



THE SPINE OWNER'S MANUAL

HOW TO AVOID BACK PAIN & LIFE ALTERING SURGERY

1st Edition | Christopher J. Centeno, MD

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Introduction

First, this book and its big brother (Orthopedics 2.0) is part of a two-decade quest to find what I have termed the <u>"Unified Field Theory."</u> Physicists have long sought a single "theory of everything" that ties together all other theories into one grand explanation of the universe. I've sought to do the same for the spine and the musculoskeletal system. Many theories abound about how to diagnose and treat spinal problems. Orthopedic surgeons have a surgical approach, family-practi ce and sports-medicine practitioners another conservative approach, chiropractors their own alternative approach, and physical therapists yet another. Within chiropractic care, physical therapy, and alternative medicine, there are literally hundreds of wholly different theories about what's wrong with the spine and how best to address these problems. Having studied many of these, I always found a kernel of truth and some interconnection between them. As the research in this area has become more robust in the past 20 years, many of these concepts can now be vetted by scientific observation rather than intuitive guesses.

This book contains my own theories based on several decades of reading and observation and offers a way to organize that information for both doctors and patients. The reader should note that while many of the components of this new theory are supported by rigorous scientific research, the whole package as I present it hasn't been studied using what doctors call <u>"level-I evidence."</u> This type of medical evidence means that randomized controlled trials have been performed and the treatment approach has been found to be effective. Having said that, most of what we do today for patients with spinal problems is not supported by level-I evidence. This includes all spinal surgeries (including fusion, laminectomy, and discectomy), chiropractic adjustments, acupuncture, massage, and most all physical therapy. These surgical and nonsurgical approaches all lack the type of rigorous scientific support (level-I evidence) that shows they are effective. In fact, when some of these procedures have been studied in controlled trials, they have often been shown to be no better than placebo surgery or no surgery. (Fusion adds nothing to the outcome of a laminectomy other than more complications.)

This is an Internet book that instead of being about fifty to one hundred pages long is really several thousand pages long. How is that accomplished? The science behind many of these concepts would create an unwieldy publication that would be too difficult for patients to read and follow. By publishing this book on the Internet, I can easily hyperlink to scientific abstracts and other references so the reader can delve deeper into any subject or simply read the basic explanations. In addition, my goal is to allow patients to submit questions and feedback so the book can be updated and improved. The Internet allows this kind of flexibility; hence, the version designation in the title (starting with version 1.0 and adding to that with each minor or major change). As the book is updated on the Internet, the version number will allow the reader to know if he or she has the



most current information. Making this book better is as simple as clicking a link. <u>To submit</u> <u>questions or ask for clarification on any part of this book, send an e-mail to the author by</u> <u>clicking here.</u>

What is Orthopedics 2.0? In particular, Orthopedics 2.0 doesn't refer to the discipline of orthopedic surgery or its successor. While orthopedic surgery may well be used as a part of Orthopedics 2.0, Orthopedics 2.0 has a bigger focus beyond just fixing one part of the spine. We focus on each part of the spine as it is interconnected to the whole. While the focus of this book is nonsurgical, there will always be situations where the best approach is surgical. What will likely occur over the next one to two decades is a slow and steady movement toward less-invasive orthopedic-type procedures. What we call "interventional orthopedics." This is identical to what's occurred in other areas of medicine, such as cardiology with fewer more-invasive open-heart surgeries and more X-ray-guided catheter procedures.

Interventional orthopedics represents the shift from spine salvage to repair. When the focus shifts to repair, the amount one needs to know increases exponentially. The pyramid at the beginning of the introduction outlines what we use to evaluate the spine. While I use stem cells in daily practice, it's important to note that helping patients is often not as simple as injecting magic stem cells. This book details the system our clinic uses to decide which procedures and therapies to apply.

The problem with repairing the spine system is its complexity. Think about your car. You know there are critical components to keep it running. The wheels have to be aligned or the car won't go straight and the tires will wear unevenly. The connections between the wheels, the axle, the drive shaft, and the engine have to be flexible and allow fluid movement. The engine, as it turns the drive shaft, has to be well oiled. As the engine cranks up to even faster speeds, the connections had better be stable or the whole thing will fly apart. Finally, your engine has miles of wiring and small computers on board to monitor the whole thing and to regulate the activity of the engine, brakes, gasoline usage, and monitoring systems.

Now think about your spine and its component parts, like discs, facet joints, ligaments, spinal nerves, and muscles. The same principles of alignment, good joint connections, stability, and sound wiring (nerves and minicomputers that impact everything from the timing of muscle firing to the information about joint position) apply. Regrettably, our surgical approach, to date, has too often just focused on bringing the car into the shop to replace a few worn parts or shave them down so they fit a little better, but it has not considered how the parts got that way. Let's look at that analogy now as it applies to a person. If a 40- or 50-something-year-old patient who runs every day is suddenly diagnosed with arthritis in the spine, could it be that for years the low-back joints were



getting worn down due to poor alignment? We'd all accept this premise at face value: a misaligned front wheel and axle could cause the right front tire to wear faster than the left. Yet, for some reason, our medical-care system often ignores why one joint wore out faster. The reason: if the plan is to fuse the joint, who cares? What if we wanted to save the joint? Would it matter more? Absolutely! This is the reason for the Ortho 2.0 approach and this book. When the shift is moved from replacement to repair, it matters how the joint got that way, if the joint is stable, if the surrounding muscles are firing correctly to protect the joint, whether the alignment is correct to support a healthy joint, and if the wiring is in order.

While stem cells are a great advance and represent a cutting-edge tool, their use without considering all of these other things doesn't get patients where they want to be, which is having a spine they can count on for many years to come. In this book, we'll look at all of the parts of the Orthopedics 2.0 paradigm listed above or, as my partner coined the term, SANS. This stands for stability, articulation, neuromuscular, and symmetry. Sans in Latin means "without"; hence, the focus of this book is to leave you sans pain.



Why Traditional Injection- Based and Surgically Based Spine Care Needs a Serious Upgrade!

If you're reading this book, you've likely been offered traditional injection-based or surgical care for your spine. You may have even tried some injections or had a surgery or two and weren't satisfied with the results. Maybe you're just exploring all your options before you pull the trigger. Let's explore why all of these options have big issues.

Before we get into how the spine is structured in more detail, I'll review it here a bit for the purposes of this discussion. The spine bones are called vertebrae and there are 5 in your low back, 12 in your upper back, and 7 in your neck. These are numbered from one at the top and increase in number as you go down the spine. Each area has its own abbreviation, with "C" meaning cervical (neck), "T" meaning thoracic (upper back), "L" meaning lumbar (low back), and "S" meaning sacral (tailbone).



There are discs between the bones that act as shock absorbers, and each of these is given a name based on the bone above and below. So L5–S1 means that this is the disc between the L5 bone and the top bone of the sacrum (S1). Each level also has two joints at the back called facet joints, and these are named the same way. There is also a spinal nerve that exits at each level, and these are named for a single nerve (like the "L5 nerve," which comes out between L5 and S1).

Pain in the neck or back can be caused when a spinal nerve is pinched by a disc bulge or a bone spur. The spinal nerve can also be irritated by either chemicals leaking out of the disc or sloppy spine bones due to loose ligaments or due to bad spinal stabilizing muscles. Pain can also come from injured or arthritic facet joints. Finally, pain can be caused by irritated muscles (called trigger points).



Now that we have that out of the way, let's review what's not quite right about what doctors have offered you for spinal pain:

• **Narcotics:** These powerful medications can help with pain, and thanks to aggressive physician marketing by big pharma, we have a real epidemic on our hands. These drugs went from being barely prescribed to being routinely handed out in ever-increasing doses to patients with back and neck pain. The problem? <u>Recent research shows that they create</u> more pain in the long run by rewiring your central nervous system.

• **Epidural Steroid Injections** (a.k.a. epidural injections, ESI): This is the single most common injection for back and neck pain in the United States and elsewhere. The injection is performed to reduce swelling around a spinal nerve due to a bulging or herniated disc or pressure on a nerve due to bone spurs. The issue is that while the high-dose steroids being used are potent anti-inflammatories, this medication type causes the local cells to become dysfunctional and die off. There's also the little problem of these medications markedly increasing the risk of fractures in older women.

• **Facet Injections** (a.k.a. z-joint injections, medial branch blocks, MBBs): This is the second most common injection for back and neck pain. The procedure is performed to reduce pain and swelling in the joints (facets) at the back of the spine. A strong anti-inflammatory and an anesthetic are injected under X-ray or sometimes ultrasound guidance. What could go wrong? The strong steroid anti-inflammatory is a steroid that can kill off cartilage cells. The most commonly used anesthetic is called bupivacaine, which is also toxic to cartilage cells. In fact, add the two of these together and you get 1+1=3, or a synergistic toxic effect.

• **Radiofrequency Ablation** (a.k.a. RFA, RF, rhizotomy, radiofrequency lesioning): When facet injections provide only temporary relief, most physicians will next perform a radiofrequency procedure, which involves burning away the nerves to eliminate the feeling of pain caused by the injured or degenerated joints. The problem is that in the low back, these nerves also control the stability muscles that protect the spine. So killing the nerves kills these critical muscles. In the neck, getting rid of the nerves also gets rid of important feedback and position sensors in the joint, which can lead to more problems in the long run.

• **Botox for Neck Pain or Headaches:** The idea behind Botox seems simple enough. You have really tight muscles that cause your headaches and neck pain; hence, injecting something to relax those muscles for a few months is a good solution. However, there's a reason these muscles are tight all the time—the deep stability muscles that are supposed to keep everything aligned are off-line, so the big outer muscles take over a job they weren't designed to handle. So injecting Botox (or its cousins Myobloc or Dysport) will just weaken these muscles that are trying to hold your head up. So while this may work in the short run, in the long run, you're left with weakened muscles where stronger ones are needed.



• **Back or Neck Disc Surgery:** It's hard to believe that we've known for two decades that surgery to get rid of a bulging disc is no better than going without the surgery. However, this procedure is still routinely performed. The problem is that cutting out a piece of the disc that's bulging out and pressing on a nerve leaves the disc weaker and more likely to herniate again. While a few patients who get this surgery really need it, don't opt for this surgery just because the above injections can't help your pain, as more-advanced solutions will be discussed in this book.

• **Back or Neck Fusion Surgery:** One way for surgeons to dramatically increase reimbursement for a surgery is to add a fusion, which is where the bones in the spine are bolted together. Does this work to improve outcomes over just cutting away bone spurs? No. <u>A recent high-level study shows that all it does is add complications.</u> Despite that, because of the increased payout for the surgeon, fusion is fast becoming the most common spine surgery performed.

• **SI Joint Fusion Surgery:** One of the latest trends is "minimally invasive" fusion of the sacroiliac (SI) joint for patients with chronic pain of the SI joint. Using X-ray guidance, the surgeon places a few screws in the area, the goal of which is to lock the joint solid. The problem is that fusing any joint causes the forces to go elsewhere—in this case to the low back and hip. <u>Read my review of the research on this procedure.</u>

The days of receiving injections of toxic substances and invasive surgery are now over for the vast majority of patients with spinal problems. Read on to find better ways to help. In the meantime, it's important that you learn how your spine works by understanding about stability, the discs and joints, the nerves and muscles, and the symmetry of it all.

Interventional Orthopedics and Its Approach in the Spine

Let's take a look at a movement in medicine that builds off of the SANS system as a whole (covered extensively in my book <u>Orthopedics 2.0</u>)—interventional orthopedics. This approach is different from traditional surgical orthopedics. So let's look at some of those differences as they relate to the spine.

Disc Herniation: The surgical approach is to remove part of the back wall of the disc, which makes it predictably weaker. The interventional orthopedics approach? Use our fourth-generation platelet lysate (growth factors) to help accelerate the body's ability to clean up the herniation and to improve blood flow and reduce swelling around the nerve. Disc Degeneration (Wear and Tear): The surgical approach is to fuse together the degenerative levels of the spine, unsurprisingly leading to more overload above and below the fusion and damage to these levels (called adjacent segment disease, or ASD). The interventional orthopedics approach? Use different types of platelet-based injections or



stem cells to help tighten ligaments and get muscles to fire to better stabilize the sloppy segments, reduce nerve swelling, improve blood supply, and help painful arthritic facet joints.

Low-Back Disc Bulge: The surgical approach is to cut out the bulge that's pressing on a spinal nerve and causing sciatica. The problem? Discs don't heal. <u>Cutting out the bulge is like sanding down a bump on a bike tire, weakening the outer rubber covering and making it more likely to fail.</u> The interventional orthopedics approach? <u>Inject platelet growth factors into the area to improve the blood supply around the disc and reduce the swelling.</u> If this doesn't work, <u>inject specially prepared and cultured stem cells into the disc bulge using a novel device to strengthen the weakened and bulged disc</u>, hopefully getting rid of the bulge.

Multifidus Muscle Pain or Atrophy: This means that the muscles that are designed to stabilize the spine and keep it adjusted to millimeter precision at all times have gotten weak and feeble. Since there's really no surgery for <u>multifidus muscle pain</u> or atrophy, the typical surgical approach is to look for discs and bones pressing on nerves around these spinal muscles and cut those out. In addition, surgery generally destroys the muscle, making the spine more unstable. The interventional orthopedics approach? Inject platelet lysate and <u>superconcentrated platelet rich plasma precisely with ultrasound guidance into the multifidus muscle in the spine.</u> This procedure utilizes the growth factors from the patient's own blood platelets to reduce nerve irritation and helps clean up any damage. Combining that with special exercises should get this critical stabilizer back on track!

Spinal Stenosis: This means that arthritis is putting pressure on spinal nerves. The surgical approach is to cut out bone, ligament, and muscle—large sections of the spine—leaving it open but unstable. <u>A fusion is often added, but the recent research shows that all this really adds is complications in that it fails to make the procedure better.</u> The interventional orthopedics approach? Tighten down the supraspinous/interspinous ligament that's connected to the ligament that often buckles into the spinal canal causing nerve compression, and use injections to further stabilize the area and get rid of nerve and joint pain.

Disc Tears and Pain: The surgical approach is to fuse the disc, with all of the problems noted above. The interventional orthopedics approach? Inject stem cells into the disc to help heal the painful tear.



Introduction to SANS



This book is designed so that if you read the next few pages, you'll have the gist of the message. The book is also rich in infographics—diagrams that allow you to use pictures and text to better understand the content. Finally, the pages that follow this section describe the concepts in more detail. So if you're really busy, start here and end here for now. Read the rest later.

The SANS concept breaks your pain problem into its root causes in four areas: stability, articulation (joint), neuromuscular, and symmetry.

If you have chronic spinal pain of almost any type, from spinal arthritis to nerve pain and sciatica, you will be forever at its mercy if you don't understand its four parts. If you understand these, however, you can take control. So pay attention!

Stability: Your spine is made up of individual segments that stack on top of each other like kids' building blocks. This is by its very nature an unstable mess. You have all sorts of systems designed to keep it all from falling apart and being sloppy.

The spine joints may have a small amount of extra motion that is literally slowly destroying them. The real shocker is that many highly trained physicians and surgeons will likely never



tell you about this instability, nor do many understand it themselves. More concerning is that it can generally be fixed with a few simple injections or exercises.



Stability means a joint that moves with the surfaces in general alignment, all the time. Ligaments are the living "duct tape" that prevents the spine from catastrophic failure. Your body also needs to make sure that the parts of the spine don't give too much when you stress it. Examples include when you walk fast, run, cut (the rapid lateral moves common in sports and exercise), or lift weights. On the other hand, muscles provide the slight adjustments that keep the joint surfaces in very precise alignment as you move. In the spine, the deep multifidus muscles fire in a symphony of movement with millisecond timing and micrometer precision to make all of this happen. How do you know if you have a problem with instability? You may or may not feel your spine popping and cracking—this tends to happen when the instability is moderate to severe. Instead you may have pain or stiffness after activity. If you don't feel these things, your spine may be more stable, but it may have more subtle problems that lead to a more rapid progression of arthritis. Click on the infographic labeled "Stability" to learn more about the general concepts.

Articulation: This is a fancy word for joint. Your spine joints (also called facet joints) allow movement in very specific ways. These small joints in your spine have a different pattern of possible motion from your shoulder, which is still different from your ankle. A joint



is generally made of two bones that come together and are cushioned by cartilage. A joint is also surrounded by a tough leathery covering called a joint capsule and is further reinforced by ligaments. If you want to stay active as you age, you need to learn more about the status of your spine joints, as healthy joints are required for maximum activity.



Everything you do injures your joints a little bit. The million-dollar question is whether they're able to keep up and repair the damage. Stem cells live in all of our joints and are like little repairmen. As we age, there are less of these repairmen, so at some point, wear and tear can exceed the ability of the joint to repair itself and arthritis begins. Many physicians have begun to supplement this repair capability using various technologies, including stem cells. Click on the infographic labeled "Articulation" to learn more about your joints.







Neuromuscular: Your nerves drive your muscles, yet this fact seems lost on much of medicine today. We'd all accept that when the nerves are severed, muscles die (think Christopher Reeve's spinal cord injury that led to his severe muscle atrophy). Yet what happens when smaller amounts of nerve irritation are present? If you have a chronically tight or weak muscle in your arm or leg, you might not think it's due to an irritated nerve, especially one in your neck or back, but it may be. When the muscles are impacted, they develop tight and weak knots known as trigger points. All the muscle strengthening in the world won't help a muscle when it has lots of these trigger points. They need to be cleaned out before the muscle can work normally. Click on the infographic labeled "Neuromuscular" to learn more.



Your individual joints need to have the same amount of full motion in all directions. If they don't they will experience more wear and tear.

The upper neck supplies information to the brain about body position. Injury to this area can cause headaches and body asymmetry.

Loss of hip movement in one direction can impact the alignment of the knee cap in it's groove..

N/mmetry

If your spine isn't straight, or doesn't have full and equal motion in all directions, then your joints will wear unevenly.

The foot can hit the ground unevenly due to irritated nerves in the back, which can lead to problems in the knee and hip.

Symmetry: Why does one joint in the spine get arthritis and not the other? Sometimes these things are due to prior injury and sometimes to lack of symmetry. Our bodies are built to work optimally only when both sides are exactly like the other. When one of our joints loses normal motion, you can bet other parts will be impacted. When the spine isn't straight or doesn't have full and equal motion in all directions, your facet joints will wear unevenly. When one leg is shorter or one ankle turns in more, again certain parts will wear out faster. Given how important left-right and front-back symmetry is, it's amazing more attention isn't paid to the problem. In order to get well, you must try to reestablish as much normal symmetry as possible. Click on the infographic labeled "Symmetry" to learn more.



In order to understand why you have chronic back pain and how best to get rid of it, you must understand and deal with all issues in the SANS system. You must address any instability issues, problems with your joints, and irritated nerves and muscle trigger points, and, finally, you must get your body as symmetrical as you can. This book is a companion book to <u>Orthopedics 2.0</u> and will explain the Orthopedics 2.0 approach as it relates specifically to the spine and the Regenexx procedures that can help address spinal issues.

Getting Personal: Why Should I Care About SANS?

While everyone wants a quick fix, what I hope to teach you in this book is that getting back to where you used to be is first about choosing the right technology to help and second about fixing all of the associated problems, like weakness in muscles, asymmetry, or poor stability. That may take some time beyond a magic-bullet shot or surgery. Here are some examples of quick fixes that don't work and some solutions

I have really bad headaches, and all my doctor wants to do is throw medications at me that make me feel foggy and "out of it."

Modern medicine loves to write scripts. Many doctors have been taught in medical school and in their practices by the companies selling the pills that they're great! However, wouldn't it be better to figure out what's causing the headaches and treat that?

What are the 7 causes of headaches and their treatments?

My head feels heavy and I have a headache...

Your neck muscles are jacked directly into the covering of the brain...

Some doctor told me all I needed was a magic injection of amniotic, placental, or fetal stem cells and I would be fixed.

How is that supposed to work? Are these even the right kind of stem cells? What are the right kinds of stem cells?

See these quick links:

<u>"Amniotic stem cells" (a.k.a. fetal or placental) have become an epic case of widespread</u> <u>consumer fraud.</u>

Fat stem cells aren't as good as bone marrow stem cells at helping orthopedic problems.



I have pain in my head, neck, upper back, or lower back/hip, but nobody is quite sure what's causing it.

There are many things that may be causing your pain outside of a slipped or bulging disc. The gold standard of figuring out which structure is causing pain is imaging-guided numbing injections. Did you know there may be abnormalities on your MRI that were missed?

See these quick links:

Diagnostic blocks may save your spine from a useless surgery.

What nobody ever told you was really wrong with your MRI.

The differences between traditional and the regenerative Orthopedics 2.0 approach to spine care are stark.

<u>Understanding your spine is crucial to understanding what can go wrong (video included).</u>

A doctor told me all I needed was an injection of fibrin to "seal the disc" in my back or neck, and he showed me good evidence of an FDA trial that proved this worked. He also had before and after X-rays of how my severely degenerated discs will be "plumped up" by injecting fibrin.

While sealing a leaky disc is an attractive concept, the "evidence" showing that this works is more of a bait and switch than real proof it's effective. In addition, those X-rays you saw were only a few minutes after injecting fibrin, and, regrettably, that doesn't last long.

Learn more at these links:

The "bait and switch research" that shows that sealing the disc will help you (the research actually shows that this doesn't work)...

Does sealing the disc really help?



Chapter 1: The Regenexx Difference





When we first pioneered low-back-disc stem cell therapy in 2005, we were the only physicians in the United States doing this type of work. The last few years have seen a bevy of clinics opening and offering stem cell therapies for pretty much whatever ails you—from arthritis to ALS to COPD to MS. A few of these clinics are legitimately trying to do a good job, but most are not. How can you tell the difference? First, let's look at the clinic types that are popping up.

1. The Miracle-Fat-Stem-Cell Clinic: These types of clinics using stromal vascular fraction (SVF) fat as their source of stem cells offer treatments for a multitude of diseases, including knee and hip arthritis. They perform a <u>small liposuction to get cells</u> and as such are usually run by a plastic surgeon who oversees a processing facility that distributes cells to other medical specialists. Some claim to be operating research studies, but when I have investigated these further, most of this is more sales than reality (e.g., one clinic system claimed to have an Institutional Review Board [IRB] research approval that turned out to have been rescinded).

In addition, on the orthopedics side of the treatments, these are <u>usually blind</u>, <u>nonspecific</u> <u>injections (without any guidance to ensure placement) somewhere in the vicinity of the</u> <u>painful joint</u>. They frequently will combine these local injections with an <u>IV infusion of fat</u> <u>stem cells</u>, 97% of which will end up in the lungs and never see the joint. As you know from the previous discussion, <u>fat stem cells don't work as well as bone marrow cells for</u> <u>orthopedic purposes</u>, so the orthopedic side of the business seems to be an afterthought to drive revenue.

2. The Little-Bedside-Machine Clinic: These clinics are often more focused on orthopedic problems, but they use an automated bedside "one size fits all" machine to process bone marrow cells and platelet rich plasma. Some of these clinics do offer guidance of the injection, but very little effort is placed on tracking patients or reporting outcome data. So the type of treatment-registry data that you've read about here isn't going to be reported by these clinics, leaving the patient to fly blind on how well these procedures work or don't work.

These machines also produce about one-tenth to one-fifteenth of the stem cells per unit volume as a Regenexx-SD procedure (based on our lab studies). They also only isolate one fraction in the bone marrow that contains stem cells and discard the other fraction (not knowing that it has valuable cells).



So what are the key components of a reputable clinic?

- A. Treatment-registry tracking of patients
- B. Guidance of the injection
- C. A focus on orthopedic problems
- D. Candidacy grading
- E. Published research
- F. A customized approach to the processing of your tissue
- G. A clinical training and certification program for affiliates

Treatment-Registry Tracking of Patients

Any new therapy that is standard of care needs to have data collected, even if it looks very promising from the standpoint of patient experience (e.g., a doctor says it has worked well in other patients). This means that standardized questionnaires are sent to the patients at set time points to see if they have less pain, more function, or had any complications with the procedure. This is a huge commitment on the part of the clinic and the doctor.

As an example, right now we use a clinical research organization (CRO) quality, customized software deployed through a nonprofit to assist us in collecting data on the patients we have treated. There are many full-time employees to collect data, several part-time supervisors, and a full-time biostatistician to analyze this data. When we want to report the data, we must enlist the help of expensive physicians to call patients who haven't responded to their questionnaires as this helps to make sure we have enough data to report. While we have a full-time biostatistician, we must also use more expensive doctor time to help our biostatistician decide what's clinically meaningful to analyze.

How can you tell if a clinic is doing this? They will have data from their patients that they have collected and reported, usually on an annual basis. Why is it important to see that clinic's data? A procedure like this may produce very different results in a different doctor's hands. In addition, the clinic will be able to tell you exactly how it collects its data, who collects it, how often, and so on. For example, a proper treatment registry collects data at set time points, like one month, three months, six months, one year, two years, three years, and so on. If all you get is a call from a nurse, like you would after any common surgery, then this isn't nearly enough.



Guidance of the Injection

How we deliver stem cells as part of interventional orthopedics makes a big difference. While delivery into an arm vein (IV) is attractive because of the low level of expertise needed to deliver cells, studies have consistently shown that adult stem cells delivered in this fashion are trapped in the lungs (<u>pulmonary</u> <u>first-pass effect</u>). Of even more concern is a recent study showing that for patients considering the use of stem cells to treat central nervous system (CNS) disorders, only about 1 in 200,000 cells injected via an IV route reaches the brain and central nervous system (1.5–3.7% made it past the lungs, 0.295% made it to the carotid artery, and 0.0005% made it past the blood-brain barrier into the brain).



At this point, until these pulmonary first-pass issues are worked out, credible stem cell delivery is local. This means placing cells directly into the tissue or into the arterial circulation that directly supplies the tissue. In addition, based on our clinical experience, for orthopedic applications (and likely for others), it's hyperlocal, meaning that placement of cells into one part of the joint may provide results; whereas, nonspecific placement into the joint may provide fewer results. <u>This means imaging guidance to place cells into joints</u> <u>is very important</u>.

A Focus on Orthopedic Problems

Figuring out how to maximize the effects of stem cells is critical. Let me give you some examples. Early on in our experience, we added in the use of a billionth of a gram of a common medication based on many research papers showing it helped stem cells create more cartilage. We then had a natural experiment where we were able to compare patients who didn't have their cells exposed to this very lowdose medication versus those who did. The graph to the left shows how much better the patients who got the medication did, and as a result, this medication became a standard part of our protocol. This is just one example of how little things about how stem cell procedures are performed can make big differences.





As a result, it's not credible for a clinic to offer therapies for 10 different diseases that have little to do with each other. Credible clinics focus in on one or two body systems and perfect their treatment protocols. This is why we've kept our hyperfocus only on orthopedics and why we continue to do the basic science needed to improve our treatments. For a short video break, click on the Regenexx video link below.



Candidacy Grading

There is no medical procedure available (including stem cells) where all patients are great candidates and expected to do well. For over a decade, we have been offering stem cell treatments, and during that time we have graded patients with regard to candidacy. These good, fair, or poor candidate grades have literally dissuaded hundreds of patients who were considered less-than-stellar candidates from undergoing the procedure.

Several years ago, after we had enough outcome data on the Regenexx-C cultured procedure, we lifted those grades as the statistical analysis didn't show that more-severe arthritis patients did any worse than patients with mild arthritis. However, we kept them in place for the Regenexx-SD procedure as the literature on platelet rich plasma (PRP) did show that while PRP worked well in mild arthritis, it often failed in patients with moderate and severe arthritis. In addition, the same holds true for knee microfracture. Again, we turned away hundreds of patients because adopting a conservative candidacy system was the right thing to do.



In 2012, our first registry analysis of Regenexx-SD showed that these candidate grades roughly followed the outcome (patients then considered "poor" candidates with severe arthritis generally had less-robust outcomes than those who were "good" candidates with mild arthritis). So we continued to try to convince many patients with more-severe arthritis not to undergo the procedure.

In late 2013, we again ran the data with more knee cases in our registry. Interestingly, as the numbers of patients being tracked increased, the association between severe arthritis and poor outcome didn't hold up, meaning that the severe-arthritis patients who chose to do the procedure anyway had about the same outcome as the mild-arthritis patients. So after many years of turning away hundreds of patients, we now feel comfortable in the statement that the Regenexx-SD patented knee stem cell procedure and its three-part treatment process works as well in severe-arthritis patients as it does in mild-arthritis patients.

So ask the clinic about your candidacy. Is it good, fair, or poor, and why? Is anyone considered a poor candidate? For example, I can tell you that based on our existing registry data, if you have hip arthritis and are <u>over the age of 55, you're less likely to do as well as someone who is younger.</u>

Published Research

In any new procedure, research should be published as the data becomes mature enough. We have always prided ourselves in <u>submitting our data to peer-reviewed journals for</u> <u>publication</u>. This takes an immense amount of work as any single publication often goes back and forth for months to a year before it's in a form that will appease reviewers. So ask if the clinic's providers have research that they've published. <u>Be careful here as</u> this is a prime area for bait and switch; I've seen clinic websites that show research that has nothing to do with the stem cell type or procedure the clinic is using. For example, showing research done by someone else on bone-marrow-isolated mesenchymal stem cells when the clinic actually uses fat stromal vascular fraction (an apples-to-oranges comparison). Ask the following: Where is your data on what you do? Where can I find publications with your name on them? Ninety-nine percent of the time, you'll find that the clinic has no such data.

A Customized Approach to the Processing of Your Tissue

Every patient is unique, yet many clinics use automated one-size-fits-all machines to process tissue because the capital and time investment is less. Most of these machines treat every sample as if it were the same, yet every sample is really quite different. Hence what comes out of the machine is often not processed based on the individual



characteristics of that tissue, so the stem cell yields are compromised. In addition, for bone marrow, all commercially available machines on the market today discard valuable stem cells that the Regenexx-SD procedure retains. Ask the clinic providers if they process your tissue by hand or if they use a small bedside machine that will treat you like a number rather than an individual.

A Clinical Training and Certification Program for Affiliates

There's a saying in medical training: "See one, do one, teach one." Regrettably, it refers to the practice of physicians often having very little experience or training before working on patients. The stem cell "wild wild west" exemplifies this problem.

Let me give you a real-world example. A few years back, I met a physician at a conference who was using bone marrow stem cells. He was describing his cell-harvest technique from the back of the hip and mentioned that he took 100 cc of bone marrow from one site. I stopped him and asked who had taught him to do it that way? It turns out that a physician who visited our clinic (and was never trained by us) in 2007 had taught him this was the correct method. In fact, what that physician had seen was us taking six to eight much smaller-volume samples because our own research and that of others had shown this increased the number of stem cells in the sample. The doctor who had visited simply didn't know why we were taking so much time to get so many samples, and the doctor simplified the technique to a single high-volume sample. In the process, he also dramatically reduced the number of stem cells. He then went out and taught this modified technique to hundreds of physicians!

As a result, the <u>Regenexx Network</u> of physicians, whom we have officially trained, is the only stem cell provider network in the world with an extensive and mandatory clinicaltraining program. First, rather than taking any willing provider, we turn down about 8 to 9 in 10 physicians who want to join, because they don't have the right skills to practice interventional orthopedics. Second, the few providers we do accept are extensively trained here in Colorado.



This includes a core-skills checklist of procedures they must know. In addition, treatment of any area (e.g., shoulder) has both a didactic education and hands-on demonstration of skills in a cadaver lab.



Regenexx Network physicians use the platelet procedures (Regenexx-PL and Regenexx-SCP) and same-day stem cell procedures (Regenexx-SD) and have in-house processing labs so they can customize the tissue processing to meet the needs of your sample. They also are part of our treatment registry. This means the outcomes we report include our own data plus the results obtained by our network providers.

I hope this detour into picking out the wheat from the stem cell chaff was helpful. Now let's explore why we believe many traditional drug approaches to joint arthritis, ligament/ tendon tears, or spine diseases have problems. We'll also explore what alternative things you can do to help your ailing body.



Chapter 2: Stability



"In all of Shakespeare's plays, no matter what tragic events occur, no matter what rises and falls, we return to stability in the end." — Charlton Heston



Take this short quiz to see if this section applies to you. If you answer any question with a yes, you may have a spinal stability problem.

- 1. My back or neck gets very sore or swollen after I exercise. Y N
- 2. I hear cracking/popping in my back or neck when I do certain activities. Y N
- 3. My back or neck feels like it's loose or moves too much. Y N
- 4. My head gets heavy by the end of the day. Y N
- 5. My arms or legs tingle or get numb when I'm active or when I sleep. Y N





What does it mean to be stable? Stable in a mechanical sense means resistance to falling apart or falling down. For your body, joint stability is a very big deal, yet you likely haven't been told the whole story. You see, you've only been told about a very unstable joint that requires surgery to fix a completely torn and retracted ligament. Yet it's the instability you don't know about that's slowly frying your spine, one

movement at a time. Discovering which areas have this kind of instability, called subfailure, may save you from spine surgery.

What is subfailure instability, and how do you know if you have it? Subfailure instability means that the levels of the spine aren't kept in exact proper alignment during movement. Why is this important? When the individual spine bones uncontrollably crash into one another or even just can't be kept in perfect alignment, the nerves can get pinched and the discs and facets joints wear down much faster. An unstable part of the spine literally experiences many times the wear and tear of a stable spine, and bone spurs form. Since stability in many joints is the number one determinant of whether that joint will have a long happy life or become "old" before its time, it's a wonder more time isn't spent assessing this component of joint health.

Let's slice and dice spine stability a little further by separating the type you've heard of and that is usually easily diagnosed from the type that will slowly destroy your joints and will likely never get diagnosed. There are two major types of instability: surgical and subfailure. Surgical instability is less common than its more prevalent cousin—subfailure instability. However, surgical instability is usually the only type that the orthopedic-spinesurgery establishment treats. This means that a spinal level is very unstable and unable to hold itself together at all. In these cases, surgery is often needed to stabilize the joint. Examples would be severely damaged ligaments in the spine, where a spinal cord injury is feared if the spine isn't surgically stabilized. A true <u>surgically unstable spine</u> may need a fusion where the bones (<u>vertebrae</u>) are fused together with additional bone. Regrettably, this isn't a perfect solution as the <u>fusion will cause other spinal segments above and below</u> to be overloaded.



Subfailure means that the ligament hasn't completely failed (torn apart like a rubber band), but instead it's only partially torn, degenerated, or just loose. This much more common type of instability often doesn't require surgery and is characterized by small extra motions in the spine just beyond the normal range. In fact, if you have this type of instability, you likely aren't aware you have this problem.

Our understanding of subfailure instability is younger and more immature, so while we have some diagnostic tests to detect this type of instability, our understanding of what is normal and abnormal is only now coming into focus. However, this type of instability is quite real, and it's a clear and long-term insidious drag on spine health.

More on Subfailure: It's All About Your Ligaments and Your Muscles

There are two types of subfailure instability: ligament and muscular. Passive ligament stability keeps our spine bones from getting badly misaligned. Think of ligaments as the living duct tape that holds the spine together. Without these ligaments, every step would cause the spine bones to experience a potentially damaging shift. On the other hand, active muscular stability is made up of the firing of deep multifidus muscles that keep the spine bones aligned as we move and represent the stability fine-tuning system. The spine building blocks tend to want to slip slightly out of alignment as the spine bends, twists, or slides, even with intact ligaments. As this happens, signals are sent to selective muscles that surround each level of the spine (multifidus) so that they adjust and correct the alignment. Without this active system, the spine would be "sloppy." This muscle firing is a muscular symphony, with microsecond precision being the difference between the poetry of beautiful movement and an asynchronous chorus of potentially damaging spine bones colliding against one another.



So what do the muscles do again? The muscles provide the fine-tuning. They act as constant stabilizers for the joint, keeping it in good alignment while we move. This small area where the joint must stay to prevent damage as we move is called the "neutral zone."

So in summary, stability is about both muscles and ligaments. Your deep multifidus muscles provide constant input to the joint to keep the spine bone alignment fine-tuned as we move. When the spine bones move too much,



the ligaments act as the last defense to prevent joint damage from excessive motion. Now let's explore how poor stability in the spine may be harming your joints, like the knee, ankle, shoulder, elbow, or hand. If this important structure isn't stable, nothing else will be quite right.

The Spine is a Marvel of Stability Engineering

The spine is advanced engineering. <u>The spinal column is a series of blocks</u> <u>that stack one upon the other and provide a base of support for the</u> <u>extremities and protect the spinal cord and nerve roots</u>. These interlocking blocks (<u>vertebrae</u>) use the same stability model as described above muscles and ligaments.





Stable Unstable

Now let's look at how the spine stays stable. What happens when you place a bunch of blocks one on top of the other? This tower of blocks gets less stable as the pile gets higher (see left). One way to stabilize this high tower of blocks would be to tape the blocks together. This would make the blocks more stable but wouldn't allow much motion. You could use more flexible tape than duct tape or Scotch tape, but again you'd either end up providing too little stability (highly elastic rubbery tape that gives a lot when you stretch it) or too much (duct

tape or Scotch tape that's more rigid).

Imagine if we had to move all the taped blocks so they could bend into a C-shaped curve (see right)? How stable would your blocks be then? The right kind of tape (ligament) would likely allow this motion, but the individual blocks would still start to shift against each other. This shifting could result in disaster, as the <u>spinal cord</u> runs through a hole (<u>spinal canal</u>) inside the blocks, and the spinal nerves exit between the blocks through a special bony doorway (<u>foramen</u>). As a result, too much movement between the blocks means nerve damage or, worse, a <u>spinal cord injury</u>. This is the dilemma of the spine—how to stack lots of blocks (about 25 high



in most people) while keeping the whole thing stable and flexible as you move and at the same time protecting the nerves. Is there a solution? Yes, the muscles provide that solution. They keep the blocks (vertebrae) aligned against one another with millimeter precision while you bend and twist. In fact, since the muscles are so important for a healthy spine, <u>it's unsettling that most physicians couldn't tell you which group of muscles</u> <u>provides this stability (it's the multifidus as you may have guessed).</u>



What are the specific ligaments that live in the spine? Look at the picture below where I have highlighted in red all the ligaments that are visible from the back view of the spine and pelvis. There are ligaments everywhere! Wouldn't you think these ligaments might be a target for therapy? Has any surgeon ever said that your ligaments were normal or abnormal? Regrettably, our medical-care system, outside of a few exceptions, ignores the ligaments that act as the duct tape to hold your spine together!

Spine Ligaments to Get to Know

I have shown the low-back ligaments below, but many of these are the same in the neck and upper back.



Supraspinous/Interspinous Ligaments: These ligaments are the "duct tape" that runs down the very back of your spine in the middle, on top of, and between the spinous processes. The spinous processes are the bony bumps you can feel in the middle of the spine along your back (thinner people will feel more of them). They control the motion of the vertebrae when you bend forward. They are commonly injured in car crashes and other trauma and may become degenerative with age.

Intertransverse Ligaments: These ligaments travel between the bony processes (transverse processes) that stick out of the side of the spine. They control motion in the vertebrae when you bend to the side. If bending that way makes things worse, these ligaments may be stretched and too loose.

Iliolumbar Ligaments: These ligaments go from the side of the L5 and sometimes L4 (bottom low-back vertebrae) to the side of the hip/pelvis area. They are major stabilizing ligaments of the most mobile part of the low back. Problems in these ligaments can refer pain down to the thigh and front of the hip. To learn more, <u>click here.</u>

Sacroiliac (SI) Joint Ligaments: These ligaments are the massive ligaments at the back of the sacroiliac joints (the joints between the tailbone and back of the pelvis/hip). Women have more issues here than men, as these can be stretched during childbirth, falls onto the butt, and trauma. The pain is often in this area at the back of the hip and can refer down the leg.



Sacrotuberous and Sacrospinous Ligaments: These ligaments anchor the bottom of the tailbone to the rest of the pelvis, and when injured, can refer pain down the leg to the outside of the foot, almost like a pinched S1 low-back nerve.

Alar, Accessory, and Transverse Ligaments:

These highly specialized upper-neck ligaments hold the head on. The alar ligaments (blue to the right) reach up from the C2 vertebra and attach strongly to the skull, while the transverse ligaments (red to the right) hold the pivot point of the C2 vertebra tightly to the C1 bone, allowing you to turn your head without getting a spinal cord injury. These ligaments can be injured in car crashes or when something hits the head. Their laxity can cause everything from dizziness to visual changes to severe headaches.



Upper Cervical Spine: Posterior to Anterior View with Ligaments

More on Muscular Stability in the Spine



If muscles help solve the problem of keeping the blocks aligned against each other while the spine moves, which muscles are these? For most of the spine, this special muscle is called the <u>multifidus</u>. These small muscles travel from vertebra to vertebra to keep the spine bones in proper position. In the 1990s, some very <u>smart scientists noted that these muscles were smaller</u> than usual in most patients with chronic low-back pain. Since that time, atrophy (when the muscles get smaller and weaker) of these and other spinal muscles is a known cause of low-back and leg pain. These muscles can be seen on almost all MRIs, and <u>numerous research</u> articles have correlated shrinkage of these muscles with back or leg pain, but, regrettably, this atrophy is almost never commented on in radiology reports!

If this muscle is such a big deal, why have you never heard about it, and why has nobody told you it's a problem? Good question! Regrettably, the vast majority of medical providers, physical therapists, and chiropractors haven't read about this research. Why? The multifidus hasn't had a champion to take it from the pages of medical journals to the



prime time of major medical conferences. This is despite the fact that to make sure your spine bones stay aligned and aren't constantly bumping into spinal nerves, these muscles have got to be firing on all cylinders all the time.

What Happens When the Muscular-Stability System Goes Down?



Remember that ligaments are only one-half of the joint stability equation—muscles are the rest of the equation. When the muscles fail to align the joint, the joint becomes sloppy. Too much movement is allowed in all the wrong directions. This causes excessive wear and tear on the joint (<u>arthritis</u>), whether the joint is a disc in the low back or a knee, shoulder, ankle, or hand joint. Obviously, the first way to help this problem would be to strengthen muscles around the joint. However, just getting the stability muscles stronger sometimes

isn't enough. Sometimes the nerves telling the muscles what to do doesn't work well. In my experience, patients who have this issue don't know they have it. In cases like this, no amount of strengthening will help until the nerve issue is addressed (see the <u>"Neuromuscular" chapter</u>).

A Dramatic Example of What Happens When the Spinal-Stability System Is Off-Line



A number of years ago, I went for a several-hour mountain-bike ride. <u>I had just read a research paper</u> <u>that discussed how factory workers showed weaker</u> <u>back-stabilizing muscles after being in a bent-forward</u> <u>posture for several hours.</u> Regrettably, that day, my own back would become an example of this phenomenon. Right after the ride, I went to pick up my son, felt a pop in my back, and collapsed. It took days to recover, and for the first 48 hours, I was in agony. In my case, the <u>lumbar multifidus</u> was "off-line" from being bent forward on my bike, which caused irritation of the

nerves. This led to a period of instability in that spinal segment, a time during which I was more likely to get injured. Essentially, that spinal segment was left completely unprotected by the spinal muscles when I went to pick up my son. My back, thus, went down in flames like the Hindenburg! This "multifidus" off-line mechanism, whether it's caused by bike riding or just something else that irritates nerves, is believed to be the cause for most acute-back-pain episodes.



I've felt the same thing occur a few more times, most notably once while dead-lifting weights in the morning (after sleeping in a flexed posture all night). These are all dramatic examples of what happens when stabilizing muscles are turned off. How do I get these muscles back online and almost immediately fix my pain rather than lying in bed for days? An epidural injection around the nerves reduces the nerve swelling and generally turns these important muscles back on. Usually, after one of these low-back meltdowns, I can exercise on a limited basis within hours of getting an epidural. However, as you'll learn later in this book, all epidurals aren't created equal.

The Heavy Head: Another Example of a Weak and Compromised Spinal-Stability System

Many patients who have been in a car crash tell us that <u>their head feels heavy</u>. How is this possible? The head is a bowling ball being balanced on the end of a stick (the spine). There are muscles at the top of the neck as well as in the front and back. These deep muscles help to hold the bowling ball on the stick. As you'll see in the section on symmetry, <u>losing the natural curve in the neck</u> can push the bowling ball forward on the stick. In addition, the articulation section will discuss how injuries to the neck joints can take the stabilizing muscles off-line. All of this can conspire to make someone feel like he or she has a bowling ball for a head!

Fusing the Spine Is Usually a Bad Idea

This past decade has seen an explosion in the number of fusion surgeries. What does this mean? The concept is simple. If some joint is unstable, you just bolt it together and put bone in the area so the joint becomes immobile solid bone. Is this a good idea? Think about the fact that your body is a finely tuned machine where millimeters of extra motion or lack of motion can cause problems. Now think about what would happen if I made a joint not move at all. It wouldn't be long before the joints above and below would begin to wear out from the extra pressure as, after all, the forces have to go somewhere.

Spinal fusion surgeries are associated with a myriad of complications and problems. In addition, as you might imagine, the joints and discs above and below the surgical site that's fused get fried very quickly. This means a patient can spend a lifetime chasing the next level as it degenerates faster because of the spinal fusion—this is called adjacent segment degeneration (ASD).

While I have seen a handful of patients, in my career, who really need and would benefit from spinal fusion, these patients, in my opinion, represent only about 5% of all the patients who are currently being fused in the United States. In fact, regrettably, treating the chronic pain suffered by spinal-fusion patients could keep a pain-management doctor in any community busy full time.



Recent Research on Spinal Fusion—Worthless for Helping Patients

Given the incredible invasiveness of bolting the spine together and the huge cost of these surgeries and the fact that we know that ASD is real, you would think there was incredible evidence that the surgery works. Nope. There was a recent publication of the world's first randomized controlled trial of lumbar fusion.

<u>In this new study</u>, almost 300 patients with low-back spinal stenosis (arthritis that's putting pressure on nerves) were randomly assigned to undergo either a traditional laminectomy or that surgery plus a lumbar fusion. **The result was not surprising—adding a fusion surgery didn't improve outcomes at two or five years after surgery!**

Let's think about that for a moment. We're exposing patients to much higher surgical risk by adding a fusion and also placing them at risk for breaking down the adjacent sites of the spine. Without any positive effect of adding the fusion, why are we doing this surgery?

Why Physical Therapy Sometimes Fails



The past one to two decades of supervising rehab programs for patients has taught me that while strengthening the muscles often helps the muscular stability of the spine, sometimes it fails. Why? **The muscles are controlled by nerves.** While much of traditional medicine has focused on big problems in nerves that can be picked up on static imaging, like <u>MRI</u>, or on electrical tests, like an <u>EMG</u> (electromyogram—a test where needles are inserted into the muscles), the more recent research shows that a lot can go wrong with nerves

<u>that is generally invisible to these tests.</u> This smaller amount of spinal nerve-root irritation can wreak havoc with the muscles by shutting parts of them down. This problem, despite the research, largely is ignored by physicians. This is because our medical-care system isn't yet focused on detecting these smoldering nerve issues.

When a part of a muscle gets shut down by nerves, it may not be responsive to strength training. These shut-down muscle areas are called <u>trigger points</u>. An easy way to get rid of these tight and weak areas of the muscle is trigger point massage, or direct pressure on the area. When this fails, a trigger point injection, or dry needling of the spot (IMS, or intramuscular stimulation), is often the answer. In our experience, this can allow that part of the muscle to work again and start helping the joint to stabilize. This concept is discussed further in the "Neuromuscular" chapter.



I can't emphasize enough that if traditional physical therapy fails, you must look at whether you have nerve and muscle problems. You may believe that you have no spinal-nerve problems, but it's more likely than not that you do and just don't know it!

Why Can't Strong Back Muscles Substitute for a Bad Ligament-Stability System?

If your back has one or more loose ligaments, maybe you can just get stronger by doing lots of physical therapy and that will suffice. After all, can't the muscles just substitute for the loose ligaments? As discussed, there are two types of stability systems, the fine-tuning is provided by the muscles while the ligaments prevent serious abnormal movements that can lead to catastrophic spinal damage. If the ligaments are stretched out a little bit, but still intact, the muscles may be able to substitute and protect the spine in most situations. However, if the ligaments are stretched or damaged so that they allow bigger abnormal motions in the spine, no amount of muscular stability will help. In the end, while having stronger stability muscles may help reduce some of the wear and tear, the spine will still get into abnormal alignments that will lead to accumulated damage. So if ligaments are stretched, it's best to tighten them, and this can often be done without surgery (see the section on regenerative medicine), though rarely, if they're completely torn and retracted, the only option may be surgery.



How Do I Know if I Have a Spinal-Stability Problem, and What Can I Do to Help It?

How do you know if you have a spinal-stability problem? Patients often complain of popping or cracking in the spine. They may at times feel sudden shifts in the spine. For example, in the neck, patients may feel that by the afternoon or evening they have a "heavy head." In other patients, there may be no perceptible sense of instability or popping/cracking in the spine, just pain or swelling after activity. These patients usually have smaller amounts of microinstability.

Regenexx Simple Muscular-Stability Tests

When you get evaluated using the SANS system, these are the quick and simple physical exam tests that will determine if you have good or poor stability in the spine and other areas of the body. Knee pain, for example, can actually be caused by a spinal issue rather than a knee issue, so this tests all areas. There are also other things that will be evaluated, like the status of your spinal-stabilizing muscles on MRI. If you have pain with any of these movements, make sure that you pay attention to the rest of these chapters.



We expect our patients to be participants in their own recovery, so we want them to take this test on a monthly basis when being treated. If you're a Regenexx patient, before your first evaluation, please take the test, print out the work sheet, and bring it to your first evaluation. If you were given a physical book, you can find the work sheet in Appendix A. If you're not a patient, you can take the test and use the work sheet to record where you are at any given time (click here for a printer-friendly link to the work sheet).

Caution! These tests may cause injury or exacerbate the conditions of patients who have a more fragile stability system or who are at a lower level in their stability. Please do not attempt these if you get injured easily. If you do decide to perform this assessment, if you get significant pain during any given test, stop immediately. You are performing these tests at your own risk.

Getting Ready: For some of these tests, you'll need an assistant. You'll also need a timer or clock/watch with a second hand (there are good stopwatch programs on most smartphones).



Lift arms above head all the way. Then have someone hold head firmly and retest. Fail if it's easier to lift the arms with head hold.

Neck–Upper Cervical (Scalenes): The first test is simple. Just lift your arms out to your sides and all the way above your head. Is this easy or hard? Can you easily get your arms up all the way? Now have someone stand behind you and hold your head very firmly. Your head shouldn't move at all as you perform the same hands-over-head maneuver. Was it easier to lift your arms with your head stabilized by someone? If you found it hard to lift your arms

over your head without someone stabilizing your head, then you may have upper-cervical instability. This means that the muscles or ligaments stabilizing your upper neck may not be strong enough. **If you experience significant pain with this maneuver, stop!**

Scoring: If you feel no difference with or without the head hold, then give yourself a 3 here. If you feel a noticeable difference and can get your arms all the way up, you score a 2. If you feel a difference and get your arms over your shoulders but not all the way up, you score a 1. If you can't get your arms over shoulder height, give yourself a 0.




TTuck chin and lift head about 2-3 inches and time. If any loss of height or chin tuck occurs, then stop timing. Normal is 38 sec for men and 29 sec for women.

Neck—Deep Neck Flexors: The second <u>neck test was developed by an</u> Australian researcher and is a bit more challenging. Lie on a flat and firm surface, like the floor, and have someone time and monitor you. First, tuck your chin fully and then lift your head 2–3 inches. Have your monitor start the timer. The clock stops when you lose any degree of the chin tuck or your head is unable to maintain the same height. Normal for men is 38 seconds and for women is 29 seconds. If you can't do this, your deep neck flexors are weak and you fail this part of the test. These are <u>important muscles that stabilize the front of your</u> neck. When these muscles are weak, patients often report headaches. If you experience signifi cant pain with this maneuver, stop! Scoring: Write down your timed score.

Neck—Extensors: The <u>third neck test was developed by a US physical therapist</u> and tests the endurance of the deep and superficial neck muscles that hold your head up. You first lie facedown on a table or firm bed, making sure your chest is stable (you may want to have someone hold you by placing downward pressure on your upper back). You then hold your head perfectly straight with the face parallel to the floor.



Lie face down with your head off the end of a table or a firm bed as shown in A. Make sure your chest is stable and hold your head straight as shown in B. Time your endurance. When your head begins to fall downward or extend due to fatigue (as shown in C and D), the clock stops.

You hold this position and time yourself. You should be able to get to 20 seconds without your neck and head bending (as shown in C) or extending (as shown in D). If you went into position C as you fatigued, your superficial neck extensors are weak. These help to hold your head up. If you went into position D, your deep neck extensors are weak, and the superficial extensors are taking over. In both of these instances, patients with weak neck-extensor muscles tend to report a heavy head or fatigued neck by the end of the day. **If you experience significant pain with this maneuver, stop!**

Scoring: Write down your timed score.





With arm out to the side with the thumb down, move the hand completely down and up in the three planes noted. Pass is at least 10 reps slowly (30 updown movements. Shoulder—Rotator Cuff: The test for the shoulder focuses on rotator cuff muscle endurance. Given that the shoulder is linked to the neck, a shoulder strength test also tests the stability of the spine. The arm is held out to the side with the thumb facing down and pointing toward the floor. The arm is then moved down toward the side and back up to shoulder height. You perform this movement in the three planes shown (behind the body, at the body, and in front of the body). Do this slowly. **If you experience significant pain with this maneuver, stop!** You should be able to do this 10 times in each

position (30 reps total). You may be fatigued. If you can't get that far without stopping, you need to strengthen the rotator cuff. These muscles help to stabilize the ball of the shoulder joint in its socket.

Scoring: If you can get through 30 reps (10 in each plane) without pain (although you may be mildly to moderately fatigued), give yourself a 3 here. If you can barely do this and experience much fatigue and effort or have pain with this test, you score a 2. If you can't get through all of this due to fatigue or pain, you score a 1. If you can't do this at all, give yourself a 0.

Core—Abdominals: There are two stability tests for the lower back, <u>both pioneered by a Japanese researcher</u>. The first is a simple sit-up maneuver where you begin by lying faceup and bringing your hips and knees to a 90-degree position as shown. Make sure you keep your neck flexed. Set a timer and hold this position. The normal time for men is 182 seconds (3 minutes), and for women it's 85 seconds (1.5 minutes). **If you experience significant pain with this maneuver, stop!** If you have a lot of pain with this maneuver, you may have a disc-pain issue as flexion places more pressure on the disc.

Scoring: Write down your timed score.



Lie on your back with your hips and knees both at 90 degrees and lift your torso off the ground. Time how long you can hold this position.





Lie on your stomach on a stiff pillow and extend your back so that your chest is off the floor and hold. Time how long you can hold this position.

Core—Low-Back Extensors: The next test starts with lying facedown on the floor and placing a firm pillow under your stomach with your hands at your side. The pillow should be firm enough or doubled up so that you can extend your back and lift your chest off the floor as shown. Your neck should remain flexed. Set a timer and hold this position. For men, the normal hold is 208 seconds (3.5 minutes), and for women it's 124 seconds (2 minutes). If you experience significant pain with this maneuver, stop! **If you have pain with extension like this, you may have either a facet or lumbar stenosis problem.**

Scoring: Write down your timed score.

Hip/Knee: There is a single test for hip and knee stability as both of these are linked. Stand and then balance on one foot (the side you want to test). You then perform a single-leg squat while trying to keep your body as straight as possible. If you're able to balance so that you're straight for about one-half of the deep knee bend and for the return back up, then you pass the test on that side. If you must tilt your body over to that side and/or your knee drives inward, you have poor hip and knee stability and fail the test. If you can't perform the half squat, you fail as well.



Normal

Abnormal

Scoring: Write down your timed score for each side as the number of seconds you can hold that side in the position shown in the picture above (Normal) versus the right (Abnormal).



Ankle: You can test the ankle from the same standing-on-oneleg position. Given that the ankle is supplied by nerves in the low back, subtle weakness here can also originate in the spine. Count to 10 while balancing on one leg, and watch your ankle. With your body straight (which is again measuring your hip stability) and hands to your sides, does the ankle roll or have to move back and forth during that 10 seconds? If so, you have an unstable ankle. If not, and the ankle remains rock solid for the 10 seconds, you pass this test. If you have a tough time with this, you may have



an ankle-stability issue. This can impact things all the way up to the knee and can even, in some patients, cause a kneecap issue.

Scoring: Write down your timed score for each side as the number of seconds you can hold the ankle stable.

What if I failed some of these tests? This means you have poor stability in these areas. This could be due to pain shutting down muscles, weak muscles, or irritated nerves that make them weaker or misfire. If it's pain, then you have to find and fix the source of that problem. If it's weak muscles, they may just need strengthening. Finally, if it's irritated nerves, no amount of getting the muscles stronger will help; you need to reduce the nerve irritation. These things will all be covered later in the book.

As you get treated, use this Regenexx Stability Test as your monthly spot-check to gage your progress. You want to increase your scores in each problem area.

What Other Diagnostic Tests Can Diagnose the Problem, and What Therapies Might Help?

Spine Diagnosis



In the spine, larger amounts of instability can be seen on either lowback or neck flexion-extension X-rays. These are tests where the patient looks down and up or bends forward and backward while X-ray films are taken. Regrettably, too often, the technicians who take these films don't push patients far enough into these positions (flexion or forward, extension or backward). If you undergo one of these tests, make sure to push yourself in these motions (without hurting yourself). <u>The research shows that such efforts can reduce</u> <u>the false-negative rates of these tests</u> (a false negative is a test that

fails to show positive even though the patient has the disease). A newer form of this test is called <u>Digital Motion X-ray (DMX)</u>. This test looks at a moving X-ray view of the spine as the patient is put through various ranges of motion. It can also be used in various peripheral joints, such as the knee, shoulder, and elbow.

Spine Therapies and Exercises

The lowest level of spinal-stability training uses ultrasound imaging or other biofeedback devices to help patients contract their <u>multifidus</u> and <u>transversus muscles</u> in the low back. The same type of program can help with neck instability. This program is called deep-neck-flexor strengthening (strengthens <u>longus colli</u> and <u>longus capitis</u>), and this therapy has <u>research to show its efficacy in helping headaches</u>.



The next level of difficulty for spinal stability is usually where most programs start. If this level of therapy makes things worse, then either it's too advanced for the level of muscle atrophy or there are other issues that have to be addressed (like damaged ligaments, irritated nerves, and painful joints). For the low back, the oldest such program is called dynamic lumbar stability, or DLS. In the neck, once patients have mastered lower-level neck-strengthening exercises, like deep-neck-flexor strengthening, we have used the <u>BTE multicervical</u> unit, which helps us identify cervical function, physical



limitations, and capabilities and allows us to record in real time cervical-spine range of motion (ROM) and isometric strength.

The following link to exercises that help build **core** strength:

Basic	<u>Transversus abdominus</u>
	Multifidus
Advanced	Spinal Stabilization at levels 1 and 2
	Spinal Stabilization at levels 3-6
	<u>Bosu Standing Squat</u>

The following link to exercises that provide low-level cervical strengthening for severely unstable patients:

Basic	Deep-neck flexors level 1
	Basic Deep-neck flexor level 2
	Basic Deep-neck flexor level 3
	Basic Deep-neck flexor level 4
Advanced	Neck and shoulder extensor stability

Neck/shoulder stability

Stretched ligaments causing unstable joints can often be helped without surgery by prolotherapy or platelet rich plasma (PRP) injections. This will be discussed further in the next chapter.



Chapter 3: Articulation



"The universe as we know it is a joint product of the observer and the observed." Pierre Teilhard de Chardin





Articulation means joint. For the spine, this means <u>the joints</u> <u>between two spine bones (the disc and facet joints)</u>. The disc and facet joints allow motion in a controlled manner. A joint has certain standard components.

Cushioning: In the spine, the middle of the disc, called the <u>nucleus</u> <u>pulposus</u>, serves as the cushion. This is not an inanimate piece of rubber, but living tissue with cells and structure. Once these components die off, the disc loses its ability to <u>provide shock</u> <u>absorption</u>.

Stability: This is provided by the joint capsule (tough outer covering of the joint) or the ligaments that help hold the joint together. In the disc, this is provided by the tough outer covering (called the annulus fibrosis).

Notice that while our entire medical-care system in orthopedic spine surgery revolves around the disc, in the SANS system, the disc is simply one part of a bigger picture where ligaments, nerves, muscles, and symmetry are equally important or more important. What are the implications of paying too



much attention to the disc? Today's modern ineffective spine surgery!

Arthritis in the spine means that the parts and pieces, like cartilage in the spine joints (facets), outer covering of the disc, and spine bones (vertebrae), are degenerating or wearing down. So how do we access the degree of arthritis? We image it using an MRI. However, do these images paint an accurate picture of why you hurt?

Imaging Insanity, or the Very Poor Correlations Between Structure and Function

Every patient I have ever met wants an <u>MRI</u>, which is a fancy picture of the soft tissues that's created by powerful magnetic fields. While our practice uses MRIs to help define pathology, what if I told you that if you placed a bet that I could tell from your MRI what was causing your pain, I could give you only 1:1 odds (50/50)? For example, several low-back studies have shown that patients with severe problems on MRI are often pain free, while other patients with severe pain often have limited structural changes on MRI of the spine.



Once we find problems on MRI, our traditional medical-care system often likes to operate on that area. Let's look a bit more at whether that's a good or bad idea.

Which Joint or Disc Is Causing the Pain?



Once a disc or facet joint is seen on MRI to have problems, we still don't know if it's causing the pain. While in joints like the fingers, it's easy to see which joint likely is causing the pain, other joints don't lend themselves to an exam.

Take, for example, the hip and the sacroiliac (SI) joint. Both can refer pain to the hip/groin area. It's very hard

to get your fingers near the hip joint as it's buried in inches of muscle. As a doctor, you can feel the back of the SI joint, but most of it is buried inside the pelvis. How do you know which one is causing what the patient reports as "hip" pain? The only way to know for sure is to "block" one or the other (inject numbing medicine to kill the pain from one). Since the accuracy of the injection is important (meaning if the doctor injects the numbing medicine into the muscles instead of the joint, you're no closer to solving the mystery), fluoroscopy (real-time X-ray imaging) or ultrasound are used to place the medication. With the facet joints in the spine, it's also difficult to tell which is the "pain generator" (the joint causing the pain).

One word of caution. Many physicians use an anesthetic known as Marcaine, or bupivacaine, to numb joints for diagnostic purposes because it lasts longer than other anesthetics. However, this anesthetic has been shown to damage cartilage, and the <u>American Academy of Orthopaedic Surgeons has warned their members not to use</u> it. Therefore, you need to ask which numbing medicine will be used. <u>If it's Marcaine, or</u> <u>bupivacaine, ask the doctor to substitute 0.25-0.5% ropivacaine as this has been shown to</u> <u>be more cartilage friendly.</u> However, also realize that this numbing medicine is often only found in maternity wards unless the doctor special orders it, so he or she may have never used it. Lidocaine, or Xylocaine, is another anesthetic that can be used. While it has less cartilage toxicity than Marcaine, it still has some negative effects.

Are There Joints in Your Spine?

Your spine has little joints about the size of a finger joint called facets. There are two at each spinal level (one on each side), which translates to 14 facet joints in your neck, 24 in your thoracic spine, and 10 in your low back. These joints limit and control motion, so, for example, this is one reason you can't turn your head around 360 degrees in Exorcist style (the cervical facets stop you at about 90 degrees of rotation).

O Regenexx®

One of the things to know about these joints is that they can be injured through trauma or get arthritis with wear and tear and cause pain and wreak havoc. For example, when they're painful, they frequently shut down the local stabilizer muscles, causing instability and more wear and tear at that level. They're also good at referring pain to various areas. Look at the facet pain-referral map, to the right, that shows where the various neck joints can project their pain. For example, if you have pain in a C2-C3 cervical facet (near the top of your neck), it will cause headaches. A C6-C7 facet joint near the bottom of your neck will cause shoulder-blade pain. Finally, a lowerback facet can also cause pain down the leg.



FIGURE 2

Pain Referral Patterns from Cervical C2-3 through C6-7 Facet Joint Injections. Shaded areas indicate areas of pain experienced by asymptomatic volunteers after injection of facet joints C2-3 through c6-7. (From Dwyer AB, Aprill C, Bogduk N. Cervical zygapophyseal joint pain patterns. I: A study in normal volunteers. Spine 1990; 15:453-457.)

Because these joints are buried deep in the spine, unlike a painful finger joint, they can be difficult for the doctor to feel. As a result, finding out which one is hurting and in need of regenerative therapies to help the painful joint can oftentimes be difficult through an exam. As a result, the doctor may use a diagnostic block to help decide which joint hurts. As above, this is just numbing the suspected joint to make sure your pain goes away.

Another common joint that can cause pain is the sacroiliac. This is the joint between the tailbone and the back of the hip. It transfers forces from the legs through the tailbone and into the spine. The joint can be injured and/or get arthritis, and it refers pain many places, including the groin, the back of the hip, or even down the leg to the thigh or ankle. The strong ligaments that hold the SI joint together and stabilize it can get loose and sloppy, leading to too much SI joint movement, which can cause pain and joint damage.

Radiofrequency: Nuking the Spine-Joint Nerves to Take Away Pain

Radiofrequency ablation has become a common way to treat painful or injured neck and back facet joints. The nerve that takes pain signals from the joint is burned away using a needle with a tip that heats up. Based on the published research, this can be an effective way to help pain for more than a year in the neck and about six to nine months in the low back. However, these nerves are there for a reason—they supply the stabilizer muscles, which then become atrophied and weak. Hence, "nuking" the nerve should only be done when there's no other good option for treatment for the patient. We've had success in treating many patients with platelet or stem cell injections into these joints, which doesn't impact the facet-joint nerves.



There's another big problem with frying the nerves to these critical joints. These nerves take pain from the joints. Hence, injured joints that can't transmit pain can injure themselves further. In medicine, this is called a "Charcot joint."

Joints in the Upper Back: Where the Ribs Attach to the Spine

In many patients who fall or get hit in a car crash, we see pain at specific spots of their upper back or chest. Oftentimes, the ribs as a cause get completely overlooked, other than a vague diagnosis of costochondritis (inflammation of the ribs) and a script for anti-inflammatories. Is there more to this problem?



The ribs attach to the spine at two joint complexes each, one in the back and one in the front. These ribs move when you take a deep breath by expanding out like a bucket handle being lifted. They move from these pivot points in the back and the front. At these spots, strong ligaments connect the ribs to the spine and the breastbone (sternum). The most frequent type of injury we see is when the metal stiffeners in the back of car seats (meant to prevent the collapse of the seat back) hit these rib anchor spots in a rear-end car crash. This often loosens the ligaments that hold the ribs, causing spot pain in the back that gets worse with breathing or twisting. The front rib attachments can get injured by the

airbag or the chest wall hitting something, causing similar pain areas. Another common cause of front rib pain is open heart surgery, where the rib joints get injured once the chest is "cracked." We usually treat these issues by tightening these ligaments with injections.

The Disc!!!

Ever since Dr. Mixter and Dr. Barr first discovered in the 1940s that the spine discs could herniate and cause nerve problems, the disc has rampaged through spine care like Godzilla knocking down buildings in Tokyo. It has devoured everything else in its path, marginalizing our knowledge of other parts of the spine simply because surgeons could operate on discs and there were insurance codes that paid good



money when they did so. So while the disc may or may not be important in why you hurt, we'll attempt to put the beast in its place in this chapter and reduce the disc hype, as, after all, the disc is just one part of the spine.



The first thing you need to understand about the disc is how it's made and what it does. It's basically a shock absorber between the spine bones that also allows motion. It has a soft inner gel (nucleus pulposus) surrounded by a tough outer fibrous covering (annulus fibrosis). As we age, the cells inside the disc begin producing less of a chemical that can hold onto water. This means that older discs can become dried out and rather inhospitable places for cells to live. Add to this that the stability of the spinal segment around the disc begins to get worse and worse due to muscle atrophy and loose ligaments, and the disc can get beat up. Tears can develop in the outer covering.

Knowing The Kind of Disc Problem You Have Is Key to Finding Regenerative Solutions

While we'll discuss things like platelets and stem cells later on in this chapter, you can learn a lot about discs by looking at what can go wrong and then what might help. We categorize "bad discs" into four different categories:

- 1. Herniated
- 2. Bulging
- 3. Torn
- 4. Degenerated



I've put together the infographic here to help you figure out which type of disc problem you have. Click on the image to see a more detailed PDF.



A *herniated disc* is when the outer covering of the structure breaks open letting the inner gel herniate out. A *bulging disc* is when the outer covering doesn't completely break open but the fibers are stretched and weak, leading the inner gel to cause bulging of the covering of the disc. A *torn disc* is when the outer covering gets a tear that doesn't result in a herniation or bulge but does cause pain due to ingrown nerves or the disc leaking nasty chemicals on the associated spinal nerve. Finally, a *degenerated disc* is one that has collapsed due to few living cells inside, causing a lack of production of the chemicals that usually plump up the disc. As the disc collapses, the whole segment gets unstable and sloppy, causing the body to grow bone spurs to stabilize the area. These bone spurs can press on nerves and cause pain.

Which Do You Have?

A great way to figure this out is to learn how to read a low-back MRI report. Here are the terms that a radiologist will use to describe each type:

- 1. Herniated—extrusion, extruded, herniated, herniated nucleus pulposus, disc herniation, disc extrusion, free fragment, sequestered fragment, sequestered
- 2. Bulging—protrusion, protruded, bulging, disc bulge, bulging disc, broad-based disc bulge
- 3. Torn—annular tear, high intensity zone, HIZ, annular fissure
- 4. Degenerated—degenerative disc disease, DDD, degenerative disc, spondylosis, decreased disc height, osteophyte(s)

Knowing Your Disc-Injury Type Is Important to Determining Which Regenerative Medicine Solution Will Work

These days, there are physicians who are offering to inject platelet rich plasma (PRP) or stem cells into low-back discs. This is all very promising technology, but, regrettably, many physicians aren't too choosy about which types of discs they treat.

Why is injecting the disc a big deal? If you need a disc injection, it's much less invasive than surgery; however, injecting the disc is never trivial. Why? The biggest issue is that a disc infection is a huge problem. It's called discitis, and while rare, if it happens, it means IV antibiotics and major surgery. Hence, if the disc injection can be avoided and the same results obtained, that's usually the way to go.



We've been injecting stem cells and PRP into discs since 2005 and were the first in the world to do that work, so we've developed experience with what works for which disc type:

1. Herniated—It's usually best to stick with a technology that doesn't inject the disc but, instead, instills isolated growth factors around the irritated nerve. This is called platelet lysate, an injection packed with natural growth factors. Why? Because an epidural injection around the nerves is less invasive than a disc injection. The focus is to <u>ramp up your body's</u> <u>own natural repair mechanisms</u>, which can get rid of the herniated disc.

2. Bulging—PRP or same-day stem cell injections into a disc have yet to demonstrate that they can get rid of disc bulges. The only technology we've seen reliably do that is <u>specially</u> <u>cultured stem cells injected in a specific fashion</u> (i.e., not same-day stem cells but ones that are grown to bigger numbers under special conditions for a few weeks). In fact, most of these patients can be managed by the above type of epidural platelet lysate injections without ever injecting the disc itself.

3. Torn—This is really the big "killer app" for stem cell and PRP disc injections. In our clinical experience, a painful disc tear can be treated reliably with a same-day stem cell injection into the disc. <u>MRIs from before and after these treatments frequently show</u> evidence that the disc tear has healed.

4. Degenerated—Regrettably, this type of disc problem doesn't respond to any type of stem cell treatment, including specially cultured stem cells. However, in our experience, it can respond to a treatment protocol focused on platelet and/or stem cell technology deployed to treat sloppy disc movement due to lax ligaments and arthritic facet joints. Click on the image below to see more information on this type of injection.





Spine Regenerative Medicine Traps

Regenerative medicine for the spine can be really confusing for patients. When we began regenerative spine care using stem cells and platelets in 2005, there were no other options and there was no textbook. We invented all of our own therapies, and we figured out what worked and what didn't through trial and error. However, there are so many options now.

One of the more interesting "traps" out there is that it's possible to regrow a degenerated disc that has collapsed and is "bone on bone" by placing platelet rich plasma, stem cells, or fibrin injections into the disc. Regrettably, it's not possible. Having said that, as discussed above, this condition can usually be managed very well by treating much more than the disc!

A good example of one of the treatments out there that's being hyped is fibrin injections into discs. You could really insert anything here, however, as what's being promised with fibrin injections is similar to what many doctors suggest will happen with other technologies like stem cells—a regrown disc from a collapsed and trashed one. Is this true? No!

What is a fibrin disc injection? Some physicians, based on the teaching of a Texas doctor, are now beginning to try to seal degenerated and torn discs by injecting fibrin glue. The type of fibrin they're using is sort of like natural rubber cement. The theory is that this glue will "seal" the small tears and holes in the disc, preventing stuff from leaking out. The reason given for doing this is that sometimes discs can leak chemicals, which can cause inflammation and swelling around the spinal nerve that passes by the disc (a.k.a. chemical radiculitis), leading to back or leg symptoms. Hence, advocates for this procedure contend that sealing the disc prevents that from happening. In addition, they argue, if you're going to be injecting stem cells inside a disc, sealing it first will keep those cells in that structure.

The before and after images of injecting really anything that gels up in the disc are impressive, whether that be fibrin, a same-day bone marrow stem cell procedure, or platelet rich plasma. Basically, the disc plumps up on X-ray images. This is because all of these things contain fibrinogen, which is what causes your blood to clot. This protein causes fibrin to be laid down, which is what gives a blood clot its structure and mojo. **However, there's little evidence that a severely degenerated disc that has lost its height stays at the new taller configuration.** Basically, once the new fibrin is dismantled by the body (which takes one to two weeks), the disc returns to its collapsed configuration. So the first thing to be cautious about is the X-rays showing the restoration of normal disc height immediately after a fibrin or other injection, as the height doesn't last!



I've written before about the <u>Pauza Discseel™ procedure</u>, having found many inconsistencies between what the website suggested and reality. The biggest was that while the website would make you believe a randomized controlled trial (RCT) had been performed that supported that the procedure was effective, in fact the only RCT performed showed that the procedure was no better than a placebo shot. In addition, the website reasons that you need to seal the disc to keep injected stem cells inside because stem cells can cause cancer. No research shows this, and the website erroneously points to a "respected Johns Hopkins study" without any such study on this topic (stem cells injected causing cancer) actually existing. In fact, our recent 2,372-patient stem-cell-safety study included patients with stem cells injected into their discs, and no such issues were found.

A good example of how confusing information on regenerative medicine can be for patients researching their options—there is a claim that states that stem cells injected into the disc can cause bone spurs to form (if they leak out). The site states that this is, therefore, one pressing rationale to seal the disc before injecting stem cells. This claim has issues on many levels, not the least of which is that the disc will only stay sealed for about a week to 10 days. However, like many things said on the fibrin disc-sealant site, there's what the words superficially suggest and then reality once you take a peek under the covers.

In this case of bone spurs, the website refers to a rabbit study where the authors tried hard to get the stem cells to make bone. Cells were cultured in a special media to force them to make bone and were not the treated rabbit's own cells, and the injury to the disc was not one that in any way replicates the real human degenerative disc disease condition. In fact, the stab model used to injure the disc before the stem cells were injected is about the same as taking a ball-point-pen-sized implement and jabbing it into your disc and sucking out all of the living cells. So unless you've been recently abducted by aliens and had your disc sucked out, the starting point for this experiment was nothing like your degenerated disc! In conclusion, nothing about this study is in any way like the same-day stem-cell-based disc injections that patients are receiving for torn and painful discs.

To illustrate why sealing the disc with fibrin is not needed, a great example is one of our patients. Ted had a multiyear history of low-back pain and had failed physical therapy and epidural steroid injections. He was so disabled that he couldn't throw a Frisbee with his kids. His MRI had a tear in the back of the disc, and his discogram (a test where dye is injected into the disc to see if it's painful) showed that the L5–S1 disc was painful. In addition, that test showed L5–S1 was a leaky disc in that the dye squirted right out. The L4–L5 disc was also leaking, but it wasn't painful.



We injected the patient's own stem cells using our patented Regenexx same-day stem cell procedure in May of 2015, and he reported 75-80% improvement. Since he wanted more, the disc was injected a second time in November of 2015. In both instances, the L5-S1 disc was injected with his own stem cells (and NOT sealed with fibrin). The patient did well and was just seen this week for his one-year follow-up. He's now reporting a 95% improvement! He's back to all activities and without pain, and never once did we entertain the idea that we needed to first seal his leaky low-back disc.

Why did this work so well if the disc was leaking? As discussed above, a same-day bone marrow stem cell procedure has its own natural fibrinogen from the patient's own body. It seals the disc all by itself. Unlike fibrin glue, which has been shown to be a very poor stem cell scaffold because it's too dense, bone marrow sets up its own natural scaffold for stem cells immediately upon detecting exposed collagen fibers, like those seen inside a torn disc.

Is there a very real downside of injectable fibrin? A few years back, we saw a woman who was a chronic dural leaker. This meant that the covering of her spinal cord (the dura) that was supposed to hold back cerebrospinal fluid (the liquid in which the brain, spinal cord, and nerve roots are suspended) would spontaneously spring leaks. She had been treated with the same type of fibrin that doctors are now using to seal the disc. There was just one little problem. Despite the product being FDA approved for being safe and free of communicable disease, she had been injected with a bad batch made from a blood donation of a patient with hepatitis C. After that therapy, she had developed the disease. While we were able to help her, using highly concentrated platelet rich plasma injections instead of fibrin, she serves as a very real reminder that there's nothing like the power of your own tissues, as even FDA-approved biologic tissues have a false-negative rate when screening disease. In addition, while this is a rare event, it's not rare if it happens to you.

So is there a pressing need for fibrin treatment for back pain to seal the disc before injecting stem cells? Not really. Certainly nothing on the fibrin disc-sealant site sounding alarm bells of why sealing the disc can prevent catastrophe is remotely credible. In addition, cases like Ted's show us that despite having a severely leaky and painful disc, without fibrin being injected, the right patients with a torn and painful disc do great. Finally, our dural leaker patient is a stark reminder that using your own tissue to heal your body should always be your first choice!



Surgery: Helpful Pain Relief, or a Slippery Slope to More-Rapid Arthritis?

Often with only a cursory exam and a picture (which you now know may not show the cause of pain), our modern orthopedic-care system schedules spine surgery.

You would think that we have a large body of scientific evidence that common orthopedic spine surgeries are effective. In fact, we don't have high levels of scientific evidence that any type of low-back or neck surgery is effective. While a few patients really need spine surgery, most who get operated on would do better without it. So why does your insurance cover this lunacy? Because of tradition. Yep, this is the silly reason why surgeons put so many patients through these barbaric procedures.



Is there a better way? Rather than cutting out tissue, our practice was the first in the United States to pioneer a new approach, starting in 2005. Rather than surgically removing or repairing the tissues, we began using the patient's own stem cells in an attempt to heal damaged tissues. There's even a nice little video that helps explain why we think this approach is better. Click on the image to play that video.

Understanding the Body's Repairmen: Stem Cells

Remember that house in your neighborhood that was inhabited by an older person who couldn't keep up with the maintenance? We'd all accept at face value that a house left unattended for years will weather and begin to slowly degrade and fall apart. Our joints and bodies are the same. A quick run around the block, a workout in the gym, or just daily use will cause microdamage in any number of tissues. Left unrepaired, these areas will begin to break down over time (just like the unattended house down the street).





So what keeps us from falling apart after just a few years? The figure to the left tells the story of the opposing forces of damage versus repair. Everything we do every day adds small (or large) amounts of damage or wear and tear on our tissues. On the other side of that coin is

repair. This is the mechanism that fixes the damage. When these two systems are equally matched (our repair system can easily keep up with the damage), you have healthy joints.



Older or Unhealthy Joint: Wear and tear exceeds repair abilities.

Turns out we have billions to trillions of tiny little repairmen in all the tissues of our body. These repairmen are called <u>adult stem cells</u>. As an example, consider an adult stem cell type called a <u>mesenchymal stem cell (MSC)</u>. These cells live in your tissues and are called into action once damage is detected. They can act as a general contractor in the repair response, giving signals to activate other subcontractor cell types that are needed for the repair job or even firing (deactivating) cells that may be causing trouble. They can also "differentiate" (turn into) the final cell type needed for the repair. For example, if the cells are repairing the cartilage of your knee, they can differentiate into these cartilage cells. When we're young, while

there may be a lot of abuse on the body, in general, the amount of repair capability (adult stem cell numbers and function) generally far exceeds the amount of damage we can inflict. <u>As we age, we have fewer of these stem cells around.</u> Even when we're younger, an area can become injured so that it doesn't allow the repairmen in the door (less blood flow, or there just aren't enough cells to affect a proper repair). At this point, the amount of damage starts to exceed the body's ability to repair.

What if we could turn that equation around? What if, despite being older, or even younger with an area that has too much damage for the local repair cells to handle, we could amplify repair in the area? As you might have guessed, this is a basic tenant of Orthopedics 2.0. The doctor's job is to increase the local repair response in nonhealing tissues so that it exceeds the existing damage or wear and tear on the area. This also includes the other side of the coin—the doctor should avoid prescribing or injecting medication that will harm or slow this repair process. In addition, the final part of the doctor's job is to reduce the local damage on the area. How is this done?



Improving the Repair Response

We can increase the repair response by dividing it into approaches with three levels of sophistication.

Level I: Microinjury

Level II: Improving the Healing Environment

Level III: Stem Cells

Level I: Microinjury



Ever since ancient times, creating a small injury to prompt healing has been seen as a good idea. For horses, this was called <u>"pin firing."</u> The technique was to take a hot poker and place it into a nonhealing ligament to cause small amounts of damage to the area, which caused the body to kick up a repair response. While barbaric, it generally worked. For centuries doctors have created small injuries in a nonhealing wound by "roughing" up the tissues.

Physicians still use this concept today for tendons, ligaments, and joint capsules. For example, in a <u>shoulder capsulorrhaphy</u>, a surgeon usually

inserts a small catheter that heats the tissue to prompt healing in a damaged shoulder capsule (the covering of the shoulder joint that helps control motion). Doctors still score ligaments with scalpels and beat up tendons with a needle (percutaneous tenotomy), all to prompt a healing response. Another example is microfracture surgery, which is a procedure used to treat a hole in the cartilage in a damaged joint. In this surgery, the doctor pokes holes in the bone to cause the cartilage to heal. Finally, the procedure known as prolotherapy is in this same category. In this procedure, rather than a mechanical injury being initiated, the physician injects a chemical irritant to cause a chemical microinjury. All of these types of treatment rely on the same concept—that we usually get only one bite of the healing "apple," and if something fails to heal completely the first time, we can create more bites at that apple simply by causing a small injury to the area.

The big advantage to microinjury techniques is that these basic procedures are simple and often inexpensive. The downside is that while many times they work well, sometimes they don't have enough oomph to produce the right type of healing or enough healing. In addition, they also tend to do better when fibrous tissue repair is what's needed. An example of this type of tissue in the spine is the ligaments I showed you in red a few pages back.



The Original Regenerative Injection Technique—Prolotherapy

Prolotherapy is an injection method where chemicals are injected to cause a small inflammatory healing reaction. In the 1940s, this was a mainstream orthopedic procedure used to treat lax ligaments and spinal pain. Heck, it even had its own pharmaceutical (Sanusol). However, in the next half of the twentieth century, prolotherapy fell out of favor. Why? Some say it was linked to the bad outcome of a single injection placed where it shouldn't be in the spinal canal. However, others place prolotherapy's demise on the fact that it had no sustainable medical business model. It was simply replaced by big surgical procedures that were far sexier and had better reimbursement through a new concept at the time—employer-sponsored medical insurance.

While we may never know what happened, the procedure was revived in the 1980s, and over the past two decades, I've seen this simple and inexpensive technique work for patients who otherwise would not have been helped. I've published on prolotherapy's ability to tighten loose spine ligaments simply through injection, and others have published on the same observation in lax knee ligaments.

If you have a loose spine ligament, prolotherapy may be a good option. However, like any regenerative injection technique, accuracy of placement matters. As a result, prolotherapy should almost always be performed under imaging guidance, especially when deep specific structures are being treated. <u>I wrote an article on this topic a few years back</u> that explains how this works.

Level II: Improving the Healing Environment

The next level of sophistication beyond just creating a healing microinjury is making the conditions in the area more conducive to healing, or <u>"anabolic."</u> You may have heard this term associated with body builders who use steroids. This is not the same use here, although body builders "build" muscle, so this is why they use "anabolic" steroids (literally "steroids that build"). Here the term means making an area pro-repair, or better able to heal.







Creating an anabolic healing environment is not a new concept in medicine and surgery. For centuries, physicians have known that some people have better innate abilities to heal, while others have fewer healing capabilities. The acronym PPP (piss-poor protoplasm) was used in my medical school training to mean a patient who, due to disease or extreme old age, was unable to heal after surgery. While surgeons have always known that some patients could have a compromised ability to heal, not much attention has been paid in how to make routine and otherwise healthy patients heal better. Doctors have always understood the basics, like good nutrition, young age, high levels of fitness, and good blood supply. About 20 years ago, that started to change in the dental community. Some dentists began experimenting with a simple concoction called <u>PRP (platelet rich plasma)</u>. The dentists used this stuff made from their patients' own blood to help dental implants heal.

PRP is a simple example of how we can improve the healing environment. Your blood has platelets that contain growth factors that help to ramp up healing. To understand how these platelets work, a paper cut will illustrate the basic points. When we cut ourselves, we bleed into the cut. The blood coagulates because of cell fragments that live in our blood called platelets. The job of the platelets doesn't stop there. They go on to release certain growth factors that stimulate local cells to heal the cut.

<u>Growth factors</u> are like espresso shots for cells. A cell works at a certain pace to do its job. If we add growth factors (like those in PRP), it's like buying all of the cells trying to repair the area a bunch of Starbucks gift cards. The cells react to the growth factors like people react to triple espresso shots: they work harder and faster. So if we use an example of a construction site where we have a few bricklayers building a new wall and we add growth factors (espresso shots), our bricklayers will build our wall faster.



As you might have guessed, Orthopedics 2.0 uses these same

concepts to promote healing. The most basic level-II procedure today is PRP, which can be mixed up from a patient blood sample in a bedside centrifuge or more preferably in a simple hospital- or clinic-based lab. PRP means that the healing platelets have been concentrated. Injecting the patient's own blood can often accomplish the same thing as it's also rich in platelets.



Not All PRP Is Created Equal

One of our focus areas since 2005 has been figuring out how to use various forms of PRP to get stem cells to grow better. This extensive experience has led us to understand that there are different "flavors" of PRP and that some of them seem to work better for kicking stem cells into high gear. How can you tell the difference? Look at the color of the PRP. Based on our lab data, red PRP doesn't work as well to promote stem cell activity as amber PRP. Most automatic bedside centrifuges used by doctors today produce this red, bloody PRP.



Why should I care? If the PRP you're being injected with is red, then it has far too many red and likely white blood cells. This will not only cause more unneeded inflammation, but it also inhibits the response of your local stem cells. You want the platelet prep being used to do the opposite-stimulate your own local stem cells! As a result, the injection should be amber in color, How do you get the better amber PRP? Make it by hand in a hood.

Why is red PRP a problem? It's rich in red and white blood cells. When it comes to energizing stem cells toward repairing more tissue, our lab experiments show that red PRP doesn't have the same "espresso shot" kick as PRP without red and white cells (amber PRP). Based on these experiments, we have created what we call a "super concentrated platelet" procedure (SCP) to maximally stimulate stem cells into action. <u>Click</u> here for an infographic that explains

<u>the issues and lab data in more detail.</u> In addition, click on the video link below to see a two-minute animation that explains these differences.



The other big difference with SCP is that since we create it by hand for every patient in a lab rather than mass manufacturing it in an automated push-button centrifuge, we have much more control over the composition of the final preparation. In particular, while most bedside cetrifuges can only concentrate to about 5-7 times more platelets than are normally present in the blood, we can concentrate SCP all the way

up to 40 times over baseline! Is getting to higher platelet concentrations better? Yes, with this type of PRP, we see more activity in local stem cells with higher concentrations of platelets. For this lab data, click here.

In our clinic, we also use next-generation level-II tools beyond PRP. These tools include platelet lysate (PL). In the case of PL, our advanced cell biology lab makes PRP from the patient's blood and then breaks open the platelets to make all of the growth factors immediately available. The difference between PRP (or SCP as we call it) and PL is the



same as between a time-release pill and an immediate-release pill. PRP has whole platelets that release their growth factors over time. PL has all of the growth factors immediately available. Based on our experience, there are specific reasons to use one or the other. For example, in our clinical experience, PL is excellent to use around nerves.

We can also produce a newer type of platelet lysate that we call PL-M and cPL-M. In our lab experiments, we noticed that despite blowing up platelet bodies and releasing growth factors, there were still some whole platelets left. This meant that there were still growth factors to be released. As a result, we developed a proprietary type of triple lysate that gets all of the available growth factors out of the platelets.

Level III: Stem Cell Therapy, or Adding in the General Contractors of the Body

While level-I therapy is about causing a little injury to prompt healing and level II about getting the local cells to work harder, level III is about adding more workers to the area. Staying with our construction-site metaphor, a general contractor (GC) is the person who pulls a construction project together. He or she hires subcontractors, like plumbers, carpenters, and electricians. Does your body have a GC cell that can help coordinate its daily repair jobs?

Yes. The GCs of your body are <u>stem cells</u>. So level-III advanced techniques use concentrated or cultured stem cells to help repair tissues. There are a number of different types of stem cells. We've all heard of <u>embryonic stem</u> <u>cells</u> that are taken from a growing embryo. While these cells are very potent stem cells, they also <u>have the nasty</u> <u>habit of forming tumors</u>. To better understand why that happens, click on my video link to the right.





Cells can also be taken from <u>cord-blood stem cells</u> or <u>adult stem cells</u>. However, while some of these cell types might be appropriate as last-ditch efforts to save someone's life, their risk of transmitting genetic disease makes them too risky for orthopedic applications. <u>As an example, in one study, an older</u> <u>rat bred to have osteoporosis donated stem cells to a</u> <u>young rat without the disease</u>. The young rat acquired osteoporosis in the bargain! Since we currently don't

possess the technology to screen donors for all inheritable diseases, the risk of using someone else's stem cells is too high for now (in my opinion).





Finally, the newest stem cell types are called induced pluripotent stem cells (iPS or iPSCs). While all of this sounds very daunting, the basic concept is that these are artificial stem cells that don't exist in nature, created from natural cells. These are created by heavy-handed genetic manipulation of the normal cells from the body (like skin cells) or by exposing these same cells to a stimulus (like putting them in an acid) and are likewise very dangerous until proven otherwise. We're much more than a decade away from seeing iPS cells used in real everyday patients. To learn more on iPS and STAP, click on the video link to the left.

Mesenchymal Stem Cells

There are many types of adult autologous stem cells, but for the purposes of this orthopedics discussion, one stands out as the best candidate for our general contractor position: the <u>mesenchymal stem cell (MSC)</u>. These cells are found in many tissues (for example, as above, they are found in bone marrow aspirate and fat). For orthopedic applications, their ability to help coordinate the repair response as well as turn into cartilage, bone, tendon, muscle, and ligament make them ideal. Other cells, such as <u>very</u> <u>small embryonic-like</u> or <u>embryonic-like stem cells</u> (VSELs or ELSCs), are also promising for orthopedic use, but not enough research has been done yet on these adult stem cell types with regard to safety to make them practical for everyday use. Also realize that there are <u>likely hundreds of classes and subclasses of adult stem cells</u> that will eventually be used for therapy. Many of these may even be combined with mixtures of other non-stem cells or tissue-engineered scaffolds to better promote healing.





Fat vs. Bone Marrow Stem Cells

Anyone perusing the Internet can see that there are two stem cell types from the same patient (autologous is the term for this) that seem ubiquitous: fat and bone marrow stem cells. When we began using stem cells in 2005, we investigated which of these we should use. At the time, there was mounting evidence that bone marrow stem cells had real utility in orthopedic applications and very little data published showing that fat stem cells were very helpful. What's amazing is that this data isn't all that different today.

Before delving into that research, let's review the five most common procedures being offered in autologous stem cells for orthopedic injuries (illustration above).



As you can see above, there are two different bone marrow procedures (same day and advanced) and three different fat procedures (two same day and one advanced). The same-day bone marrow procedure is equivalent to the Regenexx-SD procedure, and the advanced bone marrow procedure is Regenexx-C¹ where cells are cultured for two weeks. On the fat stem cell side, there's a simple adipose fat graft and an isolated stem cells procedure that are equivalent to the Regenexx-AD procedure. Finally, just like in the bone marrow procedures, you can culture the cells to get more, so there is an adipose advanced procedure.

When compared head to head, how many research papers show that bone marrow or adipose stem cells are better than the other at making cartilage or it's components?



US National Library of Medicine Search searched 5/4/15 using terms "adipose bone marrow mesenchymal stem cell chondrogenesis". End date of search was 6/21/07. Only papers showing head to head quantitative chondrogenesis considered. For example, papers that used gene expression differences, ultrastructural comparison of the cartilage repair produced in-vivo, or collagen type comparisons in micro mass.

So now that you know what's being offered, can research guide us as to which is better for orthopedic use—bone marrow or fat?

How about when the two stem cell types are compared head-to-head in the lab for producing new cartilage (like that found in the facet joints)? Thirteen papers show that bone marrow stem cells are better at producing cartilage then fat stem cells, while none show that fat stem cells are better than bone marrow.

¹ The Regenexx-C cultured stem cell procedure is only offered through independently owned and operated medical services providers operating exclusively outside the United States. These service providers are not part of or affiliated with the Centeno-Schultz Clinic or any US Regenexx Network provider. The Regenexx-C procedure is not approved by the FDA for use in the United States.



How about a comparison of how the cells are harvested? Bone marrow stem cells are taken from the patient using a bone marrow aspiration. This is where a small hole is poked through the bone with a special needle. Fat stem cells are harvested during liposuction where fat is collected through a cannula. As you can see from the graph to the right, the risk of liposuction is about 21 times more than that of a bone marrow aspirate. Again, on the safety of getting the cells from the patient, bone marrow stem cells win again.

Regenexx®

In summary, the weak data I found in 2005 on fat stem cells hasn't really gotten much stronger. This is especially true for the use of fat stem cells in the spine. Bone marrow stem cells are still our first choice for orthopedic tissues. Might that change with time? It may, but for now, we'll go with what the data says is likely to be better for our patients.

A New Type of "Stem Cell" Emerges: Placental or Amniotic Tissue

Amniotic fluid and placental tissues surround a baby growing in the womb. When the mother's water breaks, this is the fluid that ends up on the floor. The placenta is the part of the baby's sac that attaches it to the mother's uterine wall.

Amniotic and placental tissues from the birth sac have been used for a century or more as a filler for the damaged covering of the spinal cord and in eye procedures. Way back when, a surgeon would make a visit to the OB ward and pick up some of this stuff to sew into a defect. In the 1970s when FDA tissue regulations went into place, companies could



make a little money selling these tissues to hospitals and surgeons. About five years ago, some smart businessman got the bright idea to begin marketing placental tissues as an injectable regenerative medicine product. They took the birth sac and freeze dried it, chopped it up into very fine pieces, and put the powder in a bottle. <u>L</u> <u>recall one of these companies who had incredibly aggressive sales reps and tactics</u>. The companies then went to medical conferences to convince doctors to inject it for problems like tennis elbow.

Now there wasn't a shred of evidence that this stuff would help tendinitis or any other orthopedic condition, but that didn't matter—it was a "regenerative tissue." The sales were pretty good, but at some point some smart sales rep got the idea that if he told doctors that this was a "stem cell" product, that would sell more vials of the stuff. Sure enough, sales went through the roof! Never mind that freeze drying, processing, and gamma irradiation sterilization of the living membrane made sure it had no living cells of any type, let alone stem cells.

Most doctors, who were new to the concept of stem cells anyway, bought this marketing "little white lie" hook, line, and sinker. Then came amniotic/placental fluid, which was marketed the same way. This was perhaps more believable as a stem cell product, and you can find research that when it surrounds the baby in the womb, this fluid does contain some stem cells. After I was told by a knowledgeable sales rep about the rich stem cell content of amniotic fluid, we decided to test these claims.

We check all manufacturers' and reps' claims before we use products. Unlike other medical groups who frequently take payments from outside companies to perform research or be paid representatives for products, we do this at our expense so there are no potential conflicts (e.g., a manufacturer that funds a study). Also, unlike any other medical group, we also have a research lab outfitted like any university research lab in the country, so we have the ability to run these tests and separate fact from fiction.

First, as discussed, the doctors hawking this stuff buy vials of product from sales reps. So you need to determine which product the doctor is using. Second, if you go to the website of <u>any company selling amniotic powder or fluid</u>, you won't see a word about stem cells. Why? These website claims are heavily regulated, and a single claim that there are stem cells in this stuff would result in the tissue being classified as a drug product and taken off the market to undergo a decade of testing costing hundreds of millions of dollars. This is in contrast to the inexpensive FDA tissue registration required if what's being sold has no living cells. However, what the sales reps tell physicians behind closed doors is another thing. In that world, reps meet physicians who would love to "get into stem cells" but don't want the hassle of buying equipment, learning and mastering a harvest procedure, and



dedicating the staff to run the machine that isolates the stem cells. They want something that will sit on the shelf that contains stem cells that they can use when the need arises. The problem, as we found out, is that none of this stuff actually contains stem cells.

Once our lab found no living tissue in any "stem cell product" tested, the <u>Interventional</u> <u>Orthopedics Foundation did an independent assessment of these amniotic products</u> that was presented at the IOF 2015 conference. Their research also found that these products didn't contain any living tissue, let alone stem cells! So the physicians offering this therapy are injecting dead tissue and hawking it as a live stem cell therapy. So is this deliberate fraud or just a physician who doesn't know what he or she doesn't know? Most of the doctors I've met and educated about this scam had no idea that what an orthopedic sales rep was selling them was dead tissue. Hence, it's likely a doctor who bought what he or she thinks are living stem cells and who doesn't know enough basic cell biology to understand the likelihood that the sales pitch he or she heard was fiction. In addition, the doctor has no way to independently test whether these tissues contain live cells, so the doctor is wholly dependent on the sales pitch.

If you see a clinic offering amniotic, fetal, or placental stem cells, run! The doctor isn't educated enough in this field to know what he or she doesn't know, which is never a good thing for patients. If the doctor does know what he or she is injecting and is still calling it a stem cell therapy, then that's consumer fraud, which is also not a good thing for patients.

Does Where You Get Your Bone Marrow Matter?

First, when most patients hear about a bone marrow procedure, they think of a different procedure than the one we perform (a bone marrow biopsy is the one you've heard about, while we perform a bone marrow aspiration [BMA]). A BMA is a simple procedure where the area is thoroughly numbed, and then a needle is gently worked through the bone to pull out what looks like thick blood. Is this really painful? In 2007, we polled our patients and found out that about 9 in 10 thought it was no big deal.

Second, where you take the marrow can make a big difference in the number of cells you get. For example, recently some doctors who aren't comfortable using more-advanced guidance techniques have begun taking it from below the knee rather than from the back of the hip. Others have begun to take it from the front of the hip. Regrettably, <u>this knee site doesn't produce as many cells as the hip site</u>; hence, we use the back of the hip procedure (at what's called the dimples of Venus or the PSIS) that yields more stem cells.



Different Orthopedic Stem Cell Procedures

As discussed above, there are two different bone marrow stem cell procedures—one is same-day and the other is cultured. What's the difference? The same-day procedure is what it sounds like. The stem cells are isolated and used the same day. The cultured procedure grows the stem cells to bigger numbers over a few weeks.

Almost all same-day procedures being used today are performed in automatic bedside machines that remove one fraction of the bone marrow that is rich in stem cells—the buffy coat. The issue with these machines is that they often can't be very exact, so they also isolate a lot of junk cells with the stem cells. As a result, we only perform this process of extracting the stem cells by hand so it can be done with more precision.

The buffy coat has both the MSCs, discussed above, as well as hematopoietic stem cells (HSCs). While the MSCs are good for helping to repair orthopedic tissues, in our clinical experience, the HSCs are good at bringing in new blood supply and treating muscle. This may be very important for tissues like the atrophied multifidus muscle in the spine. Both of these cell types make up the Regenexx-SD procedure, which we have used to successfully treat painful low-back disc tears and injured or arthritic joints, like the facets or SI joint.

Our dedication to lab research has also allowed us to advance this same-day procedure way beyond where it began. We discovered a second fraction in the bone marrow that's very rich in stem cells and is currently being discarded by everyone else. This layer has many more stem cells per unit volume than the buffy coat, and our lab research shows that they're fast growing and very useful for orthopedic purposes. As a result, we've increased the number of stem cells we can pull out of the bone marrow by a factor of five to seven by isolating this new fraction. When we compare how many stem cells we're able to isolate in the lab per unit volume to the number that automatic bedside machines can isolate, it's 15-20 times more! More recently, we have upgraded the isolation process we use a third time, again increasing stem cell yield.

Finally, we also have a cultured stem cell procedure offered through a licensed site in Grand Cayman (a British Commonwealth country with its own medical-practice laws). In this procedure, we can grow cells to bigger numbers over one to two weeks. In addition, for the spine, we can condition and select cells in culture using patent-pending methods to allow cells to survive the rough environment of the low-back disc. As discussed above, this special process, in our clinical experience, makes the cells perfect to treat disc bulges pressing on nerves.



How Do Stem Cells Do Their Thing?

While you've likely heard that stem cells are blankslate cells that can turn into another cell to replace a damaged cell, there's really a lot more to the stem cell story that just that neat little trick. Take, for example, the concept of exosomes. These tiny little packets are excreted by stem cells and can contain snippets of mRNA (an instruction sheet to make proteins). A stem cell can use these mRNA protein instruction sheets to task another cell to make proteins on its behalf. For a short one-minute video explanation on the topic, click on the video link to the right.



Stem cells can additionally work through what's called paracrine effects. This means that the stem cell excretes chemical messages called cytokines. Think of the stem cell as a general contractor who is involved in coordinating a repair of tissue. It barks out orders to hire subcontractor cells to do some of the work. While cells can't talk, their version is releasing very specific chemicals that can attract the types of cells it needs and then releasing other chemicals to give basic instructions to the other cells. For a short video that explains this, click on the video link above. One interesting study also explained another mechanism for how stem cells may help arthritic joints (for example

Degenerativ

Joint

The adult stem cells that live in your body can act as a general contractor that can coordinate subcontractor cells to repair damaged tissue.





The stem cells detect the macrophages and cartilage breakdown products and interact with the macrophages to "deactivate" them, preventing their ability to further destroy the joint. They also instruct them to release helpful growth factors. the facet joints in your spine). In the type of catabolic joints that I describe above, there are cells that act like Pac-Man by gobbling up normal, healthy cartilage (these cells are called activated macrophages). These cells are deactivated by mesenchymal stem cells and, as such, stop eating the good cartilage.

I've also done a short one-minute video on how this Pac-Man cell inhibition works. For that, click here.



Is There Any Research That PRP or Stem Cells Can Help Disc Problems?

To date, we have the most research that your own stem cells and PRP may help the disc. There's a great deal of animal modeling—meaning that when stem cells or PRP are injected into the disc, they seem to be able to help injured animal discs heal (<u>this link</u> leads to what's currently published about stem cells and discs in the US National Library of Medicine). However, regrowing animal discs seems easier than doing the same thing in humans.

The <u>highest-level study</u> we have on the use of PRP in patients with chronic back pain caused by a torn disc showed promising results. <u>Another study</u> using same-day stem cell injections in patients with mostly the same problem also showed good results, although the study is not as high quality as the first. <u>Yet another study</u> using someone else's stem cells to treat degenerated disc pain also showed encouraging results.

However, while all of these studies show good pain relief (which is what most patients are looking for), what they don't show is as important as what they do. While some of them may show some improvement on follow-up MRIs in disc tears when the discs haven't collapsed yet, what they don't show is that a severely degenerated and collapsed disc can be regrown like new.

We observe these same things as well when using these therapies clinically. They are very good at helping patients with torn and painful discs and in those cases can show some evidence of repairing the tear, but they are less effective in treating the severely degenerated discs.

A word of caution! While there is little evidence that these therapies can regrow boneon-bone degenerated discs, there are no shortage of doctors selling stem cell therapies willing to inject these discs. As discussed above, these discs do better with what we call our degenerative disc disease procedure, which focuses on all of the issues that have caused the problem and the pain.



Can We Do More with Specially Cultured Stem Cells?

In 2006, our clinic pioneered a new approach to stem cells, culturing them to grow more. In this advanced procedure, we can obtain about 20-50 times more stem cells than in our same-day procedure. This Regenexx-C method has been the subject of <u>several safety papers published in the US</u> <u>National Library of Medicine</u>. The cultured stem cell injections were much safer than the more invasive surgical procedures they helped patients avoid.

Where might we use cultured versus same-day stem cells? In our experience, for certain applications cultured stem cells are helpful, while for others there is only a small difference. For example, <u>in patients with a disc bulge pressing on a nerve, specially cultured stem cells injected into the bulge can reduce the size of the bulge.</u> Click on the infographic to the right to see the results of this procedure.

One big advantage to cultured cells is the ability to save them for future use in cryopreservation (deep freeze). In addition, since these cells are grown in culture to bigger numbers, we can often save many treatments for the same area or other future injuries.

Can Stem Cells Be Used Outside the Disc?

We've used stem cells in many places outside the disc. This includes treatments to tighten ligaments, help atrophied stabilizing muscles, and treat beat-up facet joints. In addition, what's unique about Regenexx is that we have many non-stem cell, platelet-based therapies focused on the spine. When we discuss nerves later in the book, in the "Neuromuscular" chapter, we'll discuss these platelet-based sciatica therapies in more detail.

Bone: The Structure of the Spine

Bone is the structure that gives form to the spine. Ligaments and muscles help keep the vertebrae aligned, and the discs act as their cushion. When the discs and facet joints get too much damage, they can suffer from arthritis, where the disc or cartilage breaks down, reducing the cushion. But can the bone develop issues, too?

While we think of bone as cement, it's actually like hard plastic that gives and absorbs shock. However, it relies on the disc to help it work properly as a machine to mitigate forces. When that disc is damaged, the bone can swell, and this is called "Modic changes."





Bone and Joint Tissues Are Alive!



Bone spurs inspire fear in the hearts and minds of most patients, kind of like The Blob from a bad 1960s horror movie. When patients hear that they have them, they're immediately freaked out. However, much like the title of Stanley Kubrick's '60s movie Dr. Strangelove or: How I Learned to Stop Worrying and Love the Bomb, patients shouldn't get immediately frightened when they hear they have bone spurs as most are there to provide stability to the spine and are an important part of you! Only every once in a while do these bone spurs that you should learn to love cause problems that you should fear.

Why do bone spurs develop? When are they good or bad? Bone is made up of mature cells <u>(osteoblasts)</u> and stem cells that react to their environment. It's well known, for instance, that when the cushioning disc in the spine wears out, the bone underneath the disc makes itself thicker to handle the new forces. We know that people who don't exercise or who pursue non-weight-bearing exercise have more brittle bones, and that people who lift heavy weights have more-dense bones. So bone is alive and quickly reacts to its environment. How quickly?

As an example, for many years most physicians were convinced that bone spurs in the spine took years to form. This was based on the theory that bone was dumb, inanimate cement. However, more-recent research shows that when the lumbar discs are injured as part of an experiment, bone spurs begin to form in the one- to twomonth time frame.



The same holds true for muscles, <u>tendons</u>, and <u>ligaments</u>. They all react to increased strain forces by making themselves thicker and stronger. This ability to react quickly to increased (or decreased) demands is mediated in part by adult stem cells. The switch from seeing these orthopedic tissues as inanimate filler (bone, cartilage) or pieces of inanimate duct tape (ligament, tendon) to living tissues that react is a key concept in understanding why alignment of the joints is so important in Orthopedics 2.0.



Functional Bone Spurs?



Bone reacts to forces. I've spoken to many patients over the past few years who are planning to have surgery to remove <u>bone spurs</u>. While I can think of a few situations where this makes a lot of sense, like when a bone spur is pressing on a nerve, most times bone spurs are better off left where they are. The bone spurs that are

causing mischief I call nonfunctional bone spurs. However, in many situations, bone spurs are functional. What does this mean? Let's take the example of bone spurs in the spine that need to be left intact.

We've been conditioned to believe that all bone spurs are bad. However, bone spurs develop in the presence of instability and grow to help stabilize the spine. If we remove these bone spurs, the spine becomes unstable and the body will just place more bone spurs in this location. I call these "functional bone spurs" in that they serve a purpose, and their removal doesn't positively impact the patient.

Beware of How Traditional Orthopedic Approaches Can Hurt Tissues

Let's explore some common traditional-treatment approaches that may adversely impact how you respond to the newer biologic therapies, like platelets and stem cells. For example, steroids are commonly injected into facet joints, and patients are often prescribed nonsteroidal anti-inflammatory drugs (NSAIDs). Is this a good idea? Regrettably, the research of the past decade is increasingly showing that not only are many of these approaches ineffective, but some actually make the problems worse. Let's explore these a bit.

The Opposite of Healing: Apoptosis (Steroid Shots Are Bad News!)

What's the opposite of healing? Causing <u>apoptosis</u>, or preprogrammed cell death without any ability to heal. For many years, doctors have injected <u>high-dose steroids</u> because they quickly bring down swelling and make the area feel better.



However, study after study continues to show that these drugs when used at the high doses that physicians often inject (milligrams) cause local preprogrammed cell death (apoptosis). While causing a little cell injury is not necessarily a bad thing (as discussed above), steroids work by taking away the local repair response (inflammation and swelling) as well, and so you're left with an injured area that can't repair itself.

Doesn't the body use steroids? Yes, your body can release natural steroids into an area where the inflammation dial may be turned up too high, which turns down that inflammation dial just a smidge. How much is too much steroid? While the <u>milligrams</u> of steroid commonly injected by doctors might not seem like much, it's about 100,000 to 1,000,000 times more steroid than your body would expect to see in the area. As an example, if the amount your body uses to control joint swelling is the height of a matchbook <u>(nanogram range)</u>, the amount most doctors have been taught to inject is the height of the Empire State Building

(milligram range)! Or, as I like to tell patients, if we inject the much smaller nanogram dose, we're putting in a thumbtack with a ball peen hammer, but if we inject the much larger milligram dose, it's like putting in the same thumbtack with a sledgehammer. If you use the ball peen hammer, there won't be much collateral damage, but using the sledgehammer is bound to create problems.

Why don't we see more doctors injecting the smaller physiologic doses? For one reason, they just aren't commercially available in those dose ranges. Steroids for injection bought from a medical supply company come only in the much bigger milligram ranges. Despite injecting the much smaller doses, we usually see the same results (decreased swelling). In addition, research has shown that these smaller doses can increase the good growth factors in a joint associated with repair.

Don't I Urgently Need to Get Rid of Inflammation?

The RICE approach in orthopedics has become widespread. The concept is to get rid of inflammation through rest, ice, compression, and elevation. But is getting rid of the inflammation always the best plan?

<u>"Inflammation"</u>—you've likely heard the term in a negative way. Inflammation means swelling. <u>You've likely heard that too much</u> <u>inflammation in our arteries may be the cause of heart disease</u>. You may have heard of a rare syndrome where too much inflammation after a leg or arm injury can cause serious problems (<u>compartment syndrome</u>),








where out-of-control swelling in a confined space can lead to severe injury). All of this is true, but for this chapter, you have to understand that like anything, there is also a good side to inflammation. Without inflammation, we would never heal ourselves.

Maybe you've had a chronically swollen joint or seen people with joints that swell. The reaction from modern medicine has been to inject high-dose steroids into these joints. As stated above, since high-dose steroids are potent at reducing inflammation, this may at first seem to help. However, these ultra-high-dose drugs also destroy the natural repair response. So we now have a joint that no longer swells but also has no ability to heal itself.



Why does a joint stay swollen? Swelling is the result of your body marshaling the troops to heal an area. All of the cell types needed to build new tissue are in the swollen area: cells to clean up the damaged tissue (macrophages), cells to recognize any foreign material and deactivate invaders (white blood cells), and cells to act as general contractors in managing the repair response (mesenchymal stem cells). However, your body will keep throwing inflammation at the area (swelling) if the right signal isn't received from the newly formed repair tissue. As discussed above, if

there aren't enough stem cells to complete the construction project, the "done" signal may never be received.

An easy way to think about swelling is that it's like the heat in the oven that's used to bake a cake. After all, the term "inflammation" incorporates the Greek for the word for flame. When an area stays swollen and chronically "inflamed," it's like low-level oven heat. If you place cake batter in a 200-degree oven, you don't end up with a baked cake but with driedout mush. Why? The chemical reaction that "bakes" the cake needs higher heat, or said another way, to be more "inflamed." Turn up the oven to 400 degrees and you get a cake. The same holds true for a chronically swollen joint. The low heat of chronic inflammation isn't enough to repair the tissue, so the joint stays swollen. However, using the microinjury or platelet techniques above, we can "turn up the heat" and use much



higher-level healing inflammation to heal the tissue (or bake the cake). The same can happen when we add stem cells, which can also help to complete the repair job.



So in summary, inflammation isn't usually a bad thing in orthopedic applications. Swelling is necessary to healing. Doing things to get rid of swelling (the rest, ice, compression, <u>elevation [RICE] mantra</u>) or using anti-inflammatory drugs may have their place in certain rare circumstances to prevent things like compartment syndromes. However, in Orthopedics 2.0, the use of drugs like high-dose steroids and <u>NSAIDs (Motrin, ibuprofen, aspirin, Aleve, or other nonsteroidal anti-inflammatory drugs)</u> to halt the healing inflammatory response is generally considered a bad idea.

Getting Rid of the Steroids in Epidurals

Epidural steroid shots have become the de facto standard of care for patients with a pinched spinal nerve causing pain. This treatment involves injecting a high-dose steroid around the irritated nerve, usually using X-ray guidance to ensure accurate placement. However, steroid epidurals have been in the press lately for their negative side effects. In one study, middle-aged and older women lost huge amounts of bone with each steroid epidural. In another, the part of the brain that reacts to stress and various important hormones was knocked off-line for three weeks with each injection. Why? The amount of steroids in these shots is massive compared to what the body is used to seeing (more on this later in the "Neuromuscular" chapter). As a result, we have switched most of our epidural patients over to platelet lysate instead of these high-dose toxic steroids. See this link for an infographic that goes over our registry data showing that PL epidurals (Regenexx-PL-Disc) produce a better and more robust treatment outcome than steroid epidurals.

Our licensees outside the United States also use various activated, incubated platelet supernatants or next-generation PRP. Think of these as PRP 2.0. To make these next-generation solutions, the lab starts with PRP, activates it with various natural substances, and incubates the platelets for several hours to several days. During this time, the platelets are directed to produce certain natural growth factors that are important in specific types of healing. We then take that soup of natural growth factors and use that instead of PRP.



So, for example, we can inject a solution that is rich in the <u>growth factor TGF-beta</u> (good at healing cartilage like those found in facets) by instructing the patient's platelets to produce more of that natural substance.



Should I Take Anti-inflammatory Pills to Help My Joints and Spinal Pain? Medications That Adversely Impact Regenerative Orthopedics



Anti-inflammatory drugs have become a mainstay of orthopedic and musculoskeletal care. While we have discussed steroid medications, what about nonsteroidal anti-inflammatory drugs (NSAIDs)? These are medications that block the pathways for inflammation—with most of them, the COX pathway (cyclooxygenase). COX drugs help control swelling, but they also cause stomach ulcers by inhibiting the enzyme that helps to protect the stomach wall. Just how dangerous are these drugs?

Moore, in 2002, published that the estimated risk of death due to bleeding stomach ulcer when taking NSAIDs for more than 60 days was 1 in 1,200! While this represents only a small number of people who are very sensitive to this drug class, the overall numbers are concerning. As a result of these inherent dangers, newer drugs were designed to work against COX-2 rather than COX-1 (the

latter being more responsible for protection of the stomach wall), but these drugs had a new set of side effects. <u>These drugs (like Vioxx, Bextra, and Celebrex) all come with an enhanced cardiovascular risk (risk of sudden death by heart attack)</u>.

How do NSAIDs impact healing? Well, from a 50,000-foot view, inflammation is needed to heal, so blocking inflammation may inhibit healing. Sure enough, <u>NSAID drugs, like Motrin</u> and others, have been shown to delay healing. While most of this research has focused on fracture healing, we keep patients undergoing regenerative-medicine treatments off these drugs.

Other drugs are also notable for causing musculoskeletal problems. The antibiotic drug class that includes <u>Cipro (quinalones) has been shown to lead to tendon ruptures.</u> <u>Heartburn drugs, like Nexium, have also been linked to hip-fracture risk.</u> Cholesterol drugs have been known to cause pain and <u>harm muscles.</u> Many commonly used drugs can adversely impact regenerative-medicine healing. Our own cell-culture data implicates <u>cholesterol</u> and <u>certain blood pressure drugs</u> as causing problems with <u>mesenchymal stem cell</u> growth in culture.



If Most Drugs Are Bad News, What Else Can I Do?

Can you change your lifestyle to protect your joints and spine? The answer is likely yes. I would break this down into a few different categories: diet, exercise, supplements, prescription medications, and hormones.

Diet

Could what you're eating impact your joints and spine? We know that <u>patients with</u> <u>metabolic syndrome get more arthritis</u>, independent of their weight. What's a metabolic syndrome? This is when you gain weight (usually in middle age, but it can happen earlier), participate in limited exercise, and start to get high blood pressure. This happens because of a combination of genes, low activity levels, and a sugar/starch-based diet. Basically, excessive sugar and carbohydrate consumption leads to spikes in insulin, which eventually makes the pancreas less sensitive to insulin causing more to be produced. This "hyperinsulinism" leads to a state where insulin is always present, and this hormone is a potent blocker of fat breakdown and facilitator of fat production. All of this not only causes weight gain but also unstable blood sugar, which leads to bad chemicals that can break down cartilage.

How do you fix it? Well, since you can't change your genes, you need to change your diet. If you want an instruction manual on how to eat, get one of the diet books, like <u>Atkins, The</u> <u>Zone, The South Beach Diet</u>, or <u>The Paleo Diet</u>, that focuses on weight loss through less sugar and carbs.

How do you know if you have better blood sugar control and, hence, are doing better protecting your cartilage? There are two tests, one that you can do at home and one that requires a doctor's visit. The first is what I call the "Dark-Chocolate Test." Before you start on this diet, you'll likely think that a 70% dark-chocolate bar tastes pretty bad or at least bitter. This is because you have set the level of sugar detection for your taste buds way too high by eating too much sugar and starch. However, once you are on a low-carb diet for a few months, the 70% should taste sweet. Next, try 80%. It should taste OK if you're a real low-carb superstar. If it doesn't, be stricter with your sugar intake. When you're done with your sugar transformation, an 85% bar will taste just fine, and that piece of birthday cake will be disgustingly sweet!

The doctor's office test is called an HbA1c serum level. This is a measure of changes to red blood cells in the presence of high blood sugar levels. It can take a few months to change, so get it tested a few months into a diet. While your doctor may tell you that anything below a 6.0 is fine, it's really not. You want yours well below 5.6, and for maximum



protection, it should be below 5.1. Remember, tracking this number takes patience! As your blood sugar control improves as you eat fewer carbs, this number may take many months to drop all the way to its nadir.

Exercise

First, what are "normal" levels of exercise for cartilage protection? Second, does pounding exercise, like running, destroy joints?

We Americans and others who live in First World countries have become very accustomed to low exercise levels. My favorite candidacy review that I have ever performed was when I asked the personal physician of a Middle Eastern woman, who was a member of the royal family, whether the woman got much exercise. I was promptly told that the royal family took cars and a security detail to go around the block! While this is an extreme example, it gets the point across: we have lowered our expectations for exercise so much that we no longer know what our bodies were designed to do. Another example is the health club across the street from our office. I see people in there working out as if they are the walking dead, strolling along on an elliptical machine as if it were a Sunday stroll and not their half hour to work their body hard.

What our ancestors considered everyday activity, most of us would consider our toughest exercise day. Think about this for a moment. You wake up and haul 100 pounds of water a half mile from the river to your home. You then chop wood with a 30-pound iron axe for an hour. Then you get to walk/run 20 miles while you hunt the big game that will keep your family fed for the next week. You get the picture.

So what is "normal activity" for our conversation on "Living 2.0"? It's 30 to 60 minutes of exercise so intense that having a normal conversation is very hard. This is performed five to six days a week and combined with weight lifting. In addition, my definition of weight lifting is absolutely not what I see in the gym across the street. It's 6 to 12 reps of whatever weight causes your muscles to fail by the 6th to 12th repetition. So if its biceps curls, pick up the weight that will cause your biceps to stop working by the end of the set. Then do this twice more as a minimum biceps workout.

Why am I pushing the weights so hard? <u>Because elderly weight lifters have muscles that at</u> the cellular level look more like younger people than their old sedentary counterparts. In addition, <u>exercise increases the stem cells in your muscles</u>.

What if pain prevents you from getting to this level? Well, that's why you're reading this book. Our goal is to use regenerative therapies, diet, exercise, and specialized therapies to keep you this active well into your 70s and beyond.



How about pounding exercise, like running? Is this good or bad for your joints? Regrettably, this cartilage research is still all over the map. Some studies have shown that running is protective to knee cartilage while others have shown it's destructive. In the meantime, like anything else, everything in moderation, or get a good doctor! By this, I mean that the ideal workout mixes up lots of activities, which might include running.

Supplements

One of the most frequent questions I get asked by prospective stem cell patients is what supplements they should take to help their chances of success. While from looking at the ads on the Internet, this would appear to be a very simple question, it's actually quite complex. The issue is that while some supplements have been tested with cartilage cells, we have very few that have been tested with stem cells.

What I can say is mostly from tests with either cartilage cells or early studies in patients where objective measures of cartilage health are used (not stem cells).

• **Glucosamine** is a very common supplement that can be derived from many sources. It's basically a cartilage building block, and there's a bunch of research showing it helps cartilage. See <u>article 1</u>, <u>article 2</u>, and <u>article 3</u>.

• **Chondroitin** is the cousin of glucosamine and another common cartilage-building-block supplement. Many studies also show it helps cartilage. See <u>article 1, article 2, and article 3.</u>





• **Fish oil** is a supplement that is taken widely, but poorly understood by most who take it. While this is an important source of omega-3 fatty acids that can help reduce swelling, and there's some evidence that it may help preserve cartilage, most people take way too little to see these effects. First, you need to make sure your fish oil isn't oxidized. So if your fish oil smells fishy, ditch it and get a better brand. Second, if you pop a few pills that you bought in a grocery store with the label "Fish Oil," you're taking way under the dosage associated with big health benefits (the amounts that Greenland Eskimos consume). Using that math, it would take 20 to 30 pills per day of these garden variety capsules to match this amount. An easier way to take more omega-3s is to buy concentrated EPA/EFA brands. This is about 3,000 to 6,000 mg of omega-3 fatty acids per day. More on all of this at this link.

• **Curcumin** is an extract of the Indian spice turmeric. The research looks promising that it's an anti-inflammatory and may help preserve cartilage. In fact, one study shows that it's as good as Motrin for pain and swelling. It also seems to work better when combined with other supplements like resveratrol.



How does the supplement Curcumin stack up against NSAID drugs?

• **Resveratrol** is a powerful antioxidant and activator of the SIRT1 gene which has been associated with longevity. It's found principally in the skin of grapes, and it's thought to be one reason the French suffer fewer cardiac events. Apparently, all of that red wine is loaded with resveratrol! It also seems to help poor blood-sugar control. That middle-aged paunch caused by declining insulin control can also eat up cartilage, so resveratrol may protect joints.



There are literally a hundred other supplements touted as good for arthritis. However, our patients also want supplements that will help their stem cells, yet little stem cell research exists with supplements, which is why we had to create our own.

At first, I thought this would be a quick two- to three-month project of testing stem cells with supplements to see how they grew. As we got further into this project of determining which supplements were the best at stimulating stem cells to grow more cartilage, the science got more and more complex. We learned we needed to look at the following:

▶ How the stem cells from the bone marrow of different ages of donors responded to different supplements. Most studies in this area choose young stem cells by default since samples are often bought through a science company or taken from grad students. Yet older stem cells are different and need to be tested alongside younger cells. In addition, the stem cells from arthritis patients are likely different, and few twentysomething grad students have arthritis. As a result, we took all of our cells from real arthritis patients.



▶ How the stem cells grew when exposed to supplements. This is not so easy, as many supplements are digested into different forms before they reach the bloodstream, so we tested these forms as well.

► How well the stem cells produced cartilage components when exposed to various supplements. To do this, we needed to look at the cells as they produced these cartilage-building blocks with quantitative fluorescent microscopy. To the right is an example of what this looks like, with the bright areas representing the amount of cartilage components—collagen 2, aggrecan, and SOX9—being produced by the cells.

▶ How well these supplements were able to support the overall robustness of the cells. A cell can look great in culture, grow well, and produce cartilage, but how healthy is it when stressed? In medicine, an example is the stress test for the heart. While someone might appear healthy, when they're pushed to the limit on a treadmill, their heart can show signs of trouble. We did the same thing for cells by adding in common bad chemicals found in arthritic joints that beat up cells like IL-6 and then seeing if the supplements helped the cells recover better.

• Determining the sourcing. Most supplements come from many different sources. As an example, glucosamine can be extracted from any number of shellfish sources, pig parts, or cow parts. Is one of these better than the other at helping human stem cells?



In the end, after a year of testing, we produced a supplement we could hang our hat on: Regenexx Advanced Stem Cell Support Formula. One that was based on basic research that we felt represented what we were trying to accomplish in patients. While every other supplement you can buy is merely put together based on snippets of data taken from unrelated lab research done by others, ours was a coordinated and costly plan to find the best ingredients for our patients. Click on the video link below to see a short animation on the Regenexx Advanced Stem Cell Support Formula.



Prescription Medications

When we first began culturing cells in 2006, we were surprised to find that some patients' stem cells wouldn't grow well. After a few culture failures, we decided to try and isolate the problem, and after much trial and error, we were surprised to learn that the culprit was usually a prescription medication. For example, when we would stop the patient's medication and take new cells and reculture them, the stem cells would grow fine. We could also get the cells to grow fine when we exposed them to the serum of another person who wasn't on the drug (for research purposes only). This convinced us that many prescription medications are toxic to stem cells.

What should you avoid? Likely, the most surprising thing we encountered was that America's wonder drugs, cholesterol medications known as statins, seemed toxic to stem cells. I would take a minute to read about <u>the limited efficacy of these drugs</u> if you're on them, but only make any decisions to stop the drug after consulting your primary care physician. Other drugs that were problems included NSAIDs, like ibuprofen, Motrin, Aleve, and Celebrex. Given that <u>NSAID drugs carry significant cardiac risks</u>, <u>not taking these</u> <u>drugs makes sense</u>. We have found fish oil makes a great alternative anti-inflammatory,



and research supports this effect. Other problem drugs include <u>ACE-inhibitor blood-pressure medications</u> that are usually taken to combat hypertension due to metabolic syndrome. Rather than take these drugs, increase activities and cut the carbs to control your blood pressure. Finally, <u>steroid anti-inflammatories</u> are also a big concern as they are toxic to many different cell types, including stem cells.

Hormones

We take our hormonal mix for granted, a little like the fact that our heart beats more than 100,000 times a day. Yet if you're trying to control your weight and stem cells, these hormones can play a critical role as you age. For example, <u>serum taken from women</u> <u>after menopause</u> (when many hormones are no longer produced) directs stem cells that should turn to bone to turn into fat instead. For men, testosterone levels decline as we age. Yet <u>testosterone can activate stem cells</u> to produce more muscle and less fat. While the jury is still out on the effects of sex hormones on stem cells, in men it's pretty clear that supplementing these hormones in middle-aged and elderly patients can help overall body composition, reducing fat and <u>increasing lean muscle mass and likely improving heart</u> <u>health</u>. The same <u>has been shown for women</u>. On the other hand, <u>the willy-nilly prescribing</u> <u>of testosterone in men</u>, without strict monitoring, may be linked to an increase in heart attacks.

Should you supplement your hormones? This is a personal decision. In my experience, it makes it much easier to keep the pounds off and activity levels high as we age. On the flip side, for men, urologists have been concerned that testosterone may cause prostate cancer. However, <u>newer research suggests this likely isn't the case</u>. In addition, without strict monitoring, the excess production of blood cells may be the cause of blood clots that can cause heart attacks. For women, the Women's Health Initiative study suggested that estrogen pills could cause cancer, yet the study authors have since stated that the press misinterpreted their results. <u>They now recommend hormone therapy for younger women just after menopause</u>.

My Own Living 2.0 Story

At the age of 37, I was at the low point of my personal health. My wife and I had just gone through the trauma of having premature twins, and my exercise was sparse while my sleep was nonexistent. My weight had ballooned such that my waist had gone from a 32 while in residency to a 42. Just jumping onto our high bed made my heart race. I was an early heart attack waiting to happen. It took over a decade to completely climb out of the hole I dug, and I learned quite a few things. Had I known these at 37, I could have been in perfect shape by 40!



What did I learn along the way?

► **Carbs are key**. While for my wife's genetics, carbs aren't important, for me they are everything. I spent years on fad diets, trying to prevent my waist from ballooning and avoiding any type of shopping for pants (as that meant admitting that I could no longer fit into a size 40), none of which worked. In the end, it was understanding that limiting my sugars and starchy carbs was the answer. However, that got me only part of the way there.

▶ Understanding what the term "workout" means. Like I have said, I see people working out at the gym like they're the walking dead, and I understand that once that was me. It wasn't until I got a personal trainer that I finally understood what working out was all about. My trainer worked me like a dog, and the harder he pushed, the more I wanted. It took a few years with a trainer to understand that to stay in good shape in middle age for most of us requires working very hard. An example is the *INSANITY* workouts. For most of those 30 to 40 minutes, the exercise is tough enough that holding a conversation is difficult—you're simply pushing too hard. A kinder and gentler approach that starts slow and builds (by the same trainer) is FOCUS T25. The same applies with weights, which should be a cardio workout as well. If you're not huffing and puffing while you quickly move from one weight station to the next, you're not hitting it hard enough.

When I finally figured this out, my weight went down again. But there were still two more pieces.

▶ Hormonal control is a problem for middle-aged men and women. I spent many years being asked to lecture to doctors involved in age-management medicine as many are very interested in stem cells. I would sit through the lectures before and after mine, which were about supplementing hormones in middle-aged and elderly men and women. I have to say that at first I thought these doctors were a bit nuts. Was this really necessary? After a few years of seeing countless lecturers review the science, I was convinced that I had to be my own guinea pig. In 2010, I worked with a Denver-based age-management specialist to check my testosterone levels. They were dismally low, so I started testosterone therapy. The final piece of the puzzle clicked into place for me. Much of the extra weight that I had been carrying melted off as I already had the diet and exercise piece under good control. Unlike the recent rush to prescribe testosterone to everyone with "Low T," this program is carefully monitored. Which brings me to the last part.

▶ **Proactive injury management is important.** You can't help but get injured from time to time if you're hitting exercise hard enough to stay active as you age. I constantly look at my joint stability, levels of nerve irritation, muscular firing, and body symmetry—just as I describe in this book. All of this is to try and prevent injuries or catch problems when



they're early. Since supplementing testosterone pushes my blood hematocrit up (the number of red blood cells), I put those extra blood cells to good use. Once every two to three months, I have blood taken to reduce my hematocrit and to fuel the Regenexx-SCP and Regenexx-PL procedures. I proactively have small issues treated with these regenerative cocktails. As an example, my right knee, left shoulder AC joint, right elbow, and low back are common problem areas for me. These aren't disabling problems but are areas of constant irritation—all likely related to wear and tear and declining stem cell activity as I age. So I have my colleagues give these areas an "espresso shot" of my own healing blood platelets.

The conclusion? For me, controlling my weight, staying off prescription drugs, and staying active requires four key elements: diet, exercise, hormones, and early injury detection and management.



Chapter 4: Symmetry



"True freedom is where an individual's thoughts and actions are in alignment with that which is true, correct, and of honor—no matter the personal price." Bryant H. McGill



In most patients, there's usually more joint damage on one side than the other. Why? If someone has a genetic predisposition to arthritis and this is the only factor causing the joints to degenerate, shouldn't all joints be affected equally? In addition, osteoarthritis is more commonly seen first in the knees and hips and less often in the ankles and elbows. Why? Again, shouldn't we see all joints being impacted the same? The reason is clear: the wear and tear on our joints occurs unevenly, with some being impacted more than others or one side undergoing more wear than the other.

Why Should I Care About My Symmetry?

If you don't want to stay active and pain free as you age, then ignore this chapter. If you do want to stay active, then pay attention! Your body is designed to be symmetrical, and even slight amounts of extra force or motion, in any area, caused by an unbalanced body will cause problems. These may start small, but like a snowball rolling downhill, they become bigger and bigger with time. This is especially true of the spine, which is designed to be have opposing curves that balance each other.

Reducing the Wear and Tear



As already discussed, increasing healing ability is only one part of the Orthopedics 2.0 equation. The other half of this coin is reducing the wear-and-tear forces that destroyed the joint in the first place. As an example, placing new tires on a car with bad alignment without fixing the alignment is guaranteed to quickly wear out the tires again. This issue is often ignored in our current quick-fix treatment methodology. I've seen hundreds of patients with a specific wear pattern, like on the right medial meniscus, where the most salient questions have never been asked: how did this knee get like this, and what are we going to do to ensure that it doesn't get this way again?

The reason the issue of specific wear-and-tear patterns is mostly ignored is that it's complex. Most physicians aren't trained to understand the biomechanics of the body. The few physical therapists or other providers who have spent years of extra study learning biomechanics are often too heavily incentivized by insurance companies to take the time needed to figure out why a part keeps failing.



Let's use a simple example to illustrate this concept. The skeleton on the right has been drawn with red force arrows going down from the hips. Let's say that because the spine is slightly bent to the side that slightly more force is applied to the right side (the thicker arrow) than the left side (the thinner arrow). We take thousands of steps a day. What happens to the extra forces on the right, and how does the body handle them?

The right hip, knee, and ankle will all react. They will initially just shore up the bones, tendons, ligaments, cartilages, and muscles on that side. When this person is young, with many adult stem cells in these areas, he or she may not notice much. However, as the number



of adult stem cells begins to decrease with aging, the damage due to wear and tear at some point will begin to overtake the repair ability of these tissues and react to the extra forces. These areas (the ones that are the most vulnerable) will begin to break down.

If our only goal is to replace one of the right-sided joints with an artificial joint, that prosthesis may wear out a bit faster on that side, but this is likely not a huge issue. However, if we want to preserve that right-sided joint with Orthopedic 2.0-type procedures, we had better figure out why that joint is getting so much more wear and tear and correct that problem.

Take Five Minutes to Understand Your Symmetry

Patients are often surprised to find out that their body is no longer symmetrical. Others have noticed that a certain area has been getting tight for years. Either way, going through an intensive five-minute test of all major body joints, checking for symmetry is well worth your time. Why? As discussed above, joints that are not equal in their side-to-side or front-to-back motion don't wear evenly. If you have any areas that are tight and not symmetrical and you don't figure out what's going on and fix these, you can bet arthritis is in your future if it's not already knocking at the door. In fact, symmetry is often the single biggest thing you can work to fix that may either allow you to have an active older age or one plagued by problems and inactivity.

This is an 11-step test. You need to focus on whether you can attain the movement to the degree asked, whether the right and left are identical or different, whether the front/back is the same or different, and whether the tightness in the lettered areas is different on one side or the other. Realize the differences may be subtle but important. In addition, pay attention to which movements cause pain and where. **If you're a patient of our clinic or a Regenexx Network physician, fill out the form in appendix A and bring it to your visit.**



If any movement causes significant pain, stop immediately and see a doctor. **If you think** any of these motions may injure you, or you are prone to injuries with common or unusual movement, don't do this test, and, instead, see a physician.

Step 1: Protracted Shoulder Check



Stand normally and place your hands together. Raise them over your head as shown. Move the hands as far back as they will go. Pay attention to whether the shoulders go back equally (you may want to have someone check, or do this in a mirror). Also pay attention to whether the front of the shoulders at points A and B are equally tight or if one of them is painful. You can also do this same maneuver lying flat on your back on a firm surface. In that case, your hands should touch the floor. If they don't, then both sides of your chest are too tight, or you may have problems with the range of motion of the large shoulder joint. This step checks your ability to abduct the shoulder and also checks the tightness in the pectoralis major and minor muscles along with the front and bottom of the shoulder capsule. If this causes shoulder pain, you

may have a shoulder impingement, a problem where the structures of the shoulder are getting pinched by poor movement patterns.

Step 2: Posterior Shoulder Check

Stand normally and place one hand behind your back. Raise it up as high as it will go. Both hands should be able to go at least to the middle of your upper back with the thumb just below the shoulder blades. Notice whether both sides can go equally as high and whether you have tightness in the back of the shoulder at points A and B. Is one side tighter than the other? This movement measures the tightness in the infraspinatus muscle as well as the back of the shoulder capsule. Patients that can't do this tend to develop overload on the back of the shoulder joint where they can develop labral tears. While many surgeons would just focus on the labral tear, the real focus should be on why this shoulder can't internally rotate!





Step 3: Neck Check



Very slowly and carefully roll your neck 360 degrees (only 180 is shown). Does this cause any pain at any point? Does it roll equally well to the front, side, and back? Is one side tighter than the other? Patients who don't do well with going straight back or back and to the sides may have an arthritic or injured facet joint in the neck or

a problem with lordosis (discussed later in this chapter). If bending to the side or forward is tough, you may have tightness in the postural muscles of the neck that hold the head up, like the levator scapula, upper trapezius, or cervical extensors. Finally, if one of the front muscles is tight, it could be the stenocleidomastoid, an important muscle that helps to turn the head. You may also notice that you can't hold your head back like this or that if feels out of control if you do. This is often caused by weakness in the deep-neck flexors, like longus colli or longus capitis. Weakness in these deep-neck-flexor muscles can lead to chronic headaches. In the "Neuromuscular" chapter (next chapter), there's a strength-andendurance test that also covers this issue.

Step 4: Cervical and Thoracic Rotation Check

While standing, place your hands on your chest, and turn your head all the way over your shoulder as far as it will go. Then follow through with a rotation of your upper back as far as it will go with your feet firmly planted (they shouldn't move). Do this on both sides. Can you turn as far with your neck and your upper back on the left as on the right? Is there more tightness on one side of the neck, upper back, or lower back than on the other? Patients who can't turn their neck may have a problem with the facet joints, whereas patients who can't turn their upper back may have that issue or a problem with the normal motion of the rib cage, the rib



attachments at the spine (called rib facets), or the thoracic spinal facet joints. In addition, chronic chest wall tightness on one or both sides may also limit the rotation of the upper back.



Step 5: Hip Rotation Check

1Stand normally and place the toes of your feet together as shown. Make sure your feet are aligned and symmetrical—it's easy to cheat by placing one foot forward or back. Note whether both feet move inward equally (the motion is mostly coming from your hips). Also note the A and B points listed in the front and back of the hips. Are these areas equally tight? Is one tighter than the other? Also note the C and D points—does any of this stress or hurt your knees or one knee? Now take



your toes and rotate them out all the way. Again, be careful to make sure your feet are symmetrical (heels are together) as otherwise it's cheating. If one hip has a very different range of motion (toes don't move as far in or out, and this seems to be due to tightness in the hip), this is very concerning. The hips tend to lose range of motion quickly and almost always after the onset of moderate or severe arthritis. I would advise you to get your hip checked immediately by your physician. If you already know you have a hip problem, this means that you have serious work to do. Unlike other joints, the hip has a very limited weight-bearing area (the part of the joint where it gets the most force). When the hip loses range of motion, and when arthritis is already present, the joint will put much more pressure on already worn areas, hastening cartilage loss. Getting hip range of motion back can be a challenge. In addition, patients with poor hip range of motion tend to have a less robust response to stem cell injections. As a result, there are procedures we can use to try and improve the hip range of motion through an injection that stretches the joint capsule.

Step 6: Lateral Hip and Back Check

Stand normally and reach to the side as shown. Go as far as you can, and note points



A and B. Can you go as far on the right as you can on the left? Does one side of the lower back and/or outside hip feel tighter (point A or B)? Does that tightness extend down the side of the leg to the knee (points C and D)? This step measures the tightness in the opposite lateral lower and upper back muscles, like the quadratus lumborum and iliocostalis lumborum. It also measures the tightness of the opposite lateral hip muscles, like the tensor fascia latae and iliotibial band. If you can't bend as much to one side, there may be tightness in these muscles or the spine. If you have pain that goes down the side of the hip and leg, you may have an SI joint problem with tightness in the iliotibial band or an S1 nerve problem in your back.



Step 7: Gluteal, Lumbar, and Thoracic Extensor Check



Bend forward all the way, and try and touch your toes. Do you get pain or severe tightness in one or both hamstrings (point A)? Is one side of your buttocks tighter than the other (point B)? Is your lower or upper back tight (points C and D)? Can you get within 6 inches of the floor? Does this cause pain or perhaps tightness, numbness, or tingling in one or both legs? Is your belly in the way? Patients who have difficulty getting close to the floor have significant tightness in hip flexion (which could be arthritis) or lower/upper back flexion (which could be disc disease or extensor muscle tightness). If one hamstring is always tight despite your best efforts to stretch it, it could actually be an irritated S1 nerve in your back despite

the fact that your back doesn't hurt that much. Irritated nerves in your back can also cause one or both legs to get tingly in this position.

Step 8: Hip Flexor, Ankle Dorsiflexion, and Forefoot Check

Perform a lunge as shown, getting as far down as you can, making sure you feel a good stretch in the front of the hip on the back leg. On that leg, bend your toes so that they meet the floor flat. On the other leg, flex the ankle as much as possible. For some people, getting a good stretch may involve placing the hands all the way down to the floor. Notice the front of the ankle of the forward leg, and compare that to the other side when you perform the opposite stretch (point A). Does the buttock on the front leg have the same side-to-side tightness? On the back leg, can you flex your toes (point B), or is this restricted on one side or painful? On the same leg, is



the front of the hip (point D) equally tight on both sides, or is one side tighter? Can you get as low on each side? Patients who have difficulty at point A (front of the ankle) may have ankle arthritis or a bone spur in the front of the tibiotalar joint, restricting dorsiflexion of the ankle. Patients with problems at B (toes) may have arthritis at the metatarsophalangeal (MTP) joints in the foot. For the big and 2nd toe, this can sometimes be related to longstanding low-back problems (even though you think your back is fine). Patients with problems at C (buttocks) may have problems with hip flexion, indicating tight gluteal muscles or arthritis in the hip. Pain with this maneuver could also mean a labral tear in the front of the hip. Finally, if you have an issue at D (front of the hip), this could indicate a tight psoas muscle. This muscle goes from the front of the lumbar spine to the hip, so tightness here can be due to chronic low-back issues or trigger points in this muscle. Sometimes patients with psoas issues have trouble getting in and out of a low car.



Step 9: Knee-Extensor Mechanism Check



Stand normally and grab one foot with the same hand while bending the knee as shown. You may need to hold onto something. That's actually your first observation. If you can't easily balance like this (after a practice run), then you have significant low-back and hip-stability problems on the opposite side of the knee bend (see the <u>"Stability" chapter</u>). For the symmetry check, do the right and left knees bend equally? To really check this, make sure you stand straight while checking both sides. Does point A (quadriceps) feel the same on each side? How about the front of the hip (point B). Does either knee hurt in this position? If you have less knee bending on either side, the simplest explanation is that you have trigger points in the quadriceps muscle (see next chapter). If the front of the hip is tight, you may have issues with the rectus femoris muscle. If the knee hurts, you may have a patellofemoral problem (an issue with the kneecap in

its groove). Another common cause of asymmetry here is swelling in the knee joint due to chronic arthritis, which reduces the ability of the knee to flex.

Step 10: Adductor, Sartorius, and Gracilis Check

Lie on the floor and place one ankle over the opposite knee as shown. Next, try to get the bent knee as low to the floor as possible. Check both sides and see if they are equal in your ability to get the ankle high up on the opposite knee. See if point A (inside of the thigh) feels the same degree of tightness side to side? Can you get one bent knee farther toward the floor than the other? Tightness in



these muscles of the inside of the thigh are common in patients with chronic low-back conditions and sometimes can cause hip or inside-of-knee pain. In addition, patients with hip arthritis may notice a difference in flexibility from side to side.



Step 11: Lumbar and Thoracic Extension Check



Lie face down and prop up on your elbows, arching your back by lifting your head as high as possible and pushing your hips into the floor. Can you do this without pain? Does your lower back (point A) or upper back (point B) feel tight or hurt? Do you have pain or tightness at the back of the shoulder blades?

Patients who have tightness in the front of the hip and low back may have tight psoas muscles. Patients who have pain in their low back with this maneuver may have injured or arthritic low-back facet joints. If you have pain in one shoulder blade, there could be a problem in the joint between the shoulder blade and ribs or in the rib cage itself.

What do you do now with this information? If you have an area where your movement isn't normal, or there's a noticeable side-to-side difference in motion or tightness, there may be a few different causes. First, this needs to be looked at by a physician, physical therapist, or other musculoskeletal provider. Why? In our experience, asymmetrical motion is a leading cause of excessive wear-and-tear arthritis, so getting symmetrical and balanced motion back is absolutely critical. Second, the lack of motion may indicate problems in that joint that have yet to be addressed.

What are some common ways to treat these tight areas?

1. Do simple stretches. The longest-running stretching book on the market is Bob Anderson's. <u>See this link for Amazon</u> or <u>this one for the basic stretches.</u>

2. Clear trigger points. When I discuss specific muscles above, these are your areas to target. Many times there are knots in the muscles that when cleared will allow normal movement. The next chapter will focus more on ways to get rid of these trigger points.

3. Repair irritated nerves. Sometimes an irritated nerve won't allow normal motion in an area, as a protective response for the nerve. This topic is dealt with in the next chapter.

4. Treat joint arthritis. Sometimes an arthritic joint won't allow motion because bone spurs within the joint are blocking motion or the covering of the joint (the capsule) is too tight. For information on these issues, see the <u>"Articulation" chapter</u>.



Now That You Know You Have Problems, Let's Learn More About Symmetry

Everything starts with the spine, as your arms, legs, and head would be useless without an anchor point. That anchor is your spinal column. You might be saying, But I have a shoulder or knee problem. Why should I care about the spine? Because regardless of where you feel the pain, it's almost impossible for your spine not to be involved in some way.

A documentary on the tragedy of 9/11 provides a vivid example that may help bring this concept home. On one of the upper floors, many workers were trapped by a door to the stairs that wouldn't open. No matter how many large guys tried to ram it open, it wouldn't budge. It turns out that the violence of the airplane strike had twisted the spine of the building ever so slightly, and this compressed the doorframe against the door. These workers were eventually saved by a heroic building superintendent who guessed accurately that kicking a hole in the drywall next to the doorframe might release the pressure on the door. Your spine is the same. Small issues here can cause large problems in your shoulders, arms, hands, hips, legs, ankles, or feet.

Lordosis in the Spine, or It's All About the Curves



Lordosis is the medical term for a front-back spinal curve. A healthy body in normal standing strives to use as little energy as possible. The diagram to the left shows how the neck and low-back curves are counterbalanced so that you can stand up straight with a minimal amount of effort.

Far too little attention has traditionally been placed on these curves, but the medical and surgical community is slowly changing their position on this important issue. <u>Researchers have completed clinical trials showing</u> <u>that rehabilitation efforts to restore these curves do</u> <u>produce positive clinical results</u>. Some spine surgeons now routinely take extra time and energy to restore or preserve the normal or natural spinal curve when performing fusions and disc replacements.

Why are these so important? First, it's all about energy consumption. A way to help you understand this is in the challenges facing modern robotics. A dream of robot designers has been to design an android robot that can cook and clean your house like a personal servant. The problem was that designing a robot that could really walk like a human



turned out to be a huge engineering nightmare. However, when engineers solved that, <u>they soon realized that the human body had another secret</u>: it uses very little energy to walk, whereas their robot was an energy hog, lasting only 30 minutes on battery power. Why? The body is designed to conserve energy in movement, and it all starts with the spinal curves. These finely tuned curves allow you to stand using almost no energy. Knock one out by causing straightening and it dramatically upsets the whole system. In addition, the wear and tear on certain structures changes dramatically as well.

How would you know if you have bad spinal curves? It can be found on your MRI or X-rays, but these issues aren't often identified by the reading radiologist. It usually takes a doctor who's looking for these issues. Why is this big deal? It can fry your neck or back discs, leading to more bulges, tears, and herniations.

Each vertebra (spine bone) is built and oriented in such a way as to equally distribute forces between the front part <u>(the disc)</u> and the back part <u>(the facet joints)</u>. This only happens when the normal spinal curves (lordosis) are present (see below left).



When the curve is lost (straight spine), the forces get distributed more toward the disc, which can cause it to get overloaded. We often see this on MRIs as swelling in the vertebra around the disc (which means the disc isn't capable of handling all of these forces, and they're getting distributed to the surrounding bone). The official medical term for this MRI finding is a "Modic change."

Note the diagram to the right, which shows a loss of curve and a move of forces (large red arrows) to the front of the spine (disc).

This loss of spine curve is a problem when considering stem cell therapy or any other type of regenerative procedure on the disc because while such a procedure might be able to help keep the disc from failing, the same forces that caused the disc to fail in the first place (loss of the spinal curve, placing too much pressure on the disc) will persist. So we believe it's better to have the patient undergo therapy to restore the normal curve either before or during regenerative disc therapy.





What happens if the curve is too much? In that case, this is called a <u>hyperlordosis</u> (see figure on left), and the weight gets distributed too much to the facet joint. <u>These little joints are about the size of your finger joints and live in pairs at the back of the spine.</u> There are two at each spinal level, with one on the left and one on the right. They help to control motion, so that any specific vertebra can't get too far out of line. When they get overloaded by a big spinal curve, they can



get arthritis much more quickly and show signs of wear, such as cysts and <u>hypertrophy</u> (the body will literally make them bigger to handle the extra weight). When these joints become bigger (arthritis), they can place <u>pressure on the nearby exiting spinal nerves</u> and cause a new set of problems.

Forward Head, Kyphosis, and Lower Neck Joints

As discussed above, the spine is a delicately balanced machine where the neck and low-back curves are counterbalanced by the thoracic curve. As we age, our head and shoulders tend to hunch forward as shown below. As this happens, in order for us to see straight again, we crank our necks back farther. Because of this extra backward pressure on the neck, the facet joints in the bottom of the neck can be compressed, causing pain and more arthritis at those levels.

While performing <u>traditional facet injections</u> into these aggravated joints can help, it does nothing to solve the excess loads on the area, so the pain just returns. This is a great example of how Orthopedics 2.0 differs from pain management. The pain management approach is to put <u>steroid anti-inflammatories</u> into the joint to reduce pain. The Orthopedics 2.0 approach is to treat the joint (more likely with either low-dose anti-inflammatories or regenerative techniques), but also to work on getting weight off the joints by implementing a therapy regime that restores the proper curve, posture, or alignment.



Can Your Forward Head Cause Shoulder Arthritis?



What causes shoulder arthritis? If your shoulders have been aching as you age, you may want to pay attention. Are you in the early stages of shoulder arthritis? Is there something you can do to combat this arthritis and a forward head?

Note the diagram to the left.



When you're younger, your head and shoulders are straight and aligned with a perfectly balanced spine (neck curve points one way, upper back curve balances it by pointing the other, and your low back curves back the other way). Note that the blowup of the side and front views of the normal shoulder on the left has the ball perfectly aligned in the socket. As you get older, your head and shoulders and spine begin to move forward (due to our habit of sitting in chairs). As this progresses, the upper back part of the spine can become kyphotic (humpbacked). In fact, you may be noticing that you now have to catch yourself as it becomes more natural to be in that kyphotic (slouched) position.

What does all of this have to do with your shoulders? As you become more kyphotic (see right, above), your shoulders will begin to move forward. This has caused the ball of the shoulder joint to fall forward and down, out of the shallow socket. The new position now requires the rotator cuff (red in the front-shoulder blowup on the right above) to have to resist that new downward hanging force. In addition, it overstresses the top and bottom of the shoulder-capsule ligaments. As this progresses through the years, if you don't fix the kyphosis, the bottom part of the ball of the shoulder joint will continue to bump into the bottom of the socket. This will cause new bone spurs down there, which will further limit motion. In addition, the rotator cuff muscles will also begin to fail from all of that extra stretching. Finally, this new shoulder position will predispose you to more impingement (where the rotator cuff and biceps tendon get pinched).

So what can you do? First, in the early phases of this kyphosis, you can beat it by using constant stretching against the grain of the forward head and shoulder movement. Here are some of my personal favorites:



Lift your arms above your head and pinch your shoulder blades, feeling the stretch in vour chest. Now lift and lengthen your body while bending very slightly back. You should feel it all the way down in your abdominals.

Roll up a towel or use a foam roller and lie back and stretch the front of the chest. You can vary your arm positions up and down. You may want to have the towel or foam roller extend to your head and neck for better support.





Roll up a towel or use a foam roller and lie back and stretch the front of the chest. You can vary your arm positions up and down. You may want to have the towel or foam roller extend to your head and neck for better support.



In the later phases (once the spine bones have changed shape), you'll likely need professional help. In any case, if these stretches into extension cause significant neck, upper back, or lower back pain, you may have already had bone changes. Another possibility is that the facet joints in your spine may be arthritic or painful. Either way, you'll need to see a physician, like one of those in the Regenexx Network, if this is happening.

Are there regenerative injection treatments? If the shoulder ligaments on the bottom of the joint are lax, injections to tighten these ligaments (like the inferior glenohumeral

ligament [IGHL]) can be very helpful. In addition, if the main shoulder joint has arthritis, stem cell injections into the joint may help. The same holds true for degenerative tears in the rotator cuff, which based on our treatment registry data, respond very well to precise stem cell injections.

The Fight Against Gravity! More Things to Do at Home



We all know that aging is a fight against gravity. Yet with stem cells in the mix, we can understand it differently. For example, as discussed in this chapter, that forward head and hump some people get in their upper back is merely gravity plus bad posture

plus stem cells. What I mean by that is that any position of a joint or a part of the body held long enough will cause the local stem cells in bone to react, changing the shape of the bone.

What are some simple things to do to fight this change in bone shape? Work against it! Above is an example of using a foam roller to fight the kyphosis (head forward position) that I explained earlier. For this exercise, the <u>foam roller (the white cylinder above) is easily</u> <u>purchased as an inexpensive fitness item.</u> A rolled up towel also works well. When you first begin, lie like this to get a sense of if it hurts. If it causes pain, then stop and see a doctor. If it doesn't, then start with 5 minutes once a day and work up to <u>more-advanced exercises</u> and longer time periods (15 minutes). Any part of your body that's beginning to change can be dealt with in the same way.



Is Your Belly Frying Your Facet Joints?



Your back hurts, and like many Americans you have a belly. Could there be a connection? We discussed how changes in the curve of the spine can cause either the discs or the facet joints to get overloaded. If you look at the image to the left, note that a big belly will want to topple you forward. There's just no way to stay upright without compensating. How does that work? You'll have to naturally increase the curve in the low back to move the center of gravity of the belly backward. In doing so, as the blowup in the diagram to the left shows, your body weight gets transferred from being evenly distributed

between the discs in the front of the spine and the facet joints in the back to being more on those joints (the red arrows). Over time, this will cause your facet joints to get arthritis.

Scoliosis: A Side-Bent Spine

Your spine should be perfectly straight when viewed from the front. However, the spine can also bend to the side in a disease called scoliosis, especially in tall women. This can cause the shoulders and pelvis to become unbalanced. To the right, see how much havoc this can cause in the body, as the pelvis where the hips connect is way off. Also, it shouldn't be a surprise that when your shoulders are off, like the ones to the right, the forces on the ball and socket joint there will be very different.



What about when the side curves are more subtle? This is where many of us live—slight curves in the neck or back.

What Did I Self-Diagnose in the Symmetry Screen?

So far, we've been discussing the concept of poor stability caused by too much movement, or <u>hypermobility</u>. Equally important is <u>hypomobility</u>, or where a joint or spinal segment doesn't move enough in all directions or certain directions. This is what you self-diagnosed in the five-minute symmetry screen.

Chiropractors and osteopaths have been focusing on hypomobility for more than a century. The reason we medical doctors have given them a hard time is that hypomobility



has been traditionally hard to measure. However, there is good evidence now that this does occur. In fact, studies that specifically apply this concept (hypermobility versus hypomobility) show that patients with spinal hypermobility treated with exercise do better than patients with hypomobility. This makes sense because if you have too much mobility you need to get the muscular stability system back online with exercise or other treatments to restore muscle function. However, patients with hypomobility did poorly with stability exercise. Why? They need more mobility, not more stability. This group did better with manipulation to force these segments to move.



Hypomobility can be as damaging as hypermobility. The take-home message is that if your joints don't move normally in all directions, you have to get them to move normally, or this will place more wear and tear on the certain parts of the joint. What are some examples? If your knee won't straighten all the way, the front of the joint cartilage will wear more than the back. How about

a hip that won't turn out all the way? The inner part of the hip will wear more than the outer.

Take, for example, this simple model of a joint and the muscles that help control that joint's movement. We have a ball-and-socket-type joint with a ball sitting in a shallow socket (like the shoulder). Here we'll call them muscle A and muscle B. Both muscle A and B pull equally on the joint. When one pulls harder, the opposite muscle lengthens equally to allow the joint to move.

So as this joint moves, the ball stays in the middle of the socket. In fact, keeping the joint aligned with millimeter precision as it moves is critical.

What happens if one muscle can't release as the other pulls? Now the joint moves too much to one side, banging into the side of the socket. This is an example of muscular hypomobility, where trigger points in one muscle make it tight and weak (more on this topic at this link). How do you fix this problem? You need to loosen up the tight muscle. This is discussed further in the neuromuscular section. The same thing can happen if one part of



the joint capsule (the thick fibrous covering of the joint that helps to limit motion) is too tight or the ligaments that hold the joint together are too tight.



The Master Control System for Body Symmetry

The upper neck is a special area with respect to alignment and posture. We've all heard about the <u>balance system that involves the inner ear.</u> However, our balance system is very complex. A bit like a NASA spaceship, it has triple redundant systems, so that if one fails to provide enough information about balance, another system will automatically take over. This makes sense, since the loss of balance perception is not compatible with leading an active life and fending for yourself. Our balance system, therefore, has three inputs (as shown on the right): the inner ear, the eyes, and the upper-neck joints. For many years, the focus has only been on the inner ear, but these other two systems are equally important and play into symmetry.





What part of the upper neck seems to be the most important for body symmetry? A number of years ago, a team of researchers started killing the little nerves that took information from the C2-C3 joints. The goal was to help alleviate the pain associated with these joints. While the pain got better, these patients all became very dizzy. The researchers were confused. While they knew about the inner ear being involved in balance, they didn't know about the upper neck. This misconception still exists in medicine today, with the vast majority of doctors not knowing about "cervicogenic dizziness," or dizziness coming from the neck, despite published research on the phenomenon. Based on the research, the C2-C3 area and upper neck muscles, as well as the sternocleidomastoid muscles, all seem to be implicated.





So what does this have to do with alignment? Recently, <u>some Australian researchers have</u> <u>determined that patients with whiplash injuries</u> <u>have difficulty in determining which end is</u> <u>up.</u> Literally, their necks don't have the same <u>proprioceptive</u> ability as normal subjects. We believe that this is due <u>to injury of the upper neck</u> <u>joints, which is common in this type of injury</u>. We also see this same type of problem in patients who have injured the upper neck in the past or, for some reason, have chronic arthritis at C2–C3 or the high upper neck. These patients often have a head tilt.

When I correct the head tilt on exam so their heads are straight, they feel crooked. Why? The upper neck joints are giving bad information about normal posture. This bad data causes the patient to tilt the head, but this strange position feels normal to the patient. The <u>"righting reflex"</u> then kicks in to keep the eyes and head level while standing. This causes the patient to tilt the body to one side to compensate, as the neck stays bent. This then causes all sorts of problems with arms frequently getting numb (usually on the little-finger side of the hand) from <u>thoracic outlet syndrome</u> (the nerves in the shoulder getting pinched). In addition, notice above how this causes the spine to bend sideways and can impact areas all the way down to pelvis. It can even alter how one side of the leg/foot strikes the ground. As a result, in looking at alignment, one must always consider how it interfaces with this master balance-control system.



Can Alignment Be Impacted When the Spinal Stability System Goes Off-Line?

You've already learned about the stability system in the low back (multifidus muscle). Is there something similar in the neck, and can it impact alignment? We asked ourselves that question about 10 years ago. One of our physical therapists was tasked with seeing if the multifidus muscle in the neck showed signs of atrophy, like in the lower back. <u>He was able to prove (later as part of his PhD thesis) that this does occur.</u> When these small segmental stabilizers go off-line, something else has to kick in to stabilize the neck.

In the neck and shoulder, the muscles that kick in are the trapezius, levator scapula, and scalenes (and sometimes jaw muscles). These muscles were never designed to be



stability muscles, so they quickly get overloaded. As they get tighter to stabilize the neck, they can bring one shoulder higher. This causes the attachments of the muscles to get angry, as they were never designed to handle this type of excessive loading. This is called enthesopathy and is discussed elsewhere. In addition, the upper trapezius and scalenes both have nerves nearby or traveling through them, so when they get too chronically tight, these nerves get in on the act. For the upper trapezius, the occipital nerve can get irritated, causing headaches. For the scalenes, the brachial plexus can get pinched, again leading to thoracic outlet symptoms with numbness in the little finger of the hand (ulnar or lower trunk of the brachial plexus distribution).

The Great Adaptation Machine

Our bodies were designed to keep moving at all costs and are great at adapting to problems that may develop in the human machinery. In a preindustrial society, the amount of physical prowess it took to collect, hunt, process, and consume food was great. In such a society, the potential for injury from a runaway animal or even a rockslide was also great. The only way for us to be able to get injured and keep going was to design the musculoskeletal system to be an adaptation machine. What does this mean?

At its simplest, let's take a left-foot injury. With an injured left foot, you instinctively limp on the left and transfer more weight to the right. This takes weight off the left so it can heal. This strategy works, because we were meant to heal on the run. For example, <u>studies</u> where patients are asked to bear more weight on an injured or operated area (different from the current non-weight-bearing orthopedic-healing paradigm) show that healing with weight bearing is better than extended periods without weight on the joint.

The great adaptation machine also gets a good deal more complex. I have noticed that in chronically injured patients, the system is constantly rearranging forces to be able to offload certain areas. An example is my own minor chronic neck, upper-back, low-back, and leg symptoms. At times, my left scapula will hurt; at other times, my biceps tendon; and at others, my low back. I can feel my body rerouting forces through adaptation, from one site to the next. When the neck stabilizers go off-line, or when they are too taxed from my heavy-weight-lifting routine, the big neck muscles take over, and the upper trapezius, levator scapula, scalenes, and SCM fire up. What happens when these muscles complain too much? My body reroutes the forces to the front of the shoulder by moving the scapula forward, but this aggravates the biceps tendon. If this causes the biceps too much pain, my body reroutes those forces by turning the rib cage, which causes the low back to get torqued, and so on. This complex neuromuscular response has allowed us for millennia to continue to function with injury.



For patients and medical practitioners, this adaptation process can often be like peeling back layers of an onion. Again, at its simplest (the injured left-foot analogy), since the left foot is causing a limp, it may come as no surprise to the patient that the right foot begins to hurt as a result of excessive use. However, most patients fail to recognize the more complex adaptations. This means that they are completely unaware that the problems in all of these areas are related. In addition, physicians will often only go for the "low hanging fruit" of where it hurts today. This approach again avoids the salient question, How did all of this get this way? In addition, just treating the part that hurts will only be a temporary fix as this part will soon be overloaded again!

How Do I Know if I Have an Alignment Problem?

First, I've only scratched the surface here concerning common problems with alignment. The goal was to introduce the concept, not list all things that an experienced musculoskeletal expert would see in daily practice.

At its simplest, patients with alignment problems have one-sided pain or arthritis in the absence of specific trauma. For example, while they may have both knees that hurt, the right hurts much worse than the left. In addition, an MRI or X-ray of both knees shows one has much more severe arthritis than the other. Examples of alignment issues can often be seen when looking in a mirror or asking your friends. You may notice that one shoulder is higher than the other, or the head is slightly tilted to the right or left, or that one hip is higher. Looking at wear patterns on clothing and shoes can give more clues. For example, does one shoe wear more than the other? Does one part of the sole of one shoe wear more than the other parts? Does one part of your pants wear out faster than another? Is it easier to hold a handbag or backpack on one shoulder or the other? When you're active, are you dramatically stronger on one side versus the other (more than you would expect based on being right or left handed)?

If I Have an Alignment Problem, What Else Can I Do About It?

The good news is that there are many therapists and practitioners who specialize in alignment. These concepts really began shortly after the turn of the century, when traditional allopathic medicine was in its infancy and unable to address what seemed like obvious problems to nonphysicians. The pioneers were <u>Moshe Feldenkrais</u>, <u>Ida Rolf</u>, and <u>Matthias Alexander</u>. I was introduced to these geniuses when I realized that by the early 1990s (just out of residency) these issues were still not being addressed by physicians. The concepts I've discussed here were not part of my training <u>in physical medicine and</u> <u>rehabilitation</u>. To remedy this deficit, I took to reading the old works of these masters to try and learn what I had never been taught as a physician.



Newer systems such as <u>Pilates</u>, <u>Muscle Activation Technique</u>, <u>Myofascial Release Approach</u>, and <u>Egoscue</u> have added to the diversity of treatment methodologies that address various aspects of posture and alignment. In addition, <u>curve restoration</u> has now become a scientifically vetted medical art.

A caution, while some physical therapists have spent years learning advanced biomechanics, they are few and far between. The standard course of physical therapy education contains very little about how to identify and address common alignment problems. This is despite one of the early geniuses of muscle function actually being a physical therapist (Florence Kendall). So if you've tried and failed physical therapy, it's unlikely that you actually saw a physical therapist with proper training in the art of biomechanical and alignment analysis and treatment.

Brief descriptions of the alignment concepts follow. Click on the links to learn more.

• <u>Rolfing</u>: Sounds a bit like the vernacular for vomiting, but it's actually named after the founder of the method, Ida Rolf. The focus is on very rigorous deep-massage techniques to free up areas of muscle and fascial tightness with the goal being to restore normal posture and alignment. This is generally accomplished in 10 sessions.

• <u>Alexander</u>: Matthias Alexander was a turn of the century orator in a time before the electric amplification of voice. He figured out that certain head and neck positions allowed the speaker to project his or her voice better in an auditorium. This was later applied to "sick" performers to improve their performances. This is now a system of treatment focused on head and neck alignment popular with stage and theater performers.

• <u>Muscle Activation Technique</u>: Developed by athletic trainer Greg Roskoph and based on the concept that certain muscles can become less active based on injury and certain patterns of movement, the focus is on balancing the moving biomechanics of the body by "turning on" these inhibited muscles.

• <u>Myofascial Release</u>: Pioneered by Arizona massage therapist John Barnes, the focus is on trigger point massage to release or free up tight muscles leading to poor body alignment. There is less focus on overall body posture than in Rolfing.

• <u>Egoscue</u>: Begun by Pete Egoscue, this system focuses on activating and strengthening specific muscles with specific exercises to restore normal body alignment and posture. This system has become popular with physical therapists wanting to increase their knowledge about biomechanics.



• <u>Feldenkrais</u>: Developed by Israeli physicist Moshe Feldenkrais, the focus is on alignment in simple movements.

• <u>Curve Restoration</u>: The gurus of this now scientifically vetted field are the Harrisons, chiropractors who have been publishing their results in peer-reviewed medical journals for years. They use very specifically designed forms of specialized traction to restore the normal curvature. They have also designed home units so that patients can try to deal with this problem in a do-it-yourself program.



Chapter 5: Neuromuscular



"My countrymen should have nerves of steel, muscles of iron, and minds like thunderbolt." Swami Vivekanda



For many readers, the term "neuromuscular" is a new term. To clarify, as it's used here, it means both nerves and muscles and is often used to refer to the connection between the <u>two</u>. While the nerves in various parts of the body tell many different organs what to do, the most visible organ they direct is muscle. Your nerve says, jump, and your muscle says, how high?



Think of the <u>nerves</u> as the wires that connect the main computer (the brain) with the <u>muscles</u>. You think of a movement, and the brain generates a nerve impulse that drives muscles. Information also goes the other way, from the skin, muscles, joints, ligaments, and tendons up to the brain. This information is called <u>"proprioceptive"</u> and allows you to finely adjust your movements to what's going on in your environment. For example, if you step on something unstable, you might fall. However, that information is quickly relayed to the spinal cord

where reflex patterns stored there instantly adjust your stance.

It's easy to see how the nervous system plays a big role in the type of muscular spinal stability discussed in the first chapter. As discussed above, this type of stability during movement is made possible by proprioception, which is used to provide real-time feedback so that a moving joint stays in finely tuned alignment. For example, if the joint experiences forces that might cause it to translate or shift too much, small joint sensors detect this motion and instantly tighten muscles to counteract that abnormal motion and keep the two joint surfaces aligned (keep the joint in the neutral zone). If this didn't happen thousands of times each day, the joint would wear out much more quickly.

What happens when this system of sensors, nerve impulses, and finely tuned muscle firing goes off-line? As discussed earlier in the "Stability" chapter, all heck breaks loose. In this chapter you'll learn that when spinal nerve irritation or compression occurs, the muscular stability system for the spine goes off-line, and the spine becomes unstable. I believe the same happens in peripheral joints, like the knee. If spinal nerves are irritated in the back (note: you may not feel any back pain), the muscles that help stabilize the knee in movement can go off-line or have reduced efficiency, and as a result, the knee joint becomes unstable. So now, when the knee experiences abnormal forces like a shift, the wiring loop through the spine between the joint sensors and the muscles that protect the spine is impacted, causing an ever so slight delay. This delay leads to a joint that gets out of alignment more easily during motion and as a result, a joint that is more likely to become arthritic. Since this concept of <u>muscular activation delay has already been very</u> well documented for spinal stability (here the delay causes the vertebrae to become unstable in movement), there is no reason to believe it only applies to the spine.


One of the problems we've had as a medical community is our main and most widely available test for diagnosing nerve pathology <u>(electromyogram [EMG]/nerve conduction study [NCS])</u> is very specific for certain types of nerve injuries (such as when a nerve is

wholly or partially destroyed by trauma) but not very sensitive for other types of nerve problems. In particular, many significant problems with the nerves involve small fibers (small-fiber neuropathy), whereas the EMG/NCS test can't detect this type of pathology. In addition, the test has very poor sensitivity in detecting nerve irritation. While other more sensitive nerve tests (in particular, quantitative somatosensory tests [QST]) are commonly used in research, they are not yet widely used by physicians. So in a real way, physicians are often "flying blind" from a diagnostic testing standpoint in figuring



out when nerves are sick. In other words, based on the research, a negative EMG/NCS doesn't rule out nerve trouble. It's a test that's <u>highly specific, but with poor sensitivity</u>. The definition of a good diagnostic test is one which is highly specific and highly sensitive (capable of detecting the disease 99% of the time when it's present, and shows negative 99% of the time when the disease isn't there).

So in summary, even small amounts of spinal nerve irritation may not cause any noticeable back or neck pain, but it can wreak havoc with the muscular stability system either in the spine itself or in your joints. Since this system protects your joints during activity, when this type of nerve problem takes muscles off-line or reduces their efficiency, this will eventually lead to less protection for the joints and an earlier onset of arthritis. In addition, the diagnostic-test toolbox we have available to us today doesn't include tests that are capable of detecting this type of nerve problem, hence the reason this problem often goes undiagnosed. In addition, I believe that treating this problem is a key component of longterm joint preservation.



Take This Simple 10-Item Test for Nerve Problems

1. I have numbness, tingling, burning, or electrical sensations. Yes / No

2. I have chronic tightness that feels like pressure in my arm or leg with certain activities? Yes / No

3. I have a chronically tight muscle that just won't "let go" no matter how hard I stretch. Yes / No

4. I have pain in my wrist area whenever I reach for something. Yes / No

5. I have pain in the back or bottom of my heel that won't go away. Yes / No

6. One arm or leg always seems to be significantly weaker or smaller than the other, no matter how hard I try to strengthen it. Yes/No

7. My arm or leg feels a bit "goofy" or uncoordinated after I do certain things. Yes/No

8. I have an area of chronic pain that just won't go away no matter what I try. Yes/No

9. I have chronic neck or back pain. Yes/No

10. I had a back or neck problem years ago, but it seems to be fine now. Yes/No

If you answered yes to any of the above, you may have a nerve/muscle problem and not know it. Questions 1 about numbness is more obvious. Most people associate these symptoms with nerve problems. Question 2 isn't so obvious. When there's pressure on a nerve in the neck or back, many patients don't necessarily feel neck or back pain, but instead feel pressure in the muscle that is supplied by that nerve. Some patients describe it as feeling like a blood pressure cuff is pressing on the muscle.

Question 3 is also not obvious. That chronic hamstrings or groin tightness you've been blaming on being out of shape could actually be caused by a pinched or irritated nerve. Question 4 is really interesting. We commonly see this when there's scarring around the median and/or ulnar nerve. Reaching out to get something places the nerve on stretch, and since it's scarred, it can't move with the arm, which causes pain where the nerve is scarred down. How about question 5, the heel pain? Most patients with plantar fasciitis would think it must be caused by something in the arch of their foot. However, we see patients every day who have S1 nerve problems in their back or a pinched tibial nerve at the ankle who have this problem caused by an irritated nerve.



Question 6 may seem more like something associated with a nerve, but the weakness or atrophy (smaller muscles) I'm talking about is where one arm or leg is notably smaller, not the kind you see in paralysis. Question 7 is an extension of 6, as sometimes patients with nerve problems note that their arm or leg seems uncoordinated. Questions 8 and 9 are connected, in that a bad nerve can cause pain just as readily as it can cause numbness, tingling, or burning. Finally, it's important to note in question 10 that many patients who no longer believe they have any back or neck problems because the pain has gone away still have bad nerves that cause problems in joints, muscles, and other areas.

Arthritis Doesn't Cause Pain; Pain Causes Arthritis

<u>I saw this title come across a science news feed a few years back.</u> It hit me like a unwelcome pie in the face, as I had often suspected that something like this had been happening in my patients. The concept is simple, yet it will change the face of orthopedics and rheumatology forever. What was discovered? That irritated nerves can cause bad chemicals to dump into joints, which leads to cartilage breakdown.



Since that time, many other articles have been published confirming this link between bad nerves and bad joints (see <u>here</u>, <u>here</u>, and <u>here</u>). This discovery is equivalent to when we doctors learned that stomach ulcers were caused by bacteria and not stress (I was taught in medical school ulcers were due to stress).

How can arthritis be caused by pain? The authors created an elegant animal model that showed that nerve activation in a joint leads to bad chemicals being dumped into the joint, which leads to pain and faster onset of joint arthritis. This is a reverse of what

has traditionally been considered (i.e., that a joint is injured and begins to degrade and then causes pain). It's important to stop for a moment to consider how these scientists have turned orthopedics on its head. Again, our entire orthopedic care model is based on

the concept that injury in a joint (or accumulated injuries over a long period of time) leads to arthritis in the joint, which leads to more joint breakdown and pain. This new model reverses the old paradigm so that now it's aggravated nerves that lead to arthritis. Sound familiar? I believe this is just an extension of what we've been discussing here: problems with spinal nerve irritation lead to bad chemicals being dumped into a joint and a "sloppy" joint with poor stability, which ultimately leads to arthritis.





I have had my own knee problems caused by my back. Using this new model, my knee problems were caused by spinal nerve irritation (which I never perceived as low-back pain) causing not only a sloppy knee joint (due to parts of the big stabilizer muscles being shut down by <u>trigger points</u>) but also bad <u>catabolic</u> (breakdown) chemicals dumping into the joint. This issue was quickly fixed not by operating on my knee or even injecting magic stem cells into the knee, but by bringing the spinal and joint stability systems back online by using IMS to get rid of the trigger points. What are trigger points? Read on.

Low-Level Arthritis Pain vs. Nerve Pain

Based upon my clinical experience and this new model of nerve-related joint pain and arthritis, I would place patients into two distinct categories: what I'll call "neuropathic arthritis" versus "classic arthritis." Early on in the degenerative process, and for some patients who have more of a spinal component to their joint pain, patients are firmly

in the "neuropathic arthritis" (NA) camp. These patients have severe joint pain that is often disabling or can become disabling with certain types of



activity. I see these patients in the clinic, often very desperate because their joint pain is very intrusive. They are either completely disabled by their pain or they are unable to exercise at high levels.

In this new model of joint pain, these patients have an active spinal nerve problem manifesting as joint pain. They are often unaware that this joint pain is linked to their spine, but if you dig enough, they will usually admit to a history of spinal problems that have either (in their mind) been successfully treated (perhaps with a surgery many years in the past) or are ongoing, but the pain is low level and (in their mind) under good control. They have usually had several unsuccessful joint surgeries, which didn't work because while they have issues in the joint, they also have active issues in the spine that were unaddressed by their joint surgeries. Treating the spine in these patients can often make a huge difference.

The second camp is the traditional "classic arthritis" (CA). The CA group no longer has an active spinal component, or if they do, their arthritic joint has long since degenerated. Their pain pattern is different and matches what we know of arthritis pain. You may remember your grandparents being stiff in the morning with low-level pain that became better with activity as the joint "warmed up." Just like gramps and granny, once these patients start moving, they generally feel better. Treating the spine in this group is often too little too late, as the joint damage is done.



It's important to note that there are other factors at play in many of these patients, so this is a simplified discussion. For example, patients with joints that are unstable from a ligament standpoint may also have more pain when they are active, and patients with bad joints due to severe trauma may have less pain as their joint warms up. Like anything in medicine, the body is a very complex machine, hence the SANS approach, which looks at all components of the musculoskeletal system.

So What Can Be Done to Fix the Spinal Nerve-Joint Connection?

Despite research showing that irritated spinal nerves may be associated with joint problems, most physicians have a hard time associating joint pain with a low-back nerve problem. The first step in identifying a spinal nerve component as a cause of joint pain is simply a thorough neurologic exam. When I say a complete exam, I don't mean the "Can you feel this?" type of neurologic exam. This careful exam is focused on comparing sensation from side to side and on the same side, testing multiple different types of sensation. This includes not only light touch (the can you feel this exam) but also pain sensation (pinprick) and perhaps hot/cold sensation. The exam also recognizes that there are multiple types of pain and nerve referral patterns, including those from <u>spinal joints</u>, <u>nerve trauma</u>, and <u>muscle trigger points</u>.

If the exam shows that spinal nerve irritation may be occurring, the next step is a spine MRI. Correlations between the exam findings and the MRI are important. In addition, this correlation acknowledges that while spinal nerves can be compressed by bone spurs and herniated/bulged spinal discs, they can also be irritated by sloppy stability in the spine (see the "Stability" chapter). An MRI marker of this type of sloppy stability can be seen on MRI as multifidus muscle atrophy (see the "Stability" chapter). So even though there may be no bulging disc on the spinal nerve, significant atrophy of the deep stabilizers at this level (multifidus) combined with sensation problems at the nerve in the leg means that the spinal segment is likely sloppy from a muscular stability standpoint.

Treatment for Irritated Spinal Nerves

Your spine has discs, which act as shock absorbers. The diagram to the left shows that the <u>spinal discs can herniate their inner contents</u> (nucleus pulposus), which can place pressure on spinal nerves. This is called <u>radiculopathy</u> (if more severe) or <u>radiculitis</u> (if less severe). This has also been called <u>"sciatica,"</u> although this is not an accurate term. The traditional solution is to surgically remove the herniated portion of the disc sitting on the <u>spinal nerve</u>. In the 1940s, this was a great advance. Patients with numb and weak legs due to a bad back now had a treatment. <u>On the other hand, it began our current move toward</u> <u>invasive spine treatment</u>, a path that many have criticized.





Treatment for herniated discs remained largely surgical until studies in the 1980s showed that a few years after the disc herniation, results for patients treated with surgery weren't all that different from those who didn't get surgery. What emerged from this research was the revolutionary idea that perhaps herniated disc patients could be treated without surgery. As a result, the focus began to shift toward conservative management, which by the late 1990s included <u>epidural injections</u>. This meant placing strong anti-inflammatory medications around the painful and swollen spinal nerves. At first these injections were just adapted from pregnancy epidurals, which are usually provided to control the pain of labor, but later these evolved to more specific placement of medication between the disc and swollen nerve. These newer injections were called transforaminal injections (literally meaning through the <u>foramen</u>, or into the hole in the spine where the spine nerve exits). Transforaminal epidural injection types (interlaminar and caudal).

The conventional wisdom regarding epidural injections is that we wait until the patient has failed a significant course of medications, physical therapy, and time. While this makes some sense from a cost-savings standpoint, our newer understanding that irritated spinal nerves can lead to shrunken spinal-stabilizing muscles and instability means that earlier intervention is likely warranted. I believe that preservation of these spinal stabilizers is very important and that the goal of treatment is to bring them back online early by calming down swollen and irritated spinal nerves.

My own example is important. Several years ago, I was performing heavy dead lifts from the floor early in the morning (generally a bad idea). I felt a pop and went down on the ground. My pain was so severe that I couldn't walk, stand, move, and so forth. Had I stayed like this for any length of time, my spinal stabilizers would have quickly atrophied, leading to months of rehab to get back into shape. Instead, I had my partner perform an immediate epidural to calm these swollen nerves and was back to weight lifting three days later! This is what I mean by "early intervention."

The Paradigm Changes Again: Epidural Steroids Are Toxic

You may have seen that steroid epidurals have been all over the news of late. First, there was a <u>compounding pharmacy contamination tragedy</u> that led to many sick patients because the steroids being injected weren't sterile. Next came the revelations that <u>the steroids used in these shots caused women to lose bone mass at an alarming rate</u> (worsening osteoporosis). Other studies have shown that the <u>steroids can also shut down</u> important brain responses that help you deal effectively with stress. The conclusion? Injecting these high-dose steroids into the back or neck can cause significant side effects, but it generally works.



Is there another way to get the same or a better result that avoids steroids? Rather than big dose anti-inflammatories, we believe that the next generation treatments will be using much lower dose medications and combining these with regenerative medicine. So rather than injecting 80 mg of corticosteroid (the height of the Empire State Building), injecting smaller physiologic doses of steroids and adding in level II (or even level III) regenerative medicine solution.



To do this, we have pioneered using the growth factors from the patient's own blood platelets (platelet lysate) instead of these toxic steroids. So rather than using a sledgehammer to treat the swelling (steroids), we use growth factors that can reduce swelling and improve blood flow. Does it work? We tracked 232 patients in a registry and compared patients getting traditional steroid epidurals (85

people) and platelet lysate (Regenexx-PL-Disc procedure—147 patients). The function questionnaires (functional rating index [FRI]—a higher score is better) showed more improvement in the platelet epidural patients than the steroid epidural patients (see graph to left).

Can We Use Platelet Lysate Injections to Fix Damaged Nerves?

In medical school, I was taught that while a damaged nerve might be able to regenerate a little bit over time, most nerve injures were permanent. Recently, one of our physicians, Ron Hanson, MD, investigated whether nerves might be helped by injecting the nerve. The simplest form of this procedure is called <u>nerve hydrodissection</u>. Basically, fluid is injected around the nerve using very strict ultrasound-imaging injection protocols. This frees the nerve up from any scarring that may cause pain. Next, platelet lysate is injected to help the nerve heal. We've seen some very interesting results in select patients with everything from <u>severe chronic nerve pain (complex regional pain syndrome [CRPS])</u> to more specific <u>nerve damage</u>. We remain cautiously optimistic and will continue to move forward trying to help these patients who have few good options.



Making the Transition from the Nerve to the Muscle

You've learned about the nerves. Now you have to learn about where the nerves go the muscle. We like to think of these as separate structures, yet one look at the former Superman actor <u>Christopher Reeve</u> in his wheelchair shows they are not. A spinal cord injury in his neck shut off all nerve input to the muscles. What happened? He went from being one generation's Superman to being shriveled up. Why? The nerve and the muscle are one continuous structure. What happens to your nerves directly and immediately impacts the muscles.

The Nerve-Muscle Hotspot: Trigger Points



<u>Trigger point injections (TPIs) were first popularized by Janet</u> <u>Travell, MD</u>, one of JFK's physicians. Janet's techniques made it into popular medical culture because JFK had a bad back that often responded well to her trigger point injections with numbing medicine.

Sometime later, Canadian neurologist <u>Chan Gunn, MD</u> added a significant piece to the trigger point puzzle. Travell had noted that just using a needle without injecting anesthetic (dry needling) seemed to work just as well. To the medical

establishment of the day, this seemed like voodoo. At the time, Chinese acupuncture was largely unknown in the West, so Travell largely placed her emphasis on injecting anesthetic and anti-inflammatory medications. Gunn grew up in Korea, where a more aggressive form of Korean muscle acupuncture was common, so he moved forward with Travell's dry needling technique, substituting the much finer and less traumatic acupuncture needles for the more traumatic cutting-edge injection needles used by Travell. Gunn also theorized that the muscle trigger points that Travell thought were due to overuse were more likely caused by nerve irritation.

The science of the last 20 years supports Gunn's theory that nerve problems (autonomic and likely spinal nerve) and trigger points are closely related. By the early 1990s, while physicians who were experts at trigger point injections were few and far between, TPIs were used by a plurality of doctors treating musculoskeletal pain. Then something happened that often drives the course of medical care much more than efficacy or science—the reimbursement changed.



Prior to the mid-1990s, a physician could receive adequate compensation per site injected. After the mid-1990s, the average compensation for this procedure was reduced by about 70–90%! In addition, getting compensated by insurers became more difficult. This was all it took to relegate the art of trigger point injections to the history books. Today, because of this reimbursement collapse, finding a physician experienced in managing this type of muscle pain is like finding a needle in a haystack (pardon the pun).

I think my own personal story is important. In the late-1990s I attended a medical conference that involved days of sitting. For an unknown reason, my left knee began to ache and swell. There was no trauma to the knee. I was literally hobbling around the office, and all of my aerobic exercise came to a screeching halt. I underwent an MRI, convinced that I had somehow torn a meniscus or some cartilage. While the MRI showed the swelling and perhaps some questionable small tears in the meniscus, it didn't show a "smoking gun" cause for my severe pain.



I went to see an orthopedic surgeon who wanted to perform a diagnostic arthroscopy, likely chop out some meniscus and remove a "plica." I was desperate and convinced the MRI was missing the true cause, so I reluctantly signed on for surgery. A few days before the planned surgery, a visiting doctor from Canada was in our clinic and asked if I had tried trigger point therapy in my quadriceps muscle and low back? I said no, looking at him like he was some alien speaking in tongues. At this point, I had seen the best physical therapists in town

and failed all of their exercises, so I was desperate. I told my Canadian colleague that I would try anything.

Turns out, this visiting physician was one of those "needles in a haystack" as he was experienced in the Gunn trigger point technique (called IMS, short for intramuscular stimulation). He examined my thigh muscle (quadriceps) and my low back, pulled out an acupuncture needle, and proceeded to stick this in my low back and thigh muscles. The muscles cramped suddenly as the needles hit the trigger points (more strange than painful). After a two minute treatment, I got off the table and walked normally. That night, I went running for the first time in months, without a



twinge. I canceled my surgery and have never looked back. I was so impressed, I learned the technique and began using it in patients.



IMS has revolutionized our practice, providing relief to patients who would only otherwise be treated by much more invasive treatments. Because of reimbursement issues (insurers don't generally cover IMS, and the other form of trigger point therapy [TPI] is poorly reimbursed), the technique has remained obscure. There may be other reasons the technique has never moved to a wider physician audience, as it takes significant effort and dedication to learn how best to apply the procedure to get consistent results.

At a medical conference where both traditional Chinese acupuncture and IMS were being taught, I had insight into how my medical colleagues view this complexity. After Dr. Gunn lectured about IMS, I turned to the physician sitting to the left of me and asked, "Wow, isn't this IMS stuff great?" Her response was, "It's too complex. You have to learn where all the muscles are, what they do, where to put the needle for each one, what to avoid...With traditional Chinese acupuncture, I just look at a chart on the wall and put the needle at X marks the spot."

So while <u>traditional Chinese acupuncture</u> (placing a needle into the skin at specific Chinese chi points) has become popular, IMS has remained in obscurity. These past few years, IMS has finally taken a leap forward by being adopted by various Colorado physical therapists (PT). One of our former PTs who we had trained in IMS went through the red tape to allow physical therapists to widely practice the technique after very intense coursework. As a result, IMS is now gaining more acceptance, and more patients are getting more access to the technique through physical therapists.

Enthesopathy

We take for granted that our muscles not only contract but also have a function as shock absorbers, letting go in a controlled fashion. As an example, when you jump from a fence at a height of just four feet, your femur bone should break. Why doesn't it? The big quadriceps muscle absorbs the shock by acting an <u>eccentric contraction</u> (controlled release). When a muscle has trigger points, the biomechanical properties of that muscle change. Large sections of the muscle can lose their ability to act as active shock absorbers. We believe this leads to extra pull on the areas where the muscles attach to the bone. This causes swelling and breakdown of these areas known as _

Enthesopathy means that where a tendon attaches to bone, the tendon is aggravated and swollen. If this goes on a long time, you may see small tears in the ligament. If it gets really severe, you may see bigger tears. All of this can cause pain.





While many physicians will recognize problems in joints, and fewer will recognize trigger points in muscles, in our experience, even fewer will recognize enthesopathy. This is a problem, as many patients suffer from this problem. The good news is, a new generation of physicians armed with PRP to treat tendons and ultrasound imaging to detect problems in those tendons is finally beginning to address this important problem.

What Are Common Spinal-Associated Areas of Enthesopathy?



Head: The back of the head where the trapezius, sternocleidomastoid, and suboccipital muscles insert is frequently an area of muscle attachment overload. These muscles can irritate the greater and lesser occipital nerves, leading to headache.

Elbow: The classic medial and lateral

epicondylitis (golfer's and tennis elbow) are attachment sites for muscles that get overloaded. We see this commonly in patients who have nerve irritation in the neck or shoulder (cervical radiculopathy or radiculitis and thoracic outlet syndrome).



Upper back: The back of the ribs where the iliocostalis and quadratus lumborum muscles insert can become inflamed and lead to back pain. We see this commonly when the multifidus stabilizers are off-line.

Lower back and pelvis: The posterior superior iliac spine (PSIS) area is where many large low-back muscles take their anchor. As already described, we consider this the tennis elbow area of the low back. Patients can often point to a single spot just to the side of the upper tailbone (dimples of Venus). This spot gets overloaded as the stabilizer muscles go off-line.



Tennis Elbow of Low Back





Ischium ("Sit Bone"): This is an area where the hamstrings muscle attaches, as well as many other important deep pelvic muscles. Patient's with enthesopathy here have trouble sitting or have deep buttock's pain. This is more common in patients with a lower lumbar nerve irritation and/or problems with sacroiliac (SI) joint instability.

Posterior/Lateral Hip Girdle: This is one of the most common areas of tenderness in women (and many men) who have problems in the SI joint or with lower spine nerves. These powerful buttock muscles attach to the bone here and travel south to the back and side of the hip.





Knee–Pes Anersine: This area is a common insertion

for many muscles that travel in the front of the thigh, including the sartorius, gracilis, and semitendinosus. These patients have pain on the inside of the knee that is below the joint and is often confused with meniscus tears. We see this problem in patients who have low-back issues leading to knee pain.



Knee—Hamstrings: The hamstrings are muscles that make up the back of the leg and attach to the back of the knee. They're responsible for helping to set the position of the bottom knee bone on the top thigh bone and for helping the knee meniscus position itself correctly. They can become painful at the back inside or outside of the knee. This pain is often confused with a torn meniscus. We frequently see this happen when the S1 nerve in the low back is fired up. Patients may not recognize that a back problem is causing the pain at the back of the knee as all they may feel is that their hamstrings are chronically tight.

Foot—Plantar Fascia: At the bottom of the foot near the heel is a strong fascia that extends from this area to the base of the toes called the plantar fascia. This area helps to support the arch and acts as a shock absorber for the foot. Where this tough fibrous structure originates in the heel can develop enthesopathy. In particular, we see this happen quite a bit when patients have an S1 nerve problem in their low backs.



What Happens When Tendons Have Bigger Problems?

We've discussed what happens when tendons attached to bone are overloaded and aggravated. What happens when they fail? For partial tears, in our experience, platelet rich plasma (Regenexx-SCP) works well. For complete tears, where the pieces haven't retracted back like rubber bands, same-day stem cells tend to work well. For full-thickness retracted tears, where the ends have snapped back, surgery is likely needed (although we're working on nonsurgical options). Tendons can get partial or complete tears like ligaments. Here is a list of the most common ones we see related to the spine:

> **Iliopsoas:** This muscle starts in the low back and travels frontward toward the inside of the hip. It's responsible for helping to lift the leg. The tendon can become irritated or get partially torn if the front of the hip is overloaded, for example from too much curve in the low back. It can also be injured in kicking sports. The tendon can also snap over the hip. A common cause is problems with upper low-back nerves that provide nerve supply to these muscles.

Hamstrings: The hamstrings muscles at the back of the leg start at your buttocks area and travel downward. They actually originate from the ischial tuberosity (the bone where you sit). This is an area where they can become torn or just irritated. This can cause pain in the buttocks area as well as pain or tightness in the back of the leg. The lower lumbar nerves in the back can also cause problems in this muscle and tendon.

Iliotibial Band (ITB): This is an area at the side of the hip and leg that begins from the top of your hip and extends down to the knee. Patients with chronic instability in the sacroiliac joint tend have problems here. The ligament is rarely torn, but can be irritated.

Achilles Tendon: This is the thick tendon at the back of your ankle and heel. It connects your calf muscle to your heel, so it's responsible for pushing you off as you walk. It can get torn in middle-aged and active elderly patients, usually during sports performed beyond current fitness level. It can have partial tears inside the tendon or can tear completely by snapping back like a rubber band. It's also very common in patients who have S1 nerve problems in the low back.

Peroneal Tendons: These tendons on the outside of the foot and ankle are involved in stabilizing the foot when you're balancing on one leg or when you're on one leg while walking or running. They can become torn (commonly they can split down the middle). Usually this isn't from a specific injury, but from wear and tear. This problem can be associated with instability in the ankle ligaments as the tendons try to stabilize the foot and

ankle, something they weren't designed to do. They can also become torn when there is a chronic lower lumbar nerve problem leading to abnormal firing of the peroneal muscles.















Extensor Hallucis Longus (EHL): This is the tendon that raises your big toe. In patients with chronic foot and ankle issues, it can become misaligned so that it no longer travels across the main toe joint but is displaced more toward the inside. It can also become raised and overloaded and sore/painful, especially when walking or hiking. This tendon is commonly impacted when the nerves in the low back are chronically irritated, and as a result, the position of the foot and ankle changes. See this link for further discussion and diagrams.

How Instability and Enthesopathy Are Often Linked

You've learned about instability; now let's learn about how an unstable joint often leads to "pissed off" muscle attachments. Basically, in this case we have muscles that are pulling too hard on their bony attachments. How does this work?

Have you ever tried to stand up in a row boat? The "ground" you're standing on is unstable, as it seems to randomly move in unexpected ways. The amount of energy it takes to do anything, even reaching for an oar, is tremendous. The same happens when a joint is unstable. In the case of your body, when a joint is unstable the muscles automatically kick into overdrive to compensate and stabilize the joint. When your muscles do this for long periods of time, the places where they attach to bones can get irritated.



For example, this commonly occurs when the spinal stability system goes off-line. Look at the picture of the spine and note just some of the many spine muscles that act between spine bones (from one vertebra to the next). Imagine if all of these were working overtime because the spine bones were unstable due to a bad ligament. The places where these muscles attach would be on fire, as these attachments would be overloaded (enthesopathy). This is why many spine-injection approaches often fail, as medical practitioners fail to inject these areas in favor of nerves and joints. For example, we can inject medications into a specific structure like the facet joint or around the nerves, but if these tendons attached to these small

segmental muscles have been damaged due to years of overuse, then placing medication in a joint or around a nerve (which are different locations than these muscle attachments) won't help the pain. <u>We also frequently see this around the SI joint and hip girdle (see</u> <u>Case 1: SI Joint Shirley)</u>.



Multifidus Atrophy

<u>In the first chapter</u>, you were introduced to spinal instability. This often happens because the muscles that stabilize the spine become weak. How does this happen?

If you have numbness, tingling, or weakness in an arm or a leg, you meet the classical definition of spinal nerve root compression called <u>radiculopathy</u> or <u>radiculitis</u> (literally in English, root disease and root swelling). Getting to a diagnosis in this instance is usually routine, as long as there is something on your exam that correlates with a pinched nerve on your MRI (bone spurs or disc pressing on a spinal nerve). However, if you have more subtle signs of nerve root irritation or nothing structural on your MRI, you're less likely to get a diagnosis.



You also learned in the first chapter that your MRI may be harboring "secrets" (or at least important information that the radiologist neglected to read). If you look at your MRI, you may be able to see this problem yourself. This finding is easy to see if you think of the spinal muscles as a steak. A steak in the grocery store is a cross section of the muscles, just like the axial view of an MRI. If you find the axial view of the spine and open those images, in the low back (where this finding is commonly seen), it will look like the image to the left. The multifidus muscles will

be in the back of the spine (to the bottom of an MRI image). This area looks like a steak.

More Details on How to Read Your Own MRI Images for Multifidus Atrophy

Open your MRI CD. You will see different "sequences." These are different parameters for how the MRI was taken. There may be five to seven different ones listed. First, choose the axial T2 sequences (these might be labeled AX T2, Axial or Ax STIR, or AX T2 FSE). If you don't see "T2,[™] just pick the axial or ax sequence that looks like the image below, with a whitish circle in the middle (the fluid in the spinal canal) and lots of white to the left and right of the spine (your love handles).

These images usually start at the top of the sequence (upper part of your low back) so take a look at the dark areas in the back of the spine (the middle to bottom of the image behind the white circle). This area looks very much like a steak you would buy at the grocery store (see image above). Why does it look that way? Because a steak is a cross section through muscle, and so is this picture.





Now scroll down through the spine, paying attention to the steak part. Look for what I call "the bad steak," or where the muscles look fatty (see the image below—you want to look in the white-dashed area for whitish coloring in the otherwise dark muscles). As you start to get to areas where multifidus atrophy shows up, you'll likely be toward the bottom of the spine (higher image numbers in this sequence).

To help guide you, the illustrations below are from one of the original research papers that first described this problem (authored by Kader et al.). Note that the area in the white-dashed circle has a bit of white (fat atrophy) in the mild picture, more white in the moderate, and is 50%-or-more white in the severe. We routinely see patients in our office with 80%-90% white in this area by the time they see us. Again, look at all levels on the axial images for these white areas.



Why is this important again? Multifidus atrophy has been associated with both chronic low-back pain and leg pain. Why? The muscle acts as a stabilizer of the vertebra (as discussed in the first chapter). If it gets weaker and smaller, it can't stabilize as well, and this can lead to nerves getting irritated.



The Tennis Elbow Spot of the Low Back

Long back extensor muscles like the quadratus lumborum and ilicostails lumborum that attach to the ribs, start substituting for the mutifidus being off-line. This causes the spot where they originate (PSIS-here the blue circle area) to become chronically swollen.

After many years of examining patients, I can tell you there's always one spot that's tender in about 60% of all low-back pain patients: the posterior superior iliac spine (PSIS). Why? When stability problems manifest in the low back, we tend to see the big muscles that attach the pelvis to the ribs substitute for the smaller muscles. This includes the guadratus lumborum and iliocostalis lumborum, both of which originate at a common attachment point in the low back on the pelvic bone (see figure). This is a similar situation to the lateral and medial epicondyles of the elbow, where the muscles of the forearm attach and can cause tennis elbow or golfer's elbow. This common low-back attachment site is the <u>PSIS</u>. We consider this the tennis elbow site of the low back. This spot is the dimple (also called "dimples of Venus") just on either side of the upper tailbone. This type of chronic pain due to tendon pulling is called an <u>enthesopathy</u>, which is better explained in <u>a later chapter.</u>

Really Bad Nerve Problems...

Problems at the Neuromuscular Junction: Painful Dystonia



The <u>neuromuscular junction</u> is where the nerve talks to the muscle. As you recall, the nerve talks to the muscle to tell it what to do. The nerve says, jump, and the muscle says, how high? Think of the neuromuscular junction as a room where one person is talking to another. The nerve does the talking, and the muscle listens. If the room is quiet, the instructions barked by the nerve to the muscle can be clearly understood. If the room is very noisy or the nerve isn't speaking clearly, the instructions may be misheard or misinterpreted by the muscle.

The latter case is what happens in <u>cervical dystonia</u>. This is a nerve disease that when it's severe results in a patient with spasmodic torticollis, or where a patient has his or her head turned to one side on a permanent basis. However, like all diseases, this problem comes



in less-severe forms, where the patient just has constantly tight muscles that won't let go. This chronic tightness (called dystonia) naturally changes the alignment, which can lead to pain and other problems. I call this less-severe form, painful dystonia. We see this in patients who have had traumatic nerve injuries (often in car crash trauma). In this case, the noise in the neuromuscular junction room is turned up, so that the nerve instructions to the muscle sound like "stay tight all the time," even when that signal makes no sense for the situation. This can be treated with medications like <u>Myobloc</u> that work to turn down the noise in the room so the muscle can hear the nerve instructions. Injected in enough quantity, this medication could turn off all signals from the nerve to the muscle, but injected in smaller quantities, it only reduces the noise in the room.

Central Sensitization



Let's say you're in your car and suddenly all of the warning lights start to go off. You take the car to the mechanic, and he or she says that there may be a few things wrong here and there with the car, but the real problem is that the wiring is bad. This is central sensitization (CS) also known as complex regional pain syndrome type II, fibromyalgia, neural sensitization, and so on. In all of these conditions, it's an injury to the pain-reporting wiring of the body (the nerves and microprocessors that control them) that causes

the problem. The nerves become hypersensitive to pain. This phenomenon has been extensively published—most references are for <u>whiplash</u> or <u>fibromyalgia</u>. <u>This problem is</u> <u>also now being discussed as related to joint pain (as discussed earlier)</u>.

Patients with CS simply have a nervous system that's on fire. At its early stages, it may cause arthritis (see above), but as this gets worse, large areas of the body can be impacted. In addition, these areas don't follow normal nerve pathways like dermatomes (skin areas associated with certain spinal nerves), so oftentimes many physicians without training in this area label these patients as having "nonanatomic" sensation problems. These patients, as they progress, can't tolerate physical therapy, massage, injections, acupuncture,



IMS, and so forth. <u>Our research group demonstrated that at an early stage, trigger points</u> <u>may make the sensitization problem worse.</u> At later stages or when more severe nerve injury has occurred, cold sensitivity is common. <u>For example, for patients with traumatic</u> <u>CS, a cold summer's night (about 60 degrees Fahrenheit) is actually painful, as that's all it</u> takes to activate pain nerves.



Think about this for a second: How cold would it have to be for a normal person, you, to perceive cold as pain? Below freezing? Twenty below? These patients feel this at less than 60 degrees.



CS patients are generally the most difficult patients to treat. First, the pain sensitivity levels have to be brought down to a more normal level. One way to do this is medication. We have seen many medications for this type of nerve-related pain come and go—<u>Neurontin, Tegretol,</u> <u>Elavil (amitriptyline), Doxepin, just to name a new. They</u> all had the problem in that they didn't work for most patients. However, newer nerve pain drugs are just coming

to market, with many new ones in the pipeline. The most effective drug we have seen is the newer drug <u>Lyrica</u>. This works well in about 6 in 10 of these patients to reduce nerve pain and "put some water on the fire."

Once this is accomplished, the next step is usually to identify the problems that caused the fire. In many patients, there were specific musculoskeletal problems that led to others, which ultimately led to the fire getting out of control. Finding these specific problems and treating them can then start to provide relief. As an example, a patient labeled with "fibromyalgia" may note that his right neck and shoulder began hurting first, then his right low back, then his arm and leg. Tracing the issues back to the neck would be the way to approach this patient.

In addition, we are also encouraged with the results in using the nerve-hydrodissection technique with platelet lysate, as discussed earlier. This approach is to try and help the nerve function better by placing helpful platelet-derived growth factors near the nerve.

Neuromuscular Resources

Calming down nerves through injection often requires an expert trained in X-ray-guided procedures. Here are some resources:

- SIS (Spine Intervention Society
- ASIPP (American Society for Interventional Pain Practitioners)

Realize that almost all of these physicians still use steroid epidurals as their primary injection type. If you're interested in platelet lysate epidurals, see the <u>Regenexx Network</u> physicians at this link.



The most effective way we've seen to address chronic trigger points is either through IMS or trigger point injections. Here are some lists of where to find these "needle in the haystack" doctors and physical therapists:

- <u>A list of Gunn IMS practitioners</u>
- <u>Trigger point educational group</u>
- Physical therapists trained in IMS

Trigger points in muscles can be difficult to treat on your own, but we've seen some success with these approaches:

- <u>Electro Therapeutic Point Stimulation (ETPS)</u>
- <u>Thera Cane</u>

Enthesopathy: See <u>level 1 prolotherapy</u> and <u>level 2 PRP</u> resources discussed in the "Articulation" chapter.

Getting Out of the Hole

The goal of the rest of this book is to provide additional information for those patients who want to go further in their understanding of their problems. To go from an educated participant in their own recovery to a leader of that recovery. I've only had a few patients make it this far in their understanding. One gentleman who comes to mind was an engineer from Canada who moved to Colorado to get care from our clinic. Over time, our office visits became more of me providing advice about his next steps and what might be wrong rather than the doctor leading the patient. At first it was a bit disorienting, but later it became fun.

If you're a patient who can't do much without things badly flaring up, this section is for you. First, you need to have read this whole book to understand what's wrong. Next, you'll likely need to make some hard choices and follow the process diligently. I call this "digging out of the hole."

What is "the hole"? Your body has all or most of these systems we've talked about seriously involved. Your stability is fried, your joints are beginning to or have already given out, your nerves and muscles are shut down and on fire, and your symmetry is all catawampus. It may take you one to two years to get all of this addressed and to "climb out of the hole."



Where Do You Start?

1. You must get "off the sauce." We've seen an explosion in the prescription of narcotics by doctors <u>after pharmaceutical companies claimed that newer long-acting and very</u> <u>addictive narcotics weren't addictive.</u> Narcotics also take away your natural ability to control pain. So while they may take away the pain in the short run, they make the <u>pain signals magnified in the long run.</u> As a result, you have to decrease and eventually eliminate your use of narcotics.

2. **Reduce the pain** coming from various joints, tendons, muscles, and nerves. These problem areas must be first carefully identified. Often this will take an hour or more of hands-on exam, combining many different types of imaging, including ultrasound, MRI, and movement-based studies. Once the areas are identified, then the focus should be on precise biologic injections to ramp up the healing response in these areas.

3. Get rid of muscle trigger points caused by irritated nerves using once or twice a week IMS. This will allow your muscles to begin to participate in providing stability.

4. **Slowly work on getting more stable and stronger.** This may at first be at very low levels. For example, some patients may have to begin in the pool and then take a few months to graduate to simple and gentle land strengthening. It may take one to two years to work back up to anything resembling a big workout.

5. Fix the bad symmetry that likely got you here in the first place.

The next chapter of patient cases speaks to a practical application of SANS and how some patients at various levels of disability made it "out of the hole."



Chapter 6: Putting It All Together



"The physician can bury his mistakes, but the architect can only advise his client to plant vines—so they should go as far as possible from home to build their first buildings." Frank Lloyd Wright



My goal with this chapter is to present several patient examples relating to the spine, from less complex to more complex, so that the reader can see how the puzzle pieces fit together and interact with each other.

Case 1: SI Joint Shirley

Shirley is a 54-year-old woman who fell and had chronic pain in the back of her hip. She had seen multiple physical therapists and chiropractors, obtaining only temporary relief. Patients like Shirley who complain of hip pain are often told they have pain coming from the hip joint. An X-ray is usually taken showing some arthritis, which would be common for her age. Based on little else than the report of hip pain and the X-ray, the patient is frequently scheduled for a hip replacement without ever confirming that it's her hip joint that is causing the pain. Isn't hip pain always from the hip joint?



Common Pain Area for SI Joint Pain

To investigate the possible cause of hip pain, let's look at the pelvis. The picture to the right shows that the <u>hip joint</u> is connected to the pelvis and that the next joint up the chain is called the SI Joint (short for <u>sacroiliac joint</u>). You can see that these joints are close together. Pain from either joint can cause patients to complain of "hip pain." The hip joint proper tends to

cause more groin pain, and patients with SI joint problems tend to have more pain in the back of the hip near the PSIS area (see diagram to the left that shows a common location of SI joint pain). Truth is, either the hip or the SI joint can cause pain in the back or front of the hip, so how do we tell which is causing the complaint of hip pain?



To determine the source of the pain, we performed diagnostic numbing injections. Under X-ray guidance (fluoroscopy), we injected numbing medicine into the SI joint and hip. Injecting the hip only minimally helped her pain, while injecting her SI joint took away 70% of her pain. We had our man!





Since we had injected ultra-low-dose anti-inflammatories in her joint, this gave her some relief for a few weeks, but the pain returned. Now the question was, what caused her to have chronic SI joint pain? Because of her fall on the SI joint area, we suspected instability, which was confirmed on exam (see SANS pyramid to the right). What other clues led us to believe that she might be unstable in the SI joint? The patient had tenderness throughout the muscle attachments associated with the SI joint. The attachments of the <u>gluteal muscles</u> and <u>piriformis</u> showed signs of <u>enthesopathy</u>. At the back of the hip (greater trochanter), she was also

tender where these muscles attached. If she had just injured the joint, this didn't make sense. However, if the joint was unstable, these muscles would be working overtime to try and help stabilize the area, thus causing the enthesopathy. So in addition to treating the SI joint, these other areas would need to be treated as well. Why? While they might go away if we fixed the stability issue in the SI joint, based on clinical experience, it was more likely that they would remain as the damage at these muscle tendon attachments had been done.





So what options are there to fix an unstable SI joint? Fusing the joint surgically with screws has been used, but fusion usually just transfers forces to the joints above and below, so we wouldn't consider this option. Most pain management physicians would either consider repeatedly injecting high-dose steroids into the joint or an SI joint radiofrequency procedure.

As discussed in the "Articulation" chapter, high-dose steroids can damage the joint. Radiofrequency is where special needles or catheters are inserted to ablate the

nerves that take pain from the joint. This is covered by many insurers, <u>but these small</u> <u>nerves also provide proprioceptive input to the muscles that stabilize the joint, so nuking</u> <u>them could mean less active muscular stability</u>. In addition, this would only address the joint pain and not the pain she was having from the enthesopathy muscle areas. In our experience, the only treatment that would be regenerative rather than ablative (build up rather than destroy), tighten the ligaments to help the passive instability, and address the areas of muscle/tendon enthesopathy is prolotherapy.



After two prolotherapy treatments, pain was down by 75%. The next step was to address the likely muscular stability issues. The muscular stability system for the SI joint is the transversus abdominus. This is a tough muscle to teach patients to contract, as it's the deepest stomach muscle. However, we can easily see the muscle on ultrasound imaging, so we used this advanced imaging to help the patient have a sense of when she was contracting the muscle.

In summary, Shirley is a good case to understand how all the treatment choices and pieces fit together. Starting with identifying a pain generator then asking why she still hurts (instability) then looking at different joint treatment options and finally ending with a very specific rehabilitation component to address the muscular instability and weakness.

Fat & Skin		
		External Oblique
Fascial //		Internal Oblique
	Transversus Abdominis	
	Abdominal Contents	
Depth 8.62		

Case 2: Unstable Mable

Mable is a 45-year-old white female who was in a rear-end car crash a few months prior to our evaluation. Immediately after the crash, she noted severe and sharp right-sided upper neck and head pain. When she is seen in the clinic, she's tender over the right C1–C3 facet joints, and her right upper trapezius, levator scapula, sternocleidomastoids, and scalenes are tight. She has headaches with pressure over the upper neck. What happened to Mable and why does she still hurt after several months?

The upper neck facet joints are commonly injured in rear-end car crashes. In addition, the upper neck ligaments can be injured as well. Mable gets good relief once we inject lowdose anti-inflammatories into the right C1-C3 facet joints, but that's only temporary. Her 3.0 ultra-high-field MRI shows that there is evidence of likely stretching of the ligaments that hold the head on (alar and transverse ligaments).

I believe what happened to Mable is that once these structures were injured, the upper neck stability muscles went off-line. <u>There is good evidence in whiplash-injured patients</u> <u>that these muscles atrophy.</u> What happens next? The big neck muscles take over. These are the upper trapezius, levator scapula, sternocleidomastoids (SCMs), and the scalenes. However, they weren't designed to be neck stabilizers, so their attachments get overloaded, leading to enthesopathy. This is what causes pain at the back of Mable's head and irritates the occipital nerves that exit near the attachment of the upper trapezius and SCMs. This leads to chronic headache (as well as the referred pain from the injured C1-C3 joints, which are known causes of headache as well). In addition, Mable was unable to bear weight on these injured joints, so her neck curve was lost as her body figured out how to offload the joints.





We injected the upper cervical facet joints where the highfield MRI of her upper neck showed problems in the ligaments that hold the head in place. The enthesopathy was treated by prolotherapy injections where the muscles attached to the back of the head. Once the facet joints and upper neck muscles were calmed down, Mable was able to strengthen her neck. She also underwent curve-restoration therapy to get back the normal lordosis. Once the joint pain, enthesopathy, muscular stability, and alignment were addressed, Mable dramatically improved.

Case 3: Sensitized Sally

Sally is a 40-year-old woman who was in a rear-end car crash and developed neck and back pain within weeks of the injury. At that point, she developed severe pain and numbness in her hands and feet as well as headaches. She was diagnosed with fibromyalgia and given pain medications.

When she was first seen at our clinic, she had severe tenderness everywhere. She told me that attempts at physical therapy, massage, and trigger point injections had all caused days of severe pain. The patient was given a quantitative somatosensory test that demonstrated significant central sensitization. She was placed on Lyrica to help reduce her nerve sensitization. Once that began to reduce the severe pain, it became clear that multiple joints and muscles were being impacted by the nerve issues. Her posture had eroded, and the muscular stability in the low back was compromised.

Once the Lyrica began to help, we addressed the neuromuscular trigger points with IMS, the postural issues with Egoscue, and injected multiple joints with low-dose anti-inflammatories to reduce pain. The patient also had an epidural in the low back to reduce nerve pain and rehabilitation to get the low-back stabilizers back online. The patient's pain then began to reduce. However, due to the nature of the nerve injury, she will require long-term pain management.



Case 4: Catawampus Wayne

Are you ready for a more complex biomechanical analysis? This one demonstrates how a little injury can eventually lead to bigger problems. This patient had a serious fall from a bike about three years ago. He injured his shoulder, kidney, and hip.

When he was first evaluated for stem cell treatment of his hip, I was concerned about his low back. While stem cells in the hip helped the hip pain (he now walks faster through an airport), over the ensuing year, he continued to develop problems in his low back and leg. He was finally diagnosed with a cyst on his right L4–L5 facet joint that was pressing on a spinal nerve and giving him pain down the leg.



The facet joints are small joints in the back, and sometimes arthritis of the joints can result in a cyst (just like a swollen knee joint can develop a Baker's cyst). These cysts can press on spinal nerves, so they can be a double whammy for the patient.

The patient had the facet cyst treated with a steroid injection to pop the cyst. This helped some of the leg symptoms and severe nerve pain, but by the time I re-examined him, his back was pretty bad (unable to stand straight). This was impacting his work as a physician. While knowing he has a facet cyst is a good start in helping him, asking the question of how he got that way is important if this is going to be successfully treated without surgical fusion of this level. His case is a good example of the Orthopedics 2.0 concept. Consider the Orthopedics 2.0 pyramid to the right, in which I've filled in various portions.



To better explain, more discussion and pictures are needed. On his flexion-extension views, it was noted that he had the L4 vertebra slipping forward on the L5 vertebra. This forward slippage was at the same level as his facet cyst. Coincidence? Likely not. The way to understand this problem involves some ligaments in the back of the spine that act as the major duct tape that help keep the spine aligned. These ligaments are the supraspinous and interspinous ligaments. The image below and to the left shows the ligaments (red lines) in the back of the spine.

So before his bike accident, these ligaments were doing their job, helping to hold the spine in alignment. The picture above shows that the ligaments are holding the vertebrae in place. They act to make sure that all of the vertebrae stay aligned when you bend forward. They also control how much one vertebra is allowed to move relative to the vertebra above or below. Note that after the bike crash, the injury and tearing of these ligaments (gap in the red lines—see right) allow the L4 vertebra to move forward on the L5 vertebra. Since these ligaments help to hold the vertebrae in alignment when he bends forward, the facet joints move more than they can tolerate. This ultimately leads to excess wear and tear of these joints. Is there another part of this puzzle?







An interesting observation as he lies prone on the table is seen to the right. Why is there severe bulging of the left abdominal wall? Further questioning of the patient reveals that he also injured his kidney in the bike accident and had surgery on the left kidney. The scar can be seen in the picture above. An important stabilizer of the back is the transversus abdominus. This muscle was likely cut through to get at the kidney, resulting in the muscle weakness you see above on the left side (inability to hold in the abdominal contents).



Putting all of this together into the Ortho 2.0 triangle results in the following analysis. The transversus abdominus is a muscle that's the deepest of the abdominal wall. It attaches to the thoracodorsal fascia, and pulling on this muscle on both sides helps to allow the buoyancy of the abdominal contents to assist in offloading the weight of the upper body by literally floating it on the abdominal contents. It's also a major low-back stabilizer all by itself. The picture to the left shows that it attaches to fascia that then attaches to the back of the vertebra on both sides (spinous process). This axial view ("saw" you in half view) shows that if the pull is equal on both sides, this helps to keep the vertebra straight. However, if we cut one side of the transversus abdominus muscle (for example to get to a damaged kidney), the forces on the vertebra will be unequal, causing it to have a slight tendency to rotate (in this case to the left). This forces on the right facet joint will increase, causing more wear and tear forces on that side.



Below is an actual axial MRI image which shows the abnormal pull to the right by the transversus abdominus muscles (orange arrows) causing extra force on the right facet joint (red star). This is where the facet cyst is located.



So in summary, we believe that the damage to the ligaments in the back (supraspinous and interspinous ligaments) as well as this abnormal pull of one transversus abdominus over the other have caused the facet joints to wear out. Their response on the right (the side where we would predict the most force) is to swell to try and keep up with the wear and tear. This led to a facet cyst and then ultimately pressure on the spinal nerve. This complex example illustrates the importance of piecing together all of the parts and pieces of what caused the musculoskeletal system to fail. Many times, treating patients with musculoskeletal problems is as simple as a quick fix (in this case popping a facet cyst with a facet injection), other times it takes considerably more analysis.



About the Author



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Dr. Centeno is one of the few physicians in the world with extensive experience in the culture expansion of and clinical use of adult stem cells to treat orthopedic injuries. He is a founding member of the International Cellular Medicine Society. His clinic incorporates a variety

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Dr. Centeno has chaired multiple international research-based conferences. He also maintains an active research-based practice, with multiple publications listed in the US National Library of Medicine. Dr. Centeno has also served as editor in chief of a medical research journal dedicated to traumatic injury. He has lectured all over the world on regenerative therapies, including at the Vatican in Rome.

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