, tyrosinase to form new pigment, or that copper-hungry cells had invaded the area.

A couple of years ago I noticed some moles on my upper chest were growing, darkening and developing an irregular, granular form. I hadn't used any DHEA for a few years, just because I had lost interest in it. A friend had just been telling me about using topical progesterone and DHEA in vitamins E and A to shrink a large and growing mole on her leg. For a few weeks I watched carefully as my moles changed since their location made it easy for me to view them at very close range. After becoming nearly black, I noticed that they were fading and becoming sort of grey. Then white and creamy areas appeared, and the grey seemed to concentrate into a few blue spots. The changes were surprisingly fast. I would look at them 2 or 3 times a day, and the blue, or black, or brown spots would be in entirely different locations each time. The moles had become firm to touch, but the spots of pigmented cells appeared to be swimming wildly around within the boundaries of the moles, as if looking f or something, or escaping from something. I smeared a very light film of vitamins A and E with DHEA over the area of my chest with the moles, and the next time I looked the color was becoming regular. Within a day or two, even the texture had returned to normal.

#### **Higher Altitudes Decreases Cancer Incidence**

About ten years ago, a cancer geographer mentioned that he had found an inverse relation between altitude and the incidence of cancer around the world. A woman from Texas who was at the conference was sure his information was wrong, because the higher intensity of "radiation" at high altitudes would increase the damage to DNA that is commonly supposed to cause cancer. She decided to disprove his work by doing a study limited to the incidence of melanoma - which is supposed to be especially sensitive to ultraviolet radiation - within Texas. Her results surprised her; even melanoma incidence was lower at higher elevations.

Much of the doctrine about radiation at high altitude has been created by and for the nuclear industry. Actually, the dangerous secondary and tertiary particle showers from cosmic rays are more intense at sea-level than at high altitudes. But the greater oxygen pressure at sea-level might be the most important factor in various kinds of cancer, and other disease, since our antioxidant system can probably maintain better balance at high altitude. Mitochondria become more numerous at high altitude, and this might result from increased durability, as well as from the need for more efficient use of oxygen.

The skin is a major source of sterol formation. Skin changes in aging might indicate a special susceptibility to sterol deficiency, which could make topical application of steroid-associated materials especially appropriate.

About 50 years ago, melanosomes were believed to develop from mitochondria. Whether that was accurate or not, there are some interesting parallels between respiration and pigment metabolism. Copper, vitamin A, pregnenolone, progesterone, vitamin E and DHEA are intimately involved with both. The glucocorticoids tend to antagonize both.

## **Pigment Cells Related to Nerve Cells**

Leghorn chickens are sometimes thought of as genetic albinos, but when relieved of the stress of egg laying, they can form black feather pigment. Domesticated animals of various kinds tend to become white. Soviet fox breeders selected foxes for tameness for 20 years. The foxes bred for tameness also lost their wild coloring, and became spotted. The migration of pigment cells through the tissues of the developing embryo reflects in some way the altered balance of the whole organism. It is this distribution of cells during embryonic and fetal development, and later in life, which produces "birth marks" and moles. Pigment cells and nerve cells are closely related in embryonic development, so it is interesting to notice the similar appearance of astrocytes n the brain and melanocytes in the skin. Macrophages sometimes have a similar structure, and when stained with metals reticular macrophages and astrocytes (both have been called "metalophils") look surprisingly similar.

Metchnikov developed this theory of inflammation and immunity by phagocytosis in connection with his attempt to explain development, how ontogeny reflects - but alters - phylogeny. The recognition and preservation of self, in the immune reactions, was just an aspect of the ontogenetic creation of individuality.

When my phagocytes eat up one of my moles, and a good copy of my "original" mole is reconstituted in the same location, Metchnikov's idea of a dynamic organismic integrity, in which the phagocytes play a central developmental role, takes on new concreteness for me.

In 1983 I noticed that the whiskers on my cheeks were going white in circular as if the melanocytes were migrating centrifugally. Around the same time I noticed a flat mole, or raised freckle, was growing on my temple. Although many people have those spots removed, it is very common for people in their 80s to have a button-like black spot on the temple. My spot waxed and waned, but on average tended to get bigger and darker with the years.

In Mexico recently, a friend found that applying DHEA in vitamin E to a very big, red keloid tumor caused it to quickly fade and shrink. As I was driving home, I applied a little of the same solution to one of the mole on my temple. A day or two later I

noticed a concentric ring of brown skin on the opposite side of the mole, about a centimeter away. The next day, there were two such bands. Then a bright red line appeared at the front margin of the mole, where I had first applied the oil. Although it looked as though the edge of the mole had bled, it was actually just a thread-like region of extremely dilated capillaries. Day by day, the thread of red appeared to be

contracting around the mole. The brown rings took on a V-shape, and extended into the hair. The color of the mole became lighter for about a week, and as the red thread "contracted" the flat mole became pedunculated and mostly colorless, like a tiny mushroom. Two or three dots of color remained, apparently trapped, in the globe. After drying and shrinking for a couple of days, it fell off, and the point of attachment had become smooth, normal skin within a day.

In this case, the concentric rings of brown pigment suggested that pigment cells might be returning to their normal locations, after having lived for ten years in an island on my temple. This might suggest that this particular skin area is the last to lose its ability to produce DHEA, and that well-nourished cells can remember where they belong.

## Lipofuscin Produced in Oxygen Deprivation

Age pigment, lipofuscin, is produced in oxygen deprivation, apparently from reduced iron which attacks unsaturated fats. It has its own "respiratory" activity, acting as an NADH-oxidase. Melanin is produced by polymerization of amino acids, with copper as the catalyst. With aging, iron tends to replace copper. Melanin is an antioxidant. Thus, there is a sort of reciprocal relationship between the two types of pigment. A vitamin E deficiency relative to consumption of polyunsaturated fats, and an estrogen excess, accelerate the formation of lipofuscin. A 47 year-old woman who had only a few "liver spots" on the backs of her hands began taking large amounts of estrogen, and within a few months the brown spots had darkened and spread until most of her skin was covered with spots. When she stopped using estrogen, and applied progesterone topically, the spots disappeared.

Since both keratinocytes ("keratinophages") and "melanophages" can phagocytize melanin granules,( 2) the occasional rapid disappearance of "age pigment" makes me wonder whether pigment-loving cells can't ingest and destroy lipofuscin. To the extent that lipofuscin wastes oxygen, the phagocytes might be able to relieve some of the symptoms of oxygen deficiency in aging. (My recent experience with an 82 year-old man with emphysema who regained a normal pink color a few days after beginning to use DHEA and pregnenolone led me to wonder whether the improvement was mainly in the lungs, or in the other tissues.)

If leukocytes can direct organized attacks on moles, with reconstruction of healthier tissue at the same location, the healing of an infection, or a wound, should be seen in the context of growth, construction, and creation, instead of simply attack upon a foreign body.

If leukocytes remove age pigments and deteriorated moles by recognition of an "age antigen," as in their removal of old red blood cells, or the unnecessary tadpole tail in maturing frogs, then we have a way to begin to grasp the mechanism of ontogenic evolution through various stages and "recapitulations of phylogeny," rejecting one structure as another step forward becomes possible. (The fact that the senescence antigen appears in embryonic kidney cells, as well as in adult liver cells, indicates that it has to do with development, the transition from one stage to another, rather than simply with the "wearing out" of cells.) Melanoma sometimes spontaneously regresses, in a self-digesting manner.( 3) This might be the process that I have seen in my "autolytic" moles. The fact that the "tumor necrosis factor" is also a cachexia inducing factor (and is produced by activated macrophages) provides another perspective on the idea that the whole body might be designed to periodically melt-down, to be systematically reconstructed anew.

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