A short History of the development of Ultrasound in Obstetrics and Gynecology

Dr. Joseph Woo

[Part 1] Tread this first

L he story of the development of ultrasound applications in medicine should probably start with the history of measuring distance under water using sound waves. The term **SONAR** refers to **Sound Navigation and Ranging**. Ultrasound scanners can be regarded as a form of 'medical' Sonar.

As early as **1826**, <u>Jean-Daniel Colladon</u>, a Swiss physicist, had successfully used an <u>underwater bell</u> to determine the speed of sound in the waters of Lake Geneva. In the later part of the 1800s, physicists were working towards defining the

fundamental physics of sound vibrations (waves), transmission, propagation and refraction. One of them was Lord Rayleigh in England whose famous treatise "the Theory of Sound" published in 1877 first described sound wave as a mathematical equation, forming the basis of future practical work in acoustics. As for high frequency 'ultrasound', Lazzaro Spallanzani, an Italian biologist, could be credited for it's discovery when he demonstrated in 1794 the ability of bats navigating accurately in the dark was through echo reflection from high frequency inaudible sound. Very high frequency sound waves above the limit of human hearing were generated by English scientist Francis Galton in 1876, through his invention, the Galton whistle.

The real breakthrough in the evolution of high frequency echo-sounding techniques came when the <u>piezo-electric effect</u> in certain crystals was discovered by <u>Pierre Curie</u> and his brother Jacques Curie in Paris, France in **1880**. They observed that an electric potential would be produced when mechanical pressure was exerted on a **quartz crystal** such as the Rochelle salt (sodium potassium tartrate tetrahydrate). The reciprocal behavior of achieving a mechanical stress in response to a voltage difference was mathematically deduced from thermodynamic principles by physicist **Gabriel Lippman** in 1881. and which was quickly

Pierre Curie

1859 - 1906

thermodynamic principles by physicist <u>Gabriel Lippman</u> in 1881, and which was quickly verified by the Curie brothers. It was then possible for the **generation** and **reception** of **'ultrasound'** that are in the frequency range of millions of cycles per second (megahertz) which could be employed in echo sounding devices. Further **research and development** in piezo-electricity soon followed.



underwater echo-sounding range display systems.

Underwater sonar detection systems were developed for the purpose of underwater navigation by submarines in World war I and in particular after the **Titanic** sank in **1912**. <u>Alexander Belm</u> in Vienna, described an underwater echo-sounding device in the same year. The first patent for an underwater echo ranging sonar was filed at the British Patent Office by English metereologist Lewis Richardson, one month after the sinking of the Titanic. The first working sonar system was designed and built in the United States by Canadian <u>Reginald Fessenden</u> in **1914**. The Fessenden sonar was an

electromagnetic moving-coil oscillator that emitted a low-frequency noise and then switched to a receiver to listen for echoes. It was able to detect an iceberg underwater from 2 miles away, although with the low frequency, it could not precisely resolve its direction.

The turn of the century also saw the invention of the **Diode** and the **Triode**, allowing powerful electronic amplifications necessary for developments in ultrasonic instruments. Powerful high frequency **ultrasonic** echo-sounding device was developed by emminent French physicist <u>Paul</u> <u>Langévin</u> and Russian scientist **Constantin Chilowsky**, then residing in France. <u>Patents</u> were filed in France and the United States. They called their device the '<u>hydrophone</u>'. The **transducer** of the hydrophone consisted of a mosaic of thin quartz crystals glued between two steel plates with a resonant frequency of 150 KHz. Between 1915 and 1918 the hydrophone was further improved in classified research activities and was deployed extensively in the <u>surveillance</u> of German U-boats and submarines. The first known sinking of a submarine detected by hydrophone occurred in the Atlantic during World War I in April,1916.

Langevin's hydrophones had formed the basis of the development of naval pulse-echo sonar in

the following years. By the mid 1930s, many ocean liners were equipped with some form of



Paul Langevin 1872 - 1946

A RADAR HISTORY OF WORLD WAR II Trewels and Multaw Investments Land Boom In another development, the first successful **radio** range-finding experiment occurred in **1924**, when British physicist **Edward Appleton** used radio echoes to determine the height of the ionosphere. The first practical <u>RADAR</u> system (Radio Detection and Ranging, and using electromagnetic waves rather than ultrasonics) was produced in **1935** by another British physicist <u>Robert Watson-Watt</u>, and by **1939** England had established a chain of radar stations along its south and east coasts to detect aggressors in the air or on the sea. **World war II** saw <u>rapid developments</u> and refinements in the naval and military radar by researchers in the United States.



Such radar display systems had been the direct precursors of subsequent 2-dimensional sonars and **medical ultrasonic systems** that appeared in the late 1940s. Books such as the **"Principles of Radar"** published by the **Massachusetts Institute of Technology** (M I T) Radar school staff in **1944** detailed the techniques of oscilloscopic data presentation which were employed in medical ultrasonic research later on (see below). Two other engineering advances probably had also influenced significantly the development of the sonar, in terms of

the much needed data aquistion capabilities: the **first digital computer** (the Électronic Numerical Integrator and Computer -- the <u>ENIAC</u>) constructed at the University of Pennsylvania in **1945**, and the invention of the point-contact <u>transister</u> in **1947** at AT & T's Bell Laboratories.

Yet another **parallel and equally important** development in ultrasonics which had started in the **1930's** was the construction of pulse-echo <u>ultrasonic metal flaw detectors</u>, particularly relevant at that time was the check on the integrity of metal hulls of large ships and the armour plates of battle tanks.

The concept of ultrasonic metal flaw detection was <u>first suggested</u> by Soviet scientist <u>Sergei Y Sokolov</u> in **1928** at the Electrotechnical Institute of Leningrad. He showed that a transmission technique could be used to detect metal flaws by the variations in ultrasionic energy transmitted across the metal. The resolution was however poor. He suggested subsequently at a later date that a **reflection method** may be practical.

The equipment suggested by **Sokolov** which could generate very short pulses necessary to measure the brief propagation time of their returning echoes was not available until the 1940s. <u>Early pioneers</u> of such reflective **metal flaw detecting devices** were **Floyd A Firestone** at the University of Michigan, and **Donald Sproule** in England. Firestone produced his patented "<u>supersonic reflectoscope</u>" in **1941** (US-Patent 2 280 226 "Flaw Detecting Device and Measuring Instrument", April 21, 1942). Because of the war, the reflectoscope was not formally published until **1945**. **Messrs. Kelvin and Hughes**® in England, where Sproule



Metal flaw detector in use ** (Kretztechnik, Austria)

was working, had also produced one of the earliest pulse-echo metal flaw detectors, the M1. Josef and Herbert Krautkrämer produced their first <u>German version</u> in Köln in 1949 followed by equipment from <u>Karl Deutsch</u> in Wuppertal. These were followed by other versions from Siemens® in Erlangen, <u>KretzTechnik</u> AG in Austria, Ultrasonique in France and <u>Mitsubishi</u> in Japan. In 1949, <u>Benson Carlin</u> at M I T, and later at Sperry Products, published "Ultrasonics", the first book on the subject in the English language.







The underwater **SONAR**, the **RADAR** and the ultrasonic **Metal Flaw Detector** were each, in their unique ways, a precursor of medical ultrasonic equipments. The modern ultrasound scanner embraces the concepts and science of all these modalities.

The early development of ultrasonics is summarised <u>here</u>.

Readers are also referred to <u>an article</u> by <u>Dr William O'Brien Jr.</u>, which also looks at the early history of the developments of ultrasonics.[^]

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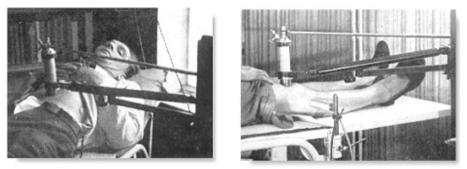
L he use of **Ultrasonics** in the field of medicine had nonetheless started initially with it's applications in <u>therapy</u> rather than diagnosis, utilising it's heating and disruptive effects on animal tissues. The destructive ability of high intensity ultrasound had been recognised in the **1920s** from the time of <u>Langévin</u> when he noted destruction of school of fishes in the sea and pain induced in the hand when placed in a water tank insonated with high intensity ultrasound; and from the seminal work in the **1930s** from **Robert Wood**, **Newton Harvey** and <u>Alfred Loomis</u> in New York and **R Pohlman** in Erlangen, Germany.

High intensity ultrasound progressively evolved to become a neuro-surgical tool. <u>William Fry</u> at the University of Illinois and **Russell Meyers** at the University of Iowa performed **craniotomies** and used ultrasound to destroy parts of the basal ganglia in patients with **Parkinsonism**. **Peter Lindstrom** in San Francisco reported ablation of frontal lobe tissue in moribound patients to alleviate their pain from carcinomatosis. **Fry** in particular had worked towards improving research and dosimetry standards, which was much needed at the time.



Ultrasound used in therapy

Ultrasonic energy was also extensively used in physical and rehabilitation medicine. Jerome Gersten at the University of Colorado reported in 1953 the use of ultrasound in the treatment of patients with rheumatic arthritis. Other reseachers such as <u>Peter Wells</u> in Bristol, England, Douglas Gordon in London and Mischele Arslan in Padua, Italy employed ultrasonic energy in the treatment of Meniere's disease.



Uses of ultrasonic energy in the 1940s. Left, in gastric ulcers. Right, in arthritis

The **1940**s saw exuberant claims made in some sectors on the effectiveness of ultrasound as an almost "cure-all" remedy, abeit the lack of much scientific evidence. This included conditons such as arthritic pains, gastric ulcers, eczema, asthma, thyrotoxicosis, haemorrhoids, urinary incontinence, elephanthiasis and even angina pectoris! Cynicism and concern over harmful tissue damaging effects of ultrasound were also mounting, which had curtailing consequences on the development of diagnostic ultrasound in the years that followed.



Karl Theodore Dussik

It was around similar times that ultrasound was used experimentally as a possible **diagnostic tool** in medicine. **H Gohr** and **Th. Wedekind** at the Medical University of Koln in Germany in **1940** presented in their paper "<u>Der Ultraschall in der Medizin</u>" the possibility of ultrasonic diagnosis basing on echo-reflection methods similar to that used in metal flaw detection. They suggested that the method would be able to detect tumours, exudates or abscesses. However they were unable to publish convincing results from their experiments. <u>Karl Theo Dussik</u>, a neurologist/ psychiatrist at the University of Vienna, **Austria**, who had begun experiments in the late 1930s, was generallly regarded as the **first** physician to have employed ultrasound in medical diagnosis.

Dussik, together with his brother Friederich, a physicist, attempted to locate brain tumors and the cerebral ventricles by measuring the transmission of ultrasound beam through the skull. Dussik presented his initial experiments in a <u>paper in 1942</u> and further results after the end of the second world war in **1947**. They called their procedure "<u>hyperphonography</u>".

transmission technique with two transducers placed on either side of the head, and producing what they called

They used a through-

"ventriculograms", or echo images of the ventricles of the brain. Pulses of 1/10th scond were produced at 1.2 MHz. Coupling was obtained by immersing the upper part of the patient's head and both transducers in a water bath and the variations in the amount of ultrasonic power passing between the transducers was recorded photographically on heat-sensitive paper as light spots (not on a cathode-ray screen). It was an earliest attempt at the concept of 'scanning' a human organ. Although their apparatus appeared elaborate with the transducers mounted on poles and railings, the images produced were very rudimentary 2-dimensional rows of mosaic light intensity points. They had also reasoned that if imaging the ventricles was possible, then the technique was also feasible for detecting brain tumors and low-intensity ultrasonic waves could be used to visualize the interior of the human body.



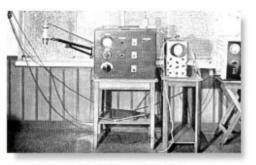
Dussik and his apparatus in 1946

Nevertheless, the images that Dussik produced were later thought to be <u>artifactual</u> by **W Güttner** and others at the Siemens Laboratory, Erlangen, Germany in 1952 and <u>researchers at the M.I.T.</u> (see below), as it had become apparent from further experiments that the reflections within the skull and attenuation patterns produced by the skull were contributing to the attenuation pattern which Dussik had originally thought represented changes in acoustic transmissions through the cerebral ventricles in the brain. Research basing on a similar transmission technique was not further pursued, both by Dussik, or at the M. I. T.. For more information read <u>Dussik</u>.

In nearby **Germany**, <u>Heinrich Netheler</u>, a physician at the Luebeck-South Hospital in **Hamburg**, was operating in **1945** a small repair facility for medical equipments at the Hamburg university hospital at Eppendorf and had a mission of developing inventive medical products. **Professor Hansen**, his superior, suggested to him in that year to develop an ultrasonic tomographic equipment for medical use basing on the concept of the RADAR. Important pioneering <u>reseach</u> <u>work</u> started at the Eppendorf University Hospital. Nevertheless, due to a lack of funds right after the war, the equipment designs had not reached the stage of actual fabrication. In the mid 1940s, German physician <u>Wolf-Dieter Keidel</u> at the Physikalisch-Medizinischen Laboratorium at the **University of Erlangen**, Germany, also studied the possibility of using ultrasound as a medical diagnostic tool, mainly on cardiac and thoracic measurements. Having discussed with researchers at **Siemens**, he conducted his experiments using the transmission technique with ultrasound at 60 KHz, and rejected the pulse-reflection method. He was only able to make satisfactory recordings of intensity variations in relation to cardiac pulsations. He envisaged much more difficulties would be encounterd with the reflection method. In the **First Congress of**

Ultrasound in Medicine held in Erlangen, Germany in May, 1948, Dussik and Keidel presented their papers on ultrasound employed in medical diagnosis. These were the only two papers that discussed ultrasound as a diagnostic tool. The other papers were all on its therapeutic use.

In France, French scientists who were in the study of ultrasonics, namely André Dognon and André Dénier and several others at the research center in Salpêtrière in Paris also embarked on ultrasound insonation experiments before the 1950s. Dénier published his theoretically work on ultrasound transmission in **1946**, among many other works on ultrasound used in therapy, and suggested the possibility of "<u>Ultrasonoscopie</u>". This was a transmission technique and recordings made on a micro-ampere meter and oscilloscope. Equipments were fabricated from 'therapy' counterparts and various electrical current values were determined on different body tissues. Attempts to display voltages as Lissajous figures on the oscilloscope were made. However the work was unsuccessful in producing useful structural images and related instruments were not constructed. André Dénier published in 1951 his book, "Les Ultras-sons -- Appliques a la Medecin". Nearly the entire book was devoted to ultrasonics used in the treatment of



Denier's ultrasonic apparatus in 1946

various diseases and only a small portion of the text was on ultrasound diagnostics.

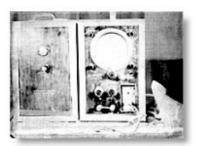


George D. Ludwig

Systematic investigations into using ultrasound as a diagnostic tool finally took off in the United States in the late 1940s. The time was apparently ripe for this to happen. The concept of applying ultrasonics to medicine had progressively matured, so were the available equipments and electronics after the war. George Ludwig, a graduate from the University of Pennsylvania in 1946 was on active duty as junior Lieutenant at the **Naval Medical Research Institute** in Bethseda, Maryland. There, he <u>began experiments on</u> <u>animal tissues</u> using A-mode industrial <u>flaw-detector equipment</u>. Ludwig designed experiments to detect the presence and position of foreign bodies in animal tissues and in pertingents to be the presence of the presen particular to localise gallstones, using reflective pulse-echo ultrasound methodology similar to that of the radar and sonar in the detection of foreign boats and flying objects. A substantial portion of Ludwig's work was considered classified information by the Navy and was not published in medical journals. Although Ludwig's work had started at a considerably earlier date, notice of his work was not released to the public domain until October 1949 by the United States Department of Defence. The June '49 report is considered the first report of its kind on the diagnostic use of ultrasound from the United States

Ludwig systematically explored physical characteristics of ultrasound in various tissues, including beef and organs from dogs and hogs. To address the issue of detecting gallstones in the human body, he studied the acoustic impedance of various types of gallstones and of other tissues such as muscle and fat in the human body, me studied the accusate impedance of methodologies and frequencies. His collaborators included **Francis Struthers** and **Horace Trent**, physicists at the Naval Research Laboratory, and Ivan Greenwood, engineer from the General Precision Laboratories, New York, and the Department of Research Surgery, University of Pennsylvania. Ludwig also investigated the detection of gallstones (outside of the human body) using ultrasound, the stones being first embedded in pieces of animal muscle. Very short pulses of ultrasound at a repetition rate of 60 times per second were employed using a combined transmitter/ receiver transducer. Echo signals from the reflected soundwaves were recorded on the oscilloscope screen. Ludwig was able to detect distinct ultrasonic signals corresponding to the gallstones. He reported that echo patterns could sometimes be confusing, and multiple reflections from soft tissues could make test results difficult to interpret. Ludwig also studied transmission through living human extremities, to measure acoustic impedance in muscle. These investigations also explored issues of attenuation of ultrasound energy in tissues, impedance mismatch between various tissues and related reflection coefficients, and the optimal sound wave frequency for a diagnostic instrument to achieve adequate penetration of tissues and resolution, without incurring tissue damage. These studies had helped to build the scientific foundation for the clinical use of ultrasound.

In the following year, Greenwood and General Precision Laboratories made available commercially the "<u>Ultrasonic Locator</u>" which Ludwig used for "use in Medicine and Biology". Suggested usage indicated in the <u>sales information</u> <u>leaflet</u> already included detection of heart motion, blood vessels, kidney stones and glass particles in the body. Ludwig's pulse-reflection methodology and equipment in his later experiments on sound transmission in animal tissues were after earlier designs from the work of John Pellam and John Galt in 1946 at the Electronics and Acoustics research laboratories of the Massachusetts Institute of Technology (M. I. T.), which was on the measurement of ultrasonic transmission through liquids. The M. I. T. was then very much at the forefront of electronics and ultrasonics research. A significant amount of physical data and instrumentation electronics were already in place in the second half of the 1940s, on the characteristics of ultrasound propagation in solids and liquids.



Ludwig's A-mode apparatus in his gallstone experiments

Among other important original findings, Ludwig reported the <u>velocity of sound</u> <u>transmission</u> in **animal soft tissues** was determined to be between 1490 and 1610 meters per second, with a mean value of <u>1540 m/sec</u>. This is a value that is still in use today. He also determined that the optimal scanning frequency of the ultrasound transducer was between 1 and 2.5 MHz. His team also showed that the speed of ultrasound and acoustic impedance values of high water-content tissues do not differ greatly from those of water, and that measurements from different directions did not contribute greatly to these parameters.

Ludwig went on to collaborate with the Bioacoustics laboratory at the M. I. T.. His work with physicist Richard Bolt (who, at the age of 34 was appointed Director of a newly conceived Acoustics Laboratory at M. I. T.), neurosurgeon H Thomas Ballantine Jr. and research physicist Theodor Hueter from Siemens, Germany were considered very important seminal work on ultrasound propagation characteristics in mammalian tissues.



Theodore Hueter

Prior to 1949, Hueter had already been involved at Siemens. Erlangen, Germany, in ultrasonic propagation experiments in animal tissues using ultrasound at frequencies of about 1 MHz, and in ultrasonic dosimetry measurements. These were started in the early 1940s by Ultrasonics pioneer Reimar Pohlman in the same laboratory. In 1948, Hueter met Bolt and Ballantine at an ultrasonic trade show in New York and agreed to join them for new research into the application of ultrasonics in human diagnosis. After a visit to <u>Dussik's</u> department in Austria with Bolt and Ballantine, the group launched a <u>formal project</u> at M. I. T. to perform experiments in through transmission similar to that of Dussik's. Their initial experiments produced results similar to that of Dussik's, and their conclusions were published in their papers in 1950 and 1951 in the Journal of the Acoustical Society of America, and Science. In further



M I T's project reported in "The Tech", 1950

experiments the team put a skull in a water bath and showed that the ultrasonic patterns they had been obtaining from the heads of selected subjects could also be obtained from an empty skull. They noted that ultrasonic mapping of the brain tissues within the human skull was prone to great error due to the large bone mass encountered. Efforts were made to compensate for the bone effects by using different frequencies and circuitries, but were only marginally successful at that stage of computational technology.

The M. I. T. research project was subsequently terminated in 1954. They wrote in their paper: "It is concluded that though compensated ultrasonograms (sound shadow pictures) may contain some information on brain structure, their are too sharply "noise" limited to be of unqualified clinical value". The findings had prompted the United States Atomic Energy Commission to conclude that ultrasound will not be useful in the diagnosis of brain pathologies. Medical research in this area was somewhat curtailed for the several years that followed, and enthusiasm was dampened at the Siemens laboratories in Germany to carry out further developments in imaging with ultrasound. At M .I. T. nevertheless, in the course of these pursuits, much basic data essential for tissue characterization and dosimetry were assembled and proved useful for later diagnostic work on other body regions. They had also demonstrated very importantly that interpretable **2-dimensional images** was not impossible to obtain. These efforts had paved the way for the subsequent development of 2-D ultrasonic image formation. M. I. T.'s research had also benefited from interactions between the various groups at **Champaign-Urbana**, **Minnesota** and **Denver**.

By the mid 1950s, bibliographic listing of work on ultrasonic physics and engineering applications had totalled more than 6,000. Ultrasonics was already extensively deployed in non-destructive testing, spot welding, drilling, gas analysis, aerosol agglomeration, shear processing, clothes washing, laundering, degreasing, sterilization and, to a lesser extent, medical therapy. Hueter and Bolt's book "SONICS - techniques for the use of sound and ultrasound in engineering and science" published in 1954 became, for example, one of the important treatises in ultrasonic engineering.



In 1956, D Goldman and Hueter pulled together all the then available data on ultrasonic propagation in mammalian tissues for publication in the Journal of the Acoustical Society of America. The earliest journal devoted entirely to the application of ultrasonics in medicine was "Der Ultraschall in der Medizin" published in Germany. Articles prior to 1952 were entirely on aspects of ultrasound used in therapy. Much of the academic activity at M. I. T. were published in the *M. I. T. quarterly progress reports* and the *Journal of the Acoustic Society of America*. After the mid-1950s, due to its ineffectiveness, the transmission technique in ultrasonic diagnosis was abandoned from medical ultrasound research worldwide except for some centers in Japan, being replaced by the reflection technique which had received much attention in a number of pioneering centers throughout Europe, Japan and the United States.

Smaller and better transducers were being assembled from the newer piezoceramics barium titanate after the mid 1940s. They were replaced by lead zirconate-titanate (PZT) when it was discovered in 1954. PZT had a high electro-mechanical coupling factor and more superior frequency-temperature characteristics. The newer transducers had better overall

sensitivity, frequency handling, coupling efficiency and output. The availability of very high input impedance amplifiers built from improved quality **electrometer tubes** in the early 1950s had also enabled engineers to greatly amplify their signals to improve sensitivity and stability.

The 'newer' uni-directional pulse-echo A-mode devices developed from the reflectoscope/ metal flaw detectors were soon employed in experiments on medical diagnosis by bold and visionary pioneers around the world. Such were the cases with Douglas Gordon, JC Turner and Val Mayneord in London, Lars Leksell (in 1950), Stigg Jeppson and Brita Lithander in Sweden, Marinus de Vlieger in Rotterdam and Kenji Tanaka and Toshio Wagai in Japan for their pioneering work in the examination of brain lesions. These devices were also employed by Inge Edler and Carl Hellmuth Hertz in Lund in cardiac investigations in 1953, and followed on by Sven Effert in Germany in 1956, Claude Joyner and John Reid at the University of Pennsylvania in 1957 and Chih-Chang Hsu in China, designing their own A- and later on M-Pennsylvania in 1957 and Chin-Chang Hsu in China, designing their own A- and later on m-mode equipment. Similarily A-mode devices were used in ophthalmologic investigations by Henry Mundt Jr and William Hughes at the University of Illinois in 1956, Arvo Oksala in Finland in 1957 and Gilbert Baum and Ivan Greeenwood in 1955. These uses were all in the



Inge Edler

1950s and largely predated clinical applications in the abdomen and pelvis. Researchers in Japan were also actively investigating and producing similar ultrasonic devices and their diagnostic use in neurology, but their findings have only been sparsely documented in the English literature (see below).



John Julian Wild, an English surgeon and graduate of the Cambridge University in England, immigrated to the United States after World War II ended in 1945. He took up a position at the Medico Technological Research Institute of Minnesota and



John Wild c. 1953

started his investigations with ultrasound waves on the thickness of the bowel wall in various surgical conditions, such as paralytic ileus and obstruction. Working with **Donald Neal**, an engineer, Wild published their work in **1950** on uni-directional A-mode ultrasound investigations into the thickness of surgical intestinal material and later on the properties of gastric malignancies. They noted that malignant tissue was more echogenic than benign tissue and the former could be diagnosed from their density and failure to contract and relax. Wild's original vision of the application of ultrasound in medical diagnosis was more of a method of tissue diagnosis from the intensity and characteristics of different returning echos rather than as an imaging technique. Between 1950 and 51, he also collaborated with <u>Lyle</u> <u>French</u> at the department of Neurosugery in making diagnosis of brain tumors using ultrasound, although they had not found the method to be very helpful.

Donald Neal was soon deployed to regular naval services at the naval air base after the Korean war. John Reid, a newly graduating electrical engineer, was engaged through a grant from the

National Cancer Institute as the sole engineer to build and operate Wild's ultrasonic apparatus. The device which they first used was an ultrasonic instrument which had been designed by the U.S. Navy for training pilots in the use of the <u>radar</u>, with which it was possible to practise 'flying' over a tank of water covering a small scale map of enemy territory. "We have a **tissue radar machine** scaled to inches instead of miles by the use of ultrasound". **Wild** and **Reid** soon built a linear hand-held B-mode instrument, a formidable technical task in those days, and were able to visualise tumours by sweeping from side to side through breast lumps. The instrument operated at a frequency of 15 megahertz. In 1952 they published the Landmark paper: "<u>Application of</u> <u>Echo-Ranging Techniques to the Determination of Structure of Biological Tissues</u>". In another paper Reid wrote about their first scanning equipment:



John M Reid c.1970s

' The first scanning machine was put together, mechanically largely by John with parts obtained through a variety of friends in Minneapolis. I was able to modify a standard test oscilloscope plug-in board. We were able to make our system work, make the first scanning records in the clinic, and mail a <u>paper</u> off to Science Magazine within the lapsed time of perhaps ten days. This contribution was accepted in early 1952 and became the <u>first publication</u> (to my knowledge) on intensity-modulated **cross-section** ultrasound imaging. It appeared even before Douglass Howry's paper from his considerably more elaborate system at the end of the same year.



Wild and Reid's setup reported in Electronics in 1955

In May 1953 they produced real-time images at 15 megahertz of cancerous growths of the breast. They had also coined their method 'echography' and 'echometry', suggesting the quantitative nature of the investigation. By 1956, Wild and Reid had examined 117 cases of breast pathology with their linear real-time Bmode instrument and had started work on colon tumour diagnosis and detection. Analysis of the breast series showed promising results for pre-operative diagnosis. Malignant infiltration of tissues surrounding breast tumours could also be resolved.

Wild and Reid had also invented and described the use of A-mode trans-vaginal and trans-rectal scanning transducers in 1955. Despite these, Wild was not commended for his unconventional research methods at the time. His results were considered difficult to interpret and lacked overall stability. Intellectual and financial support for Wild's research dwindled, and legal disputes and politics also hampered further governmental grants. His work was eventually supported only by private funds which ran scarce and his data apparently received much less recognition than they deserved.

John Reid completed his MS thesis in 1957 on focusing radiators. In addition he had importantly verified that dynamic focusing was practical. After leaving Wild's laboratory he pursued his doctoral degree at the University of Pennsylvania. From 1957-1965 he worked on echocardiography, producing and using the first such system in the United States, with cardiologist **Claude Joyner**.

Visit John Wild's own site on his discoveries and current activities.

Read also: "The scientific discovery of sonic reflection of soft tissue and application of ultrasound to diagnostic medicine and tumor screening" by John J Wild (Press Release at the Third Meeting of the World Federation for Ultrasound In Medicine and Biology, 1982).



COUND.WAVE PORTRAIT IN THE FIESH

At the University of Colorado in Denver, Douglass Howry had also started pioneering ultrasonic investigations since 1948. Howry, a radiologist working at the Veteran's Administration Hospital, had concentrated more of his work on the development of B-mode equipment, displaying body structures in a 2-dimensional and sectional manner "comparable to the actual gross sectioning of structures in the pathology laboratory". Published works from the **M I T** Radar school staff served as initial reference metazia on tophicipuse in data reconstration material on techniques in data presentation.

He was able to demonstrate an ultrasonic echo Douglass Howry c 1967 interface between structures or tissues, such as that between fat and muscle, so that the individual structures could be outlined. Supported by his nephrologist friend and colleague Joseph Homles, who was

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Howry's somascope reported in the LIFE magazine in 1954

then the acting director of the hospital's Medical Research Laboratories, Howry produced in 1951 with William Roderic Bliss and Gerald J Posakony, both engineers, the 'Immersion tank ultrasound system' *, the first 2-dimensional Bmode (or PPI, plan position indication mode) linear compound scanner. Two dimensional cross-sectional images were published in 1952 and 1953, which convincingly demonstrated that interpretable 2-D images of internal organ

structures and pathologies could be obtained with ultrasound. The team produed the formal motorized 'Somascope', a compound circumferential scanner, in **1954**. The transducer of the somascope was mounted around the rim of a large metal immersion tank filled with water . The machine was able to make compound scans of an intra-abdominal organ from different angles to produce a more readable picture. The sonographic images were referred to as '<u>somagrams</u>'. The discovery and apparatus were reported in the Medicine section of the LIFE Magazine® in 1954.



Joseph H Holmes 1902 - 1982

The 'Pan-scanner' *, where the transducer rotated in a semicircular arc around the patient, was developed in **1957**. The patient sat on a modified dental chair strapped against a plastic window of a semicircular pan filled with saline solution, while the transducer rotated through the solution in a semicircular arc. The achievement was commended by the American Medical Association in 1958 at its scientific meeting in San Francisco, and the team's exhibit was awarded a Certificate of Merit by the association.

The work of **Douglass Howry**, **Joseph Holmes** and his team is necessarily the most important pioneering work in B-mode ultrasound imaging and contact scanning in the United States that had been the direct precursor of the kind of ultrasound imaging we have today. Pioneering designs in electronic circuitries were also

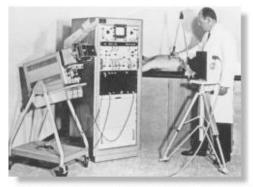


The pan-scanner in 1957

made in conjunction with the development of the B-scan, these included the pulse-echo generator circuitry, the limiter and log amplification circuitry and the demodulator and time gain compensation circuitries.

The Howry/ Holmes systems, although capable of producing 2-D, accurate, reproducible images of the body organs, required the patient to be totally or partially immersed in water, and remained motionless for a length of time. Migration to lighter and more mobile versions of these systems, particularly with smaller water-bag devices or transducers directly in contact and movable on the body surface of patients were imminently necessary.

Read notes and see more pictures from Gerald Posakony on the early Howry scanners here.



The articulated arm scanner that Wright and Meyerdirk built in 1962, the earliest of such design in the U.S.

Homles, together with consultant engineers William Wright and Ralph (Edward) Meyerdirk, and support from the U.S. Public Health Services and the University of Colorado, continued to fabricate a new prototype <u>compound contact</u> <u>scanner</u>, which had the transducer in direct contact with the patient's body and suspended on moving railings above the patient. The apparatus and the usuage of ultrasound scanning were reported in the May 22 issue of the TIME Magazine in 1964.



Time Magazine article May, 1964

After working on the project for about 2 years, the team finally came up with an innovative multi-joint articulated-arm compound

contact scanner with wire mechanisms and electronic position transducing potentiometers. The transducer could be positioned by hand and moved over the scanning area in various directions by the operator. In 1962, with blessing from Holmes, Wright and Meyerdirk left the University to form the Physionics Engineering® Inc. at Longmont, Colorado, to produce and market their scanner.

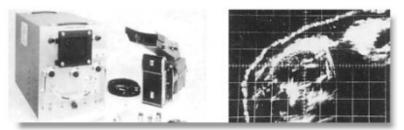
In 1963, the first hand-held articulated arm compound contact B-mode scanner (pictured on the left) was commercially launched in the United States. The launch was reported in the Longmont "*Daily Times-Call*" in 1963. This was the start of the most popular design in the history of static ultrasound scanners, that of the articulated-arm scanning mechanism.

Physionics® was acquired by the Picker Corporation in 1967. Picker continued to produced improved versions of the design right into the 1980s.

Much of the later work in clinical ultrasound was followed up by Homles and his colleagues, Stewart Taylor, Horace Thompson and Kenneth Gottesfeld in Denver. The group published some of the earliest papers in obstetrical and gynecological ultrasound from North America. Douglass Howry had moved to Boston in 1962 where he worked at the Massachussetts General Hospital until he passed away in 1969.



The Daily Times-Call Report, 1963



Earliest Wright-Meyerdirk scanner console with one of the first images from a practical commercial articulated-arm scanner. Portability was also emphasized.

In Japan, at about the same time as Wild and Howry's development, **Kenji Tanaka** and <u>Toshio Wagai</u>, surgeons at the Juntendo University, Tokyo, together with <u>Shigeru</u> <u>Nakajima</u>, director of the Japan Radio Company, **Rokuro Uchida**, physicist and chief engineer, had also started looking into the use of ultrasound in the diagnosis of intracranial disease in collaboration with the Nihon Musen Radiation and Medical Electronics Laboratory which had later become the <u>ALOKA</u>® Company in <u>1950</u>, headed by **Uchida**. Nakajima and Uchida built Japan's <u>first ultrasonic scanner</u> operating in the Amode in <u>1949</u>, modified from a metal-flaw detector. <u>Yoshimitsu Kikuchi</u>, professor in ultrasonics from the **Tohoku University** in <u>Sendai</u> also assisted in their research. Together, the team started their formal ultrasound research in **1952**.



B-mode experiments at Juntendo, 1955

They published 5 papers on <u>ultrasonic</u> <u>diagnosis in brain diseases</u> in that year and many other papers in the ensuing few years. In **1954**, Tanaka published an



Toshio Wagai c.1979

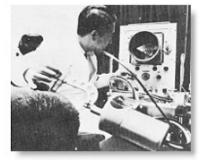
important review entitled "Application of ultrasound to diagnostic field", and investigations had started with other body organs. By **1955**, experiments and fabrication with **B-mode scanning** had started using a similar scope modified from the original A-mode machine coupled with a <u>linear moving transducer gantry</u>. This was shortly developed into the <u>water-bag</u> scanners.

▶ Also read the **Preface and introduction (history)** to Tanaka's book "Diagnosis of Brain Disease by Ultrasound" published in 1969 for a short history of his pioneering work in the 1950s.

The **M I T** hosted a historical conference in Bioacoustics in **1956** and those who attended included **Wagai, Kikuchi, Dussik, Bolt, Ballantine, Hueter, Wild, Fry** and **Howry**. Many of them met each other for the first time and important views concerning methods and instrumentations were exchanged at the meeting.

Kikuchi was very active in equipment designs, and by **1957** he was able to demonstrate the "<u>one-point contact-sector scanning tomography</u>" using the plan-position indication (PPI) B-mode format, which had a resemblance to a '**radar display**'. This development, which was at around a similar time as the pioneering work of <u>Howry</u> in Denver and <u>Ian Donald</u> in Glasgow (see below), had a similar concept of "position-referenced contact scanning".

Aloka® produced japan's first commercial medical A-scanner, the <u>SSD-2</u> and the water-bag B-scanner, the <u>SSD-1</u> in **1960** (pictured on the right). The application of ultrasound in Obstetrical and Gynaecological diagnosis started around 1956 with the A-scan basing on a <u>vaginal approach</u> and later Bscans at around **1962** basing on the use of the "<u>one-point contact-sector</u> <u>scanner</u>" in the PPI format. Early commercial water-bag scanners were being produced by Aloka® and Toshiba® in the early 1960s.





The water-bag B-mode scanning system, the SSD-1, from Aloka in 1960

their work after the first two papers presented at scientific meetings.

Masao Ide at the Musashi Institute of Technology in

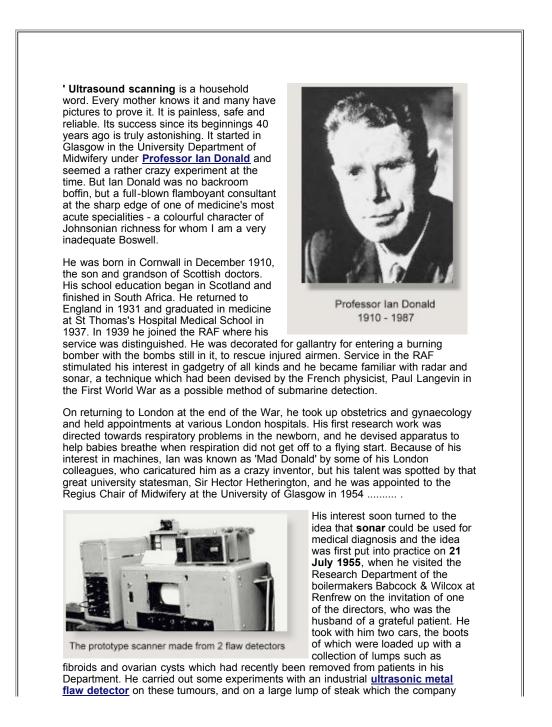
Kikuchi's one-point contact sector tomograph in 1957

Toyko, working with **Wagai** and others launched important pioneering research on the bioeffects of ultrasound. <u>William Fry</u> hosted another conference on ultrasonics in **1962** at the University of Illinois which served as a very important meeting point for researchers from the United States, Europe and Japan.

Michio Ishihara at the National Sanatorium Kiyose Hospital in Tokyo and **Hajime Murooka** at the department of obstetrics and gynecology, Oomiya Red Cross Hospital, Saitama, delivered the first paper on ultrasound diagnosis of gynecological masses in the Japanese language at the 19th Kanto District Meeting of the Japanese Obstetrical and Gynecological Society in **1958**, basing on the A-scan. Murooka had earlier in 1957 received instructions from **Wagai** on the A-scan methods at the Juntendo University. They described A-scan echoes in cancer of the cervix and also in the presence of different causes of uterine enlargement. **Wagai** published a review article in the use of ultrasound in Obstetrics and Gynecology in **1959**. The Murooka's group apparently did not continue Also read a short History of the development of Medical Ultrasonics in Japan.

J ohn Wild was back in England in 1954 to give a lecture on his new discovery and this was attended by <u>Val</u> <u>Mayneord</u>, Professor of medical physics at the Royal Cancer Hospital (now the Royal Marsden) who had also been experimenting with the Kelvin & Hughes® MK IIB metal flaw-detector in neurological diagnosis. Among the audience was <u>Ian Donald</u> who was then Reader in Obstetrics and Gynaecology at the St. Thomas Hospital Medical School in London and was about to take up the appointment of Regius Chair of Midwifery at Glasgow University. **Donald** was quick to realize what ultrasound had to offer.[#] Wild, while returning to Minnesota, had mainly concentrated his investigations on the diagnosis of tumors of the breast and colon using 15 MHz probes which had tissue penetrations of only up to 2 cm. In 1956, Wild published his landmark paper on the study of 117 breast nodules, reporting an accuracy of diagnosis of over 90 percent. Despite that, the ultrasonic method of tissue diagnosis which he so popularised did not reach the point of wide acceptance. Pioneering work in ultrasonic diagnosis in the field of **Obstetrics and Gynaecology** however, soon took off in **Glasgow**, **Scotland**.

The following is an excerpt from an <u>article</u> in the University of Glasgow publication 'Avenue' No. 19: January 1996 entitled '<u>Medical Ultrasound ---- A Glasgow Development which Swept the World</u>', by <u>Dr. James Willocks MD</u>, who had best described the circumstances of **Donald**'s early work :



had kindly provided as control material. (No one had the appetite for the steak afterwards!) Later he formed a link with the Kelvin & Hughes Scientific Instrument Company, and particularly with a young technician called Tom Brown. Quite by accident, Tom Brown had heard the strange tale of a professor who was attempting to use a metal flaw detector to detect flaws in women. He telephoned Professor Donald and suggested a meeting, and it was not long before Donald and Brown together with **Dr John MacVicar**, later Professor of Obstetrics & Gynaecology in the University of Leicester, plunged into an intensive investigation into the value of ultrasound in differentiating between cysts, fibroids and any other intra abdominal tumours that came their way.

Early results were disappointing and the enterprise was greeted with a mixture of scepticism and ridicule. However, a dramatic case where ultrasound saved a patient's life by diagnosing a huge, easily removable, ovarian cyst in a woman who had been diagnosed as having inoperable cancer of the stomach, made people take the technique seriously. 'From this point', Ian Donald wrote, 'there could be no turning back'. Results eventually appeared in print in The Lancet of 7 June 1958 under the arid title

Investigation of Abdominal Masses by



pages from the 1958 paper

Pulsed Ultrasound'. This was probably the most important paper on medical diagnostic ultrasound ever published. Ten years later all doubt had been cast away and Ian Donald was able to review the early history of ultrasound in a characteristic, forthright manner. 'As soon as we got rid of the backroom attitude and brought our apparatus fully into the Department with an inexhaustible supply of living patients with fascinating clinical problems, we were able to get ahead really fast. Any new technique becomes more attractive if its clinical useful department with an extent of the department with the patients. usefulness can be demonstrated without harm, indignity or discomfort to the patient Anyone who is satisfied with his diagnostic ability and with his surgical results

is unlikely to contribute much to the launching of a new medical science. He should first be consumed with a divine discontent with things as they are. It greatly helps, of course, to have the right idea at the right time, and quite good ideas may come, Archimedes fashion. in one's bath.'



In 1959 Ian Donald noted that clear echoes could be obtained from the **fetal head** and began to apply this information. I became involved shortly afterwards, and indeed was given the project to play with on my own. At the Royal Maternity Hospital, Rottenrow, there was no separate room to examine the patients and not even a cupboard in which to keep the apparatus, so my colleague, the physicist Tom Duggan, and I pushed it about on a trolley and approached patients in the wards for permission to examine them at the bedside. Glasgow women are wonderful method of fetal head measurement to assess the size and

James Willocks

growth of the foetus. When the Queen Mother's Hospital opened in 1964 it became possible to refine the technique greatly. My colleague Dr. Stuart Campbell (now Professor at King's College Hospital, London) did this and fetal cephalometry became the standard method for the study of fetal growth for many years.

Within the next few years it became possible to study pregnancy from beginning to end and diagnosis of complications like multiple pregnancy, fetal abnormality and placenta praevia (which causes life threatening haemorrhage) became possible. Professor Donald had gathered around him a team of talented young doctors and technologists, including the research engineers John Fleming and Angus Hall, who were engaged by the University when the Kelvin Hughes company was closed in 1966.

John Fleming has continued at the Queen Mother's Hospital as the technical genius behind all developments, and is also in charge of the valuable historical collection about diagnostic ultrasound. Practically all apparatus is now Japanese in origin, but the contribution of Scottish engineering to the development of medical ultrasound should never be forgotten.



lan Donald was also aware of the work of Howry in the United States and Kikuchi in Japan in the early 1950s, and had referenced these pioneers alongside with the work of Wild and Reid in his Lancet paper in 1958. Donald had felt that it was his fortune to have started with these historical A-mode and B-mode instruments instead of the apparatus that Wild and Howry had used, as these involved high frequency transducers (and hence associated with poor penetration into tissues) or a <u>water-bath arrangement</u> which could both become deterrants to further development in a medical setting ^{##}. Aside from this, **Donald** had on many occasions remarked that a lot of his developments in ultrasound was from a stroke of



accident, coincidence and luck. The '**full bladder**' was one, which he only discovered in 1963. That the fetal head, being a symmetrical skull bone could be easily demonstrable and measured accurately by a beam of ultrasound in an A-scan was another, as was the opportunity of meeting up with a number of important **administrators** on the way and working with the very bright engineer <u>Tom Brown</u> from Kelvin & Hughes®.

Brown, at the age of 24, invented and constructed with Ian Donald the prototype of the world's first **Compound B-mode** (plan-position indication, PPI) contact scanner in **1957**. The transducer operated at 2.5 MHz. The prototype was progressively improved to become the **Diasonograph®** manufactured commercially by **Smith Industrials of England** which had taken control of the **Kevin and Hughes Scientific Instrument Company** in 1961.

For a detailed account of the pioneering development of the prototypes, read an important unpublished paper by **Tom Brown** entitled <u>Development of</u> <u>ultrasonic scanning techniques in Scotland 1956-1979</u>.

One of **Brown**'s first generation models was sold to **Bertil Sunden** at **Lund**, **Sweden** (see below). The console design of the **Diasonograph**® came from **Dugald Cameron** who was then an industrial design student at the Glasgow school of Art. Brown also invented and patented an elaborate and expensive **automated compound contact scanner** in **1958** and it was at the



The prototype scanner that Donald and Brown built in 1957

machine's exhibition in London in 1960 that Ian Donald met for the first time <u>Douglass Howry</u> from the United States who had been using the much larger size water-tank circumferential scanner for several years (see above). **Donald** nevertheless had quoted in his <u>1958 paper</u> in the Lancet **Howry**'s work in B-mode scanning. The meeting had also influenced **Howry** and his team into producing a <u>similar compound contact scanner</u> like the Donald's although this had rapidly evolved into the <u>multi-joint articulated arm</u> version.

A brief description of the working of the prototype compound contact scanner (which eventually developed into the **Diasonograph**®) was given by Donald and Brown in **1958**, the same concept and design were extended into the later commercial models:



The Diasonograph produced in 1963

" A probe containing both transmitting, and receiving transducers is mounted on a measuring jig, which is placed above the patient's bed. The probe is free to move vertically and horizontally and, as it does so, operates two linear potentiometers, which give voltage outputs proportional to its horizontal and vertical displacements from some reference point. The probe is also free to rotate in the plane of its horizontal and vertical freedom, and transmits its rotation via a linkage to a sine-cosine potentiometer. The voltage outputs from this system of potentiometers control an electrostatic cathode-ray tube, so that the direction of the linear time-base sweep corresponds to the inclination of the probe, and the point of origin of the sweep represents the instantaneous position of the probe. The apparatus is so calibrated that the same reflecting point will repeat itself in exactly the same position on the cathode-ray tube screen from whatsoever angle it is scanned, and likewise a planar interface comes to be represented as a consistent line.

The echoes picked up by the probe are displayed on three oscilloscope screens: an A-scope display, a combined B-scope and PPI display on a long-persistence screen for monitoring: and a similar screen and display of short persistence with a camera mounted in front of it. The probe is moved slowly from one flank, across the abdomen to the other flank being rocked to and fro on its spindle the whole time to scan the deeper tissues from as many angles as possible."

The **automated scanner** which Brown originally designed to overcome the effects of motion variables did not catch on well, while the **Diasonograph**® was sold to many other parts of Britain and Europe including Sweden, London and Bristol, the place where another ultrasound pioneer, <u>Peter NT Wells</u>, a medical physicist, had been developing a different version of the <u>multi-joint articulated arm scanner</u> (basing on the **Diasonograph electronics**), independently from his American counterpart.



The NE 4102

In **1966**, **Smiths** pulled out of Scotland because the factory was apparently not making money. At the same time the US Supreme Court ruled against Smiths in favour of **Automation Industries** (formerly the **Sperry Company**) of Denver on the question of the so-called **"Firestone patents**" (Floyd Firestone's patent on flaw-detection devices in 1942, see above). As part of the settlement, Smiths undertook to withdraw both from the industrial and medical applications of ultrasound, and Automation acquired title to the collection of Smiths' patents on these subjects. This included the **Brown** patents on 2-D contact scanning.



NE 4102 scanning gantry

Smiths sold the medical business to **Nuclear Enterprises (G.B.) Ltd**. in Edinburgh, which took over the manufacturing of the **Diasonograph**® (see Tom Brown's <u>Recollections</u>). Ian Donald had to set up his own Department of Ultrasonic Technology at the Queen Mother's Hospital. He had <u>John Fleming</u> and **Angus Hall** back to help him. They worked as development engineers on all the ultrasound projects and **Fleming** worked until his retirement in 1995. He is co-ordinator of the <u>BMUS historical collection</u> and oversees the ultrasonic equipments at the Hunterian Museum, University of Glasgow. By 1968, **Brian Fraser** and **Alan Cole** at **Nuclear Enterprises** revamped the mechanical and valve design and redeveloped a new electronic system using semiconductor technology. The resulting "**NE 4102**" became a very popular instrument, and was used in most British hospitals and many European ones.



Donald in front of the NE 4102

For a detailed account of the pioneering development of medical ultrasonics in Glasgow, Scotland, read the biography of <u>Tom Brown</u> and an important unpublished paper by Brown on the <u>Development of</u> <u>ultrasonic scanning techniques in</u> <u>Scotland 1956-1979</u>.

Visit the pictorial presentation: <u>Scenes</u> from the History of Ultrasound from the <u>British Medical Ultrasound Society</u> (BMUS) Historical Collection. Co-ordinator: Mr. John Fleming.

Read also Peter Well's article on the

History of the development of ultrasonography.

Joseph Holmes and <u>Ian Donald</u> had subsequently become good friends across the Atlantic and Ian Donald and John Fleming were invited to speak on their experiences at the <u>International Conference</u> at Pittsburg hosted by Homles and others in **1965**. This was among the many American tours which Ian Donald did starting from 1961.

He spoke about Homles in a speech he gave in 1967 to the <u>World Federation for Ultrasound in Medicine and Biology</u> (WFUMB), 'I think Joe Holmes has done more than anyone to pull us all together from our several pathways'. Holmes became the founding editor of the Journal of Clinical Ultrasound in 1973.

Over in continental **Europe**, <u>Bertil Sunden</u> in Lund, Sweden, had started investigations in 1958 with Alf Sjovall, his professor in Obstetrics and Gynecology, on early pregnancies using an A-mode echoscope (a <u>Krautkramer®</u> reflectoscope USIP 9). The study on the application of ultrasound in Lund had already started formally in 1953 in cardiology and neurology (see above). Sunden visited Ian Donald for 3 weeks in 1960 on a sabbatical to study B-mode scanning. His work at Donald's department had resulted in the shipment of the <u>first generation Diasonograph®</u> to Lund, with which he produced his doctoral thesis on the use of ultrasound in Obstetrics and Gynecology, and reported his experience on 400 cases of pelvic pathologies. He also studied the possible harmful effects of ultrasound on pregnant rats, and did not find any. Sunden's thesis published in the Acta Obstet Gynaecol Scand in 1964 represented the earliest and the most comprehensive publication in Obstetrical and Gynecological ultrasonography at that time.



Donald, Takeuchi and others.

Pittsburg, 1965

Bertil Sunden C. 1972

Read also: <u>A short history of the development of ultrasonography in Lund</u>, <u>Sweden</u>.

At around the same time, **N D Selezneva**, a disciple of the famous Soviet scientist, <u>S Y Sokolov</u>, published his work in <u>ultrasonography in Gynecology</u> in the former **USSR** in the early 1960s. **R A Khentov**, **R A Khestova** and **I A Skorunskii** from the Central Institute of Advanced Training in Medicine, Moscow followed on with a large number of Russian publications in Obstetrics and Gynecology from **1965** onwards, using A-mode and later on B-mode equipments made at the USSR Scientific Research Institute of Medical instruments and Equipment. Almost ninety-nine percent of these publications were nevertheless in the Russian language.

▶ Read <u>further notes on early developments in Obsterical and Gynecological ultrasonography in the Soviet</u> <u>Union</u>

The Ultrasonic Boom



Alfred Kratochwil c. 1966

The increase in the research and application of ultrasound in Obstetrics and Gynecology appeared to **boom from 1966** onwards (see chart below) when there was an upsurge of centers and people in Europe, the United States and Japan that had begun to embark on studies in the application of ultrasound diagnosis in this specialty. **A-** and **B- mode equipment** were both in use including the first 'fast Bscanner', the <u>Vidoson</u>® from **Siemens**® (see **part 2**) used by **D**

Hofmann and Hans Holländer at the Wilhelm University in Münster, Germany.



<u>Alfred Kratochwil</u> at the Second University Frauenklinik, Vienna, Austria started working on placental localisation with

KretzTechnik AG in Zipf, Austria. He soon learned of Ian Donald's work with the B-scan and quickly collaborated the company to develop a similar device. The model 4100 originally designed for ophthalmologic use was adapted to carry an articulated-arm gantry (pictured below) for the abdominal B-scan mode. The articulated-arm design he found, was easier to manipulate than the Glasgow counterpart. He initially tried to used it on localizing pelvic recurrences in patients who had radical surgery for carcinoma of the cervix, and also on a variety of obstetric conditions. As early as 1972, **Kratochwil** had, among other endeavours, successfully demonstrated the visualisation of ovarian follicles with static B-mode ultrasound.

An early articulated-arm compound static scanner from KretzTechnik, Austria** A vaginal scanning device is seen on the left

Kratochwil soon became one of the most prolific users of the instrument and worked on areas such as the breasts and other surgical conditions, where he also published a number of important early papers. Since 1968 he developed training courses in ultrasound in Vienna and his department was visited by many hundreds of radiologists and obstetricians to learn about the applications of ultrasonography. Kratochwil was probably the most productive of all the investigators in Europe and was instrumental to the constantly improving designs at KretzTechnik AG.

Read a Short history of Kretztechnik AG, Austria. ⊩

Hans Henrik Holm, a urologist, started the ultrasound laboratory at the Gentofte Hospital in Copenhagen, Denmark in 1964, and with Jorgen Kristensen, Allen Northeved, Jan Pedersen, Jens Bang among others had established a strong research team. Holm also designed

their version of an articulated-arm scanner which subsequently was taken up for commercial production at Smith Kline Instrument® in the United States. The Copenhagen center had in time become a leading center in Interventional ultrasound, even up to this day.

Read a short history of the early development of ultrasonography in Copenhagen, Denmark.

And so it was that the early pioneers in diagnostic ultrasound from the **United States**, **Japan**, **United Kingdom**, **Austria**, **Germany**, **Sweden**, **Switzerland**, **Denmark**, **France**, **Poland**, **Holland**, **USSR** and **China** have all started with the A-scan basing on the **metal flaw detector** or a modification of the instrument. Many had first started their investigations in neurology, cardiology and ophthalmology, and only later on did they apply ultrasonic techniques to the abdomen and pelvis.

In Germany, at around 1950 both Siemens® and Krautkrämer® had started to make flaw-detecting equipment. Located close to the steel industry Krautkrämer® provided better service than Siemens® and soon dominated the market. After W Güttner and others had shown the impracticality of the transmission technique in 1952, Siemens had lost interest in diagnostic ultrasound. Around the end of 1956 the company decided to stop producing flaw-detection equipment completely. It was Inge Edler and Carl Hertz in Lund who adapted three of the Siemens® flaw detectors for cardiac investigations in 1957 (see above), and these were introduced back into hospitals in Germany. After a lapse of almost 10 years, the company developed the first fast B-scanner, the **Vidoson** in 1967, suitable for gynecological and abdominal examination (see **Part 2**). Germany was nevertheless one of the more 'prolific' of the European countries in terms of centers in early ultrasonic applications and research, with publications coming from Muchen, Erlangen, Bonn, Heidelberg, Berlin, Frankfurt, Freiburg and Bochum.

Read a history of the development of ultrasonography at Siemens, Germany.

Vienna in Austria, as noted above was 'historically' important because of the company <u>Kretztechnik</u> AG which produced some of the best and most advanced machines in the world at that time. The **B-scan**, basing on more sophisticated instrumentation emanating from radar sciences quickly evolved and replaced the A-scans. Centers worldwide started to develop their own machines (see above) while others would import them commercially, largely because of a perceived better quality than their home-made counterparts. For example, in the late 1960s some Finnish centers used Physionics/ Picker® machines from the United States and French and Italian centers used scanners from Nuclear Enterprise® and KretzTechnik AG. Smith Kline Instruments® scanners were used in Spain, Aloka® models in Brussels and the Siemens® Vidoson was employed by a number of centers outside of Germany.

Read a short history of the development of ultrasound in Obstetrics and Gynecology in France.



d LL

The "First World Congress on Ultrasonic Diagnostics in Medicine" was held in Vienna in **1969** and the second in Rotterdam in 1972 where an increasing number of papers in this specialty was presented. These meetings identified and brought together an international group of clinicians and scientists who started to contribute heavily towards the developments of ultrasonic instrumentation and methodology. In Europe, <u>Alfred Kratochwil</u> (1966), (Austria), D Hofmann (1966), <u>Hans Hollander</u> (1966), <u>Manfred Hansmann</u> (1966), (Germany), Malte Hinselmann (1968) (Switzerland), <u>Salvator Levi</u> (1967) (Brussels), <u>Hans Henrik Holm</u> (1967), Jens Bang (1967) (Denmark), <u>Georges Boog</u> (1969), Francis Weill (1969) (France), <u>I</u>



Roszkowski I (1968), Jerzy Groniowski (1968), (Poland), Paavo

Salvator Levi

Pystynen (1966), Pekka Ylöstalo (1971), Pentti Jarvinen (1968), <u>Pentti Jouppila</u> (1970) (Finland), J Hernandez (1970), R Montero (1970), Fernando Bonilla-Musoles (1971) (Spain), Bruno Damascelli (1967), L Roncoroni (1967), Alberto Zacutti (1968), C Brugnoli (1968), Achille lanniruberto (1970) (Italy), E Kalamara (1972), M Bulic (1972), <u>Asim Kurjak</u> (1973) (Yugoslavia, now Croatia), <u>Juriy Wladimiroff</u> (1974) (Netherlands), M Falus (1969), M Sobel (1969), L Kun (1973), P Bosze (1973) (Hungary), among many others, soon followed up with their many publications in obstetrical and gynecological sonography, although much of what was published was not in the English language. [The year in parenthesis denoted the year in which publications in Obstetrics and Gynecology from the particular author first appeared in the literature]. The delegates of 13 European ultrasound societies met in Basel, Switzerland in 1972 to form the <u>European Federation of Societies for Ultrasound in Medicine and Biology</u> (EFSUMB).

Read a brief history of the development of medical ultrasonics in Poland.



Peter NT Wells

In the United Kingdom, <u>Ellis Barnett</u>, <u>Patricia Morley</u>, <u>Hugh</u> <u>Robinson</u>, <u>Usama Abdulla</u> in Glasgow, <u>Peter Wells</u> in Bristol, A C Christie in Aberdeen, E I Kohorn, <u>Stuart Campbell</u> in London (see Part 2), <u>Hyton Meire</u> and <u>Pat Farrant</u> in Middlesex, and <u>Christopher Hill</u> at the Royal Marsden continued to make very important contributions in many areas.

Barnett and Morley's book in **1974**: "*Clinical Diagnostic Ultrasound*" was the first book (including publications from the United States) devoted to abdominal B-mode ultrasonography. **Peter Wells** in particular, was the single most important contributor to the advancement of ultrasound technology in Britain. <u>Stuart Campbell</u> eventually became one of the



Stuart Campbell

world's most well-known researcher and teacher in the field of Obstetrical and Gynecological ultrasound. The **British Medical Ultrasound group** was formed in **1969** by members of the Hospital Physicists Association and the British Institute of Radiology. The group later changed its name and became officially the **British Medical Ultrasound Society (BMUS)** in **1977**.

Read the early history of Obstetrical and Gynecological ultrasound in Finland.



Back in the **United States**, **J Stauffer Lehman**, in Hahnemann, Philadelphia was instrumental in the early 1960's to the continuing development of ultrasound technology in the **United States**. His association with **Smith Kline Instruments®** had been catalytic to the company's production of water-bag and contact B-mode scanners on top of their existing line of A- and M- mode equipments for echocardiography. The <u>LIFE® magazine</u> made an introduction to Ultrasound scanning at <u>Lehman's laboratory</u> in their <u>January</u> and <u>September</u> issues in **1965**. The **Family Circle® magazine** also reported on the medical use of ultrasound in their <u>October 1966</u> issue.



Barry Goldberg

Lehman's equipment was nevertheless cumbersome and expensive to fabricate and later on a smaller company, **Hoffrel** took up the

production of his machine. After the expiration of SKI's contract, Lehmann turned to use the <u>articulated arm</u> scanner originally invented and produced by the <u>Physionics Inc</u> in Longmont, Colorado (<u>later on acquired by</u> the <u>Picker</u> <u>Corporation</u> and further expanding its development).

Barry Goldberg joined Lehman in 1968 and expanded the research. He published extensively on a variety of subjects including echocardiography and interventional ultrasonography and was on record the first to describe fetal cephalometry in 1965 outside of Britain and Europe. George Evans, then a young Radiologist, was responsible for organizing the service and several important research projects. With his team was <u>Marvin Ziskin</u>. Together they have introduced ultrasound to the Radiological community in the United States and convincing them of the technique's clinical value. Lajos von Micsky, working at the St. Luke's Medical Center in New York, was also one of the important pioneers of <u>abdominal</u> as <u>well as endoscopic sonographic equipment</u>. He established a bioacoustical laboratory at the center in 1963 and devised many innovative abdominal, trans-vesical, rectal and trans-vaginal scanners.

Also read an article "Obstetric US imaging: the First 40 Years" by Dr. Barry Golberg.

Articulated arm scanners such as the PortaScan from Physionics Inc® produced in the mid-1960s had become the most popular format in compound contact B-scanners in the United States and throughout the world. Other earliest manufacturers of similar devices included the UniRad Corporation®. Newer machines soon followed from manufacturers in the United States and worldwide. These included the <u>Picker</u>® Laminograph 102, the KretzTechnik AG <u>Combison 1</u> and 2, the Nuclear Enterprise® Diasonograph 4102 (pictured above), the Aloka® <u>SSD-10 compound contact scanner</u> (pictured below) and the Toshiba® TSL systems. Jan C Somer and <u>Nicolaas Bom</u> in the Netherlands introduced the phased-array and <u>linear-array transducers</u> respectively in 1968 and 1971 (see Part 2).



Louis M Hellman, Mitsunao Kobayashi, Ross Brown, George Leopold, Roy Filly, Roger Sanders, Arthur Fleischer, Kenneth Taylor, Fred Winsberg, John Hobbins and William Cochrane were among those who produced a substantial amount of work from the early 1970s on the application of ultrasound relating to Obstetrics and Gynecology and had contributed much to moving the modality forward. Winsberg had a particular interest in realtime scanners and he was the first to use the German



History of Ultrasound in Obstetrics and Gynecology, Part 1



in the mid-70s, evolved from

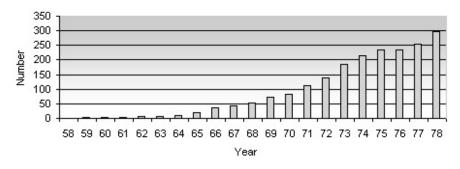
the Physionics Porta-scan

Vidoson® real-time scanner (see part 2) in North America (at the McGill University in Montreal, Canada) in 1970. One of the very earliest textbooks in sonography in the English language aside from Bertil Sunden's thesis was from Kobayashi, Hellmen and Cromb: "Atlas of Ultrasonography in Obstetrics and Gynaecology" published in 1972.



The <u>American Institute of Ultrasound in Medicine (AIUM)</u> which was founded in 1952 by a group of physicians engaged primarily in the use of ultrasound in physical medicine only started to <u>accept members</u> in the diagnostic arena in 1964. Diagnostic ultrasound has since then become the mainstream application in the organization. The "First International Conference on Diagnostic Ultrasound" was held in Pittsburgh, Pensylvannia in 1965 and was well attended by most of the ultrasound pioneers.

The *Journal of Ultrasound in Medicine*, the official journal of the **AIUM**, was inaugurated in **1982** replacing the **Journal of Clinical Ultrasound** as the association's main vehicle of communication with it's members. <u>George Leopold</u> was its founding editor. By the mid-1970s important producers of articulated compound B-scanners in the United States included the <u>Picker Corp</u>®, Smith Kline Instruments®, the UniRad Corporation®, Searle Ultrasound®, Rohe Scientific®, Litton Medical Systems® and Metrix Inc®. A list of manufacturers of static compound contact scanners as at **1975** can be found <u>here</u>.



The number of publications in the world literature each year on the application of ultrasound in Obstetrics and Gynecology rose from 1(Ian Donald's paper) in **1958** to 296 in **1978**. In the first 10 years, most publications were of a general descriptive nature and had similar titles to the effect of "The use of ultrasonography in Obstetrics and Gynecology".^{ref}



Hisaya Takeuchi

In Japan, Shigemitsu Mizuno, <u>Hisaya</u> <u>Takeuchi</u>, Koh Nakano and Masao Arima followed up the ultrasound work at the Juntendo University in Tokyo, and experimented with new versions of the <u>A-mode transvaginal scanner</u>. The first ultrasound scan of a 6-week gestational sac by vaginal A-scan was reported in the Japanese language in 1963. From 1962, the group worked extensively with the **water-bag** Bscanner, the Aloka <u>SSD-1</u> and was very active in many areas and producing a huge number of research publications, ranging from early pregnancy diagnosis to cephalometry to

placentography. They also reported on a large series of pelvic tumors in 1965, and in the following 2 years switched from the water-bag contact scanner to the articulated-arm compound contact scanner, the <u>SSD-10</u>. Another group consisting of **T Tanaka**, **I Suda** and **S Miyahara** started researches into the different stages of pregnancy in 1964.



Articulated-arm compound contact scanner the SSD-10 from Aloka used at the Juntendo University in late 1960's. The same model was used by Salvator Levi in Brussels in 1968

Shigemitsu Mizuno, <u>Hisaya Takeuchi</u> and their team also demonstrated in **1965** an **endovaginal scanner** for pelvic examination using the plan-position indication (PPI) B-mode format. The device was mannually rotated and the resulting display was very similar to a circular military 'radar" display. Used either transrectally or transvaginally, It was capable of producing some meaningful pictures of the pelvic organs. See <u>Hisaya Takeuchi</u> for a list of early work from the group.

The Japan Society of Ultrasonics in Medicine was officially formed in 1962. In the 1970s important work started at the Tottori Uinversity, Toyko under Kazuo Maeda, particularly on doppler fetal cardiotocography and at the University of Toyko under Shoichi Sakamoto. Toshiba® produced their first A-mode scanner, the SSA-01A and the compound contact B-scanner, the TSL system in 1967. Hitachi® produced their first A-mode (the EUA-1) and B-mode scanner (EUB-1) in 1971 and 72 respectively.

Also read a short history of the development of Medical Ultrasonics in Japan.



In the **Republic of China**, **Shih An** founded in **1958** the **Shanghai Ultrasonic Medical Research group** at the Sixth People's Hospital of Shanghai and his team included Tao-Hsin Wang and Shih-Yuan An. In the same year they started ultrasonic investigations using a modified



B-mode scanner produced at Wuhan, China in the early 1960s

metal flaw detector (the <u>Chiang Nan Type I</u>) manufactured at the chiang Nan Ship Building Plant. The group collaborated with investigators from the Shanghai First and Second Medical Colleges, namely Shih-Liang Chu, Hsiang-Huei Wu, Chih-Chang Hsu (Zhi-Zhang Xu) and Kuo-Juei Yu. They published in **1960** their <u>preliminary report</u> on the application of diagnostic ultrasound in various clinical conditions. This article which was published in Chinese in the 'Chinese Medical Journal' was not known to the west until two years later when their follow-up publication "<u>The use of pulsed ultrasound in clinical diagnosis</u>" appeared in the foreign language edition of the same journal. In these articles the diagnosis of <u>hydatidiform mole</u> with A-mode ultrasound was described, supposedly the first time in world

literature, where they demonstrated a significant increase in the number of small echo spikes between the proximal and distal uterine walls.

Further work in Obstetrics and Gynecology came from <u>Xin-Fang Wang</u> and **Ji-Peng Xiao** at the Wuhan Medical College (now Tongji Medical University) in **Wuhan**, **China**. In **1963**, the group reported on the sonographic findings in 261 abnormal pregnancies and in **1964** described fetal <u>M-mode echocardiography</u> which was probably the earliest of such reports in the medical literature⁵⁰. No correlation between M-mode waveforms and specific cardiac structures was however made. **Yong-Chang Chou** who had also been pioneering A-mode ultrasound diagnosis since the late 1950s at the **Shanghai Sixth People's Hospital** published a similar report in the next issue of the same journal (May, 1964).

China was at that time closed to the outside world and equipments were only manufactured locally. Apart from the Amode scanners, <u>B-mode equipments</u> were produced from a radar factory in **Wuhan**. One of the more important designs came from <u>Zhi-Zhang Xu</u> of the Shanghai Research group working at the ZhongShan Hospital. Regrettably progress was completely brought to a halt by the <u>Cultural Revolution</u> in 1966 and did not resume until the late 1970's.

In **Taiwan**, Republic of China, ultrasonic investigations started at the **National Taiwan University** in 1966, where J P Hung and Y C Chen used the Aloka® SSD-2C in the detection of mid-line shifts in head injuries and brain tumors. In the following year, their department and Obsteticians Hsi-Yao Chen and S M Wu had switched to the use of the B-mode **SSD-10** from **Aloka**® and published papers on B-mode cephalometry in **1971** in the chinese language. The Society of Ultrasound in Medicine, Republic of China (SUMROC) was founded in **1984**.

Also read a history of the Early development of ultrasonography in China. (partly in chinese).



George Kossoff c. 1975

Down under in Australia, the Ultrasonic Research Section at the National (formerly Commonwealth) Acoustic Laboratory in Sydney was established in 1959, with the objective of creating a center of technical expertise in the field of medical ultrasound. The section was headed by it's chief physicist <u>George Kossoff</u>. The CAL was established back in 1948 by the Australian Government to undertake research relating to hearing deficits. An ultrasonics committee was set up in 1955 under the chairmanship of Norman Murray. Murray visited Joseph Holmes' laboratory in 1958 and was impressed with the use of ultrasound as a diagnostic tool.



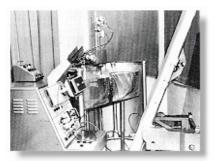
William Garrett

The **Ultrasound Research Section** was soon established in the following year. Working in conjunction with **William Garrett**, a

The original echoscope was replaced with a **Mark II** version in **1969**, which had already

incoporated

diagnostic method for placental localization, Kossoff introduced the water-coupling <u>CAL echoscope</u> in **1959** and perfected it in **1962**, which was also modified for breast scanning. His team also included **David Robinson**, who joined the Institute in 1961. They published their first obstetric scans at the Ultrasonics symposium in Illinois in the following year.



The CAL echoscope (MK I)

basic gray scaling in the images, before the invention of the 'scan-converter'.

The group reported **gray-scale obstetric scans** in **1971** at the International Biological Engineering meeting in Melbourne and then at the World Congress of Ultrasonic Diagnosis in Medicine in Rotterdam in **1973**. **David Carpenter** joined the Section in 1968, and headed the Engineering Research subsection. **Stanley Barnett**, a physiologist who



In 1968, **Garrett**, **Robinson** and **Kossoff** published one of the earliest papers in fetal anatomy "Fetal anatomy displayed by ultrasound" using the water-bath **CAL echoscope** that had brought out the role ultrasound would play in the diagnosis of **fetal malformations**. In **1970** they published <u>one</u> <u>of the ealiest papers</u> on the diagnosis of fetal malformation, reporting a

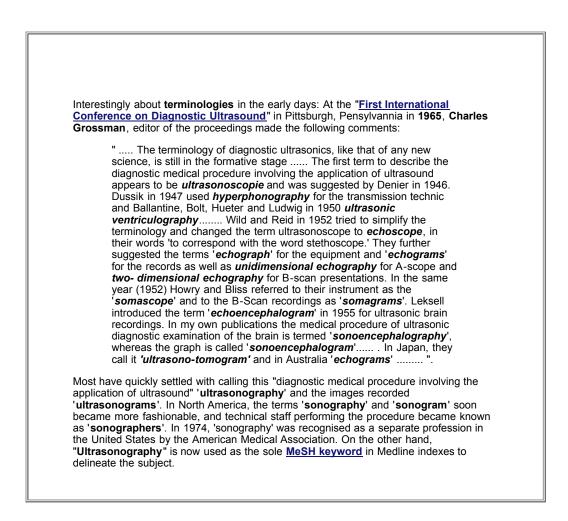
case of fetal polycystic kidneys at 31 weeks of gestation.

The UI Octoson

subsequently published extensively on ultrasound bioeffects joined the Section in 1970. Kossoff and his team developed sophisticated **annular dynamic phased-arrays** in **1974** which was installed in the mark II water-coupling echoscope.

In 1975, they constructed the UI Octoson, a rapid multi-transducer water-bath scanner which had then incoporated the new scan-converter, improved annular array transducers and more powerful computing electronics that had allowed for superior compound scans to be completed in less than 1 second. The scanning mechanism of the Octoson is completely immersed in the coupling tank and the patient, lying prone, is examined from below. (see also Part 2)

Also read An historical look at ultrasound as an Australian innovation by Kaye A Griffiths.



Go to [Part 2] and [Part 3]

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Acknowledgements:

Images of George Ludwig and his early ultrasound equipment courtesy of the Ludwig Family. Reproduced with permission. Image of the Denver somascope reproduced with permission from <u>Mr. GJ Posakony</u>. Images of the Denver pan-scanner and Dr. Toshio Wagai were reproduced with permission from <u>Dr. Barry Goldberg</u>, Chairman of the Archives Committee of the <u>AIUM</u>. Pictures of this, as well as other early scanners can be found in the Eastman Kodak Health Sciences publication, "Medical Diagnostic Ultrasound: A retrospective on its 40th anniversary" by Drs. Goldberg and Kimmelman published in 1988. Image of Dr. John Wild courtesy of Dr. Wild. <u>Dr. William O'Brien Jr.</u>, Professor, Bioaccoustic Research Laboratory, Department of Electrical and Computer Engineering, University of Illinois. ** Courtesy of <u>KretzTechnik</u> **AG**, Zipf, Austria. Images of the NE 4102 reproduced with permission from Dr. RG Law, from his book 'Ultrasound in Clinical Obstetrics', John Wright and Sons Ltd, Pristol 1980.

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Pictures of Professors Bertil Sunden and <u>Salvator Levi</u> courtesy of Professor Levi.
^ Courtesy of the Department of Ultrasonics, Polish Academy of Science.

ref raw data from "Ultrasound in Biomedicine - Cumulative Bibliography of the World Literature to 1978" by Drs. Denis White, Geraldine Clark, Joan Carson and Elizabeth White. Pergamon Press 1982.

³ The story of the early development of sonar in Glasgow was vividly narrated in the article "Sonar -- the Story of an Experiment" by Professor Ian Donald which appeared in Ultrasound in Medicine and Biology, vol 1 pp109-117, 1974.
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 Every effort has been made to ensure accuracy in dates, persons and events.
 It is not possible to include all the names who have contributed significantly to the advancement of Obstetrical and Gynecological sonography, some who may have been less well-known than the others and some who may not have published so extensively in the English language Apologies are extended to those whose contribution has not been fully credited in this article.

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A short History of the development of Ultrasound in Obstetrics and Gynecology

Dr. Joseph Woo



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Pushing ahead, new technology and new techniques

The A-mode scan had been used for early pregnancy assessment (detection of fetal heart beat), <u>cephalometry</u> and <u>placental localization</u> in Europe, Britain, United States, Japan, China, USSR, Poland and Australia in the early 1960s, the measurement of the <u>biparietal diameter</u> (BPD) having been invented by <u>Ian Donald</u> in **1961** and further expanded in his department by <u>James Willocks</u>, basing on improvements in the 'bright-up' markers and the electronic caliper system. The measurements were done 'blindly' without actually seeing the structures under study. Visualising the **gestational sac** by **B-mode** ultrasound was first described by the **Donald** and **MacVicar** team in **1963**. In 1965, they were able to demonstrate a 5-weeks gestational sac. The **Gestational sac diameters** in the assessment of fetal maturity was described by Lou **M** Hellman and **M** Kobayashi in **1969** and by <u>Pentti Joupilla</u> (Finland), <u>Salvator Levi</u> (Brussels) and **E** Reinold (Vienna) in **1971** in relation to early pregnancy complications. Kobayashi also described the ultrasonic appearance of **extra-uterine pregnancy** using bi-stable B-mode ultrasound in **1969**. Kenneth Gottesfeld in Denver reported in **1970** a large series of patients where fetal death in utero was diagnosed solely on bistable ultrasound scan.



Stuart Campbell

The ability to recognise and confirm the presence of fetal cardiac action in early pregnancy was considered to be one of the **most indispensible** use of ultrasonography (and still is). Although detection of fetal hreartbeat by the <u>A-scan</u> and audio doppler ultrasound (the first "<u>Doptone</u>" was invented in **1965**, see section below on doppler) had been variously reported by early groups such as <u>Wang</u> (1964, M-mode from 10 weeks), <u>Kratochwil</u> (1967, vaginal A-scan from 7 weeks), <u>Bang and Holm</u> (1968, A- and M-mode from 10 weeks), it was not until **1972** that <u>Hugh Robinson</u> in Glasgow, basing on improved instrumentation reported a practically useful 100% detection of fetal cardiac action from **7 weeks onwards**. The fetus was first located with B- scan ultrasound and the heartbeat obseved with a directed beam in A- and M-mode (also see below). This breakthrough has profound implication in the management of **early pregnacy bleeding** and **threatened miscarriages**.

B-mode <u>placentography</u> was successfully reported in **1966** by the **Denver group** in the United States and the **Donald group** in **1967** (<u>Usama Abdulla</u>).

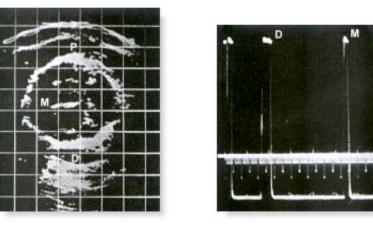
Ultrasonic diagnosis of **molar pregnancies** was described as early as **1963** by the same group.

Stuart Campbell's landmark publication in **1968** "An improved method of fetal cephalometry by ultrasound" described the use of both the A- and B-mode scan to measure the fetal biparietal diameter. This elegant and practical 'maneuver' had quickly become standard practice in an ultrasound examination of the fetus for the next 10 years. Operating the static scanner skillfully and effectively has also become a crafted art. In **1971**, with improvements in the caliper system, Campbell and Newman published normograms for the biparietal diameter from the 13th weeks of gestation and has made cephalometry a standard tool for the assessment of fetal growth and maturity. Many early paper in cephalometry followed in the late 1960s



Localisation of the placenta another indispensible use of ultrasonography

such as those from **Boog** in France, **Khentov** in the USSR, **Zacutti** and **Brugnoli** in Italy, **Kratochwil** in Austria and **Pystynen** and **Ylostalo** in Finland.



The early bistable oscilloscopic B-scan image at the level of the BPD and the A-scan tracing showing cephalic (P and D) and midline echoes (M). The distance between the 2 cephalic echoes is the BPD. Without scan converters on-screen (oscilloscope) measurements on the B-mode image are not possible. Very accurate measurements can however be made using the A-scan calipers.



The concept of blighted ovum started with advent of the B-scan Two years later in **1973**, measurement of the fetal <u>crown-rump length</u> was described by <u>Hugh Robinson</u> in Glasgow who was then a research registrar. Life size magnification of the images had become possible with the newer machines which enabled accurate measurements to be made on early embryos. In **1972**, the Scottish group, basing on the ultrasonic findings, expounded the concept of **'blighted ovum'** in Obstetrics, first described by **Ian Donald** in 1967, which had changed considerably the concept and management of pregnancies with vaginal bleeding in the first trimester.

<u>Horace Thompson</u> in **Denver** has described in **1965** measurement of the <u>thoracic circumference</u> (**TC**) as a method for studying fetal growth. The measurement had an accuracy of within 3cm in 90% of the patients. Thompson

also introduced the idea of **fetal weight estimation** basing on the TC and estimates were found to be accurate to within 300 grams in 52% of patients. The resolution of images at that time however, did not in general allow for very accurate measurement of the fetal trunk. <u>Manfred Hansmann</u> in Bonn, Germany re-introduced the <u>thoracic circumference</u> in 1972 and similarily correlated it with the birth weight of the fetus. In the same year he also described intrauterine transfusion under ultrasonic guidance. William Garrett and David Robinson in Sydney had also reported on measurement of the fetal trunk area as a means to assess fetal size. The group used a water-bath Echoscope which by 1970 had already incorporated some degree of gray scale capabilitiy allowing for better visualization of the fetal trunk.



Horace Thompson

With the static B-scan, <u>Campbell</u> reported the diagnosis of an anencephalic fetus at 17 weeks in **1972** and the diagnosis of spina bifida in **1975**. Both had appeared as landmark papers in the Lancet. They were the first cases of these conditions in which a correct diagnosis by ultrasound had effectively led to a termination of pregnancy. In **1975**, the **Campbell** group introduced the measurement of the <u>abdominal circumference</u> which has since then remained the most important single parameter to assess fetal weight and nutrition. Circumference measurements of the fetal trunk is considered superior to diameter measurements as the former is less affected by the change in shape of the fetal body. It was not until **1979** that **Margi Mantoni** and **Jan Fog Pederson** in **Copenhagen** first described the visualization of the **yolk sac**, using the static B-scan.

Also read a brief discussion on the historical aspects of the fetal trunk circumference measurements.

It was round about this time in the *History of Ultrasonography* (the mid-70s) that several important developments in ultrasound instrumentation took place.

Gray scale and the Scan Converter



Pioneering designs in <u>electronic circuitries</u> were made in conjunction with the development of the B-scan, these included the pulse-echo generator circuitry, the limiter and log amplification circuitry and the demodulator and time gain compensation circuitries. Early Bscanners employed threshold detection which registered echoes on a phorsphorous coated oscilloscope screen as dots of light. A '**storage**' or '**bi-stable**' cathode-ray tube was used. Echoes above a certain amplitude are displayed as dots of **constant** intensity and echoes of a lesser amplitude below the threshold were not depicted. The images were often recorded unsatisfactorily on ordinary 35mm photographic film. The situation was improved with the advent of the black and white "peel-apart" **polaroid® instant** film which had become



available from 1959 onwards in both England, Europe and the United States. Although there was good representation of size, shape and position, the images did not depict differences in echo amplitude. It was apparent that some sort of gray-scaling was imminently necessary to expand the diagnostic capability and accuracy of a B-scan.

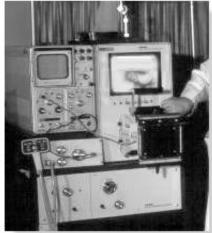
The early B-scan with the bistable oscilloscope **

The most important innovation in ultrasound imaging subsequent to the

invention of the compound contact scanner was the advent of the scan converter. The cathode ray tube had a low dynamic range of about 16 decibels. Early attempts at creating a 'gray scale' was through the use of a short-persistence oscilloscope and varying the time of shutter exposure in the photography. This could manage roughly 4 shades of gray in the final picture but the process was difficult to control and results unpredictable. In later developments prior to the appearance of the scan converter, echoes were compressed using sophisticated logarithmic compression amplifiers to accomodate the maximum amount of information into the range and a useful degree of grey-scaling could be managed with this principle.

Machines such as that developed in Glasgow in the mid-1950's were actually grey-scale ready from the outset. The function of the signal processor was to provide a degree of time-domain pulse shaping, in an attempt to separate echoes arriving closely-spaced in time and secondly to enable the display to record the very large dynamic range of signals (at least 60 dB, a range of signal amplitudes of more than 1000 to 1) which

were received, without going into 'hard limiting' at the top end, or suppression of small echoes at the bottom end. The focus at that time was rather more on



Scan converter incoporated in early American model. Image is being displayed on the TV monitor on the right Note the black-on-white display format

"spatial noise" reduction by signal integration as it was on accommodating the large dynamic range of the received signals.



Videotaping became possible with the advent of the scan converter

The analog scan converter which was hailed as a profound "invention" in the field of image processing and reproduction, used a silicon oxide/silicon target that acted as a capacitance matrix and was then raster-scanned by an electron beam that 'read' it and displayed the information on a standard television monitor unit. By doing so, computerprocessor technology, which was just up and coming at about the same time, could be applied to process the signal. Images could then be scaled, calipers moved and applied on-screen (something that had changed entirely the way measurements are made), gray-scaling applied to the images and the resultant image recorded on a variety of media including videotape, emulsion films and thermal printer devices.

The application of true gray scaling had evloved from the work of the Kossoff group at the Ultrasonic Institute in Sydney (formerly the National Acoustic Laboratories), Australia.



George Kossoff c. 1975

George Kossoff, chief physicist and director of the Ultrasonic Research Section, had been inventing and refining ultrasound apparatus for a variety of purposes including ophthalmic applications since 1959. Together with William Garrett, a gynecologist, George Radovanovitch and David Carpenter, two brilliant engineers, they published their new scan converter with **gray scale** capabilities in **1973**, basing on work which they had already started in **1969**. They demonstrated that the clinically relevant echoes that affected the magnitude of reflected echoes were those that came from the internal texture of soft tissues, and in gray-scale imaging they optimised the signal processing to display this textural information. By about '73-'74 other centers in Britain and Europe have also published on their version of gray-scale equipments. In 1975 the Kossoff group constructed and demonstrated the UI Octoson®, a rapid water-bath scanner employing 8 annular dynamic phased-array transducers which achieved it's scans by a combination of mechanical rocking and sequential pulsed-echo operations. The machine



produced very impressive images at that time compared with the European counterparts.

A similar mechanism was evaluated at the Royal Marsden Hospital in England at the same time under Kenneth Taylor and David Carpenter, who was visiting engineer from the Kossoff group. Together with Christopher Hill and VR McCready the group published their experience with gray scale imaging in 1973 and demonstrated their version of the compound gray-scale contact scanner. (Taylor moved to Yale University in 1975). Gray scale equipments had soon become widely available commercially by 1976. Gray scale sonography, as Kossoff had put it "had the shortest transition phase between development and acceptance" because improvements in the quality and 'interpretability' of the images were truely dramatic. The addition of gray scale had been instrumental at that point in time to the evolvement of the measurement of the fetal abdominal circumference, first discribed by the Campbell group at King's College Hospital; and to the assessment of fetal malformations and gynecological pathologies.



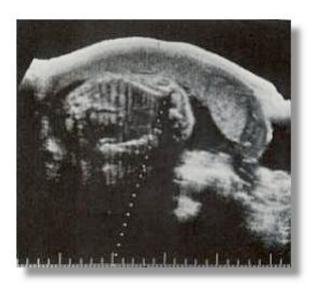




A bistable bscan image of the maternal abdomen showing abdominal circumference and placenta using a compound contact scanner (Diasonograph®) without grayscale in the late 1960s.

▶ B-scan image with gray scale of a similar section of the maternal abdomen showing abdominal circumference and placenta using the Nuclear Enterprise® NE 4102 in the late 1970s

A gray scale Octoson® image of the abdominal circumference and placenta in the late 1970s. The Octoson® produced superior images as compared to articulated arm scanners but loosed out on mobility and flexibility.



A gray scale longitudinal scan of a section of the fetal trunk and placenta made with the very popular Picker® 80L static scanner in the early 1980s. Despite the very good images that could be obtained with these machines. they were soon replaced by the new real-time scanners.

With advancement in computer electronics, the analog scan converter was soon being replaced by <u>digital scan converters (DSC)</u>. <u>Albert Waxman</u> and others at **Searle Ultrasound** produced one of the earliest DSCs in **1976** with a 256 by 240 pixels memory. Position and velocity data were fed through a <u>PDP-11</u> mini-computer (Digital Equipment Corporation) and logic circuits sampled the data from the analog front end and wrote it to the correct place in DRAM, and at the same time reading out the data continuously for display on a grayscale video monitor.

The digital scan converter provided for enhancement in peak detection operation for larger pulse echo amplitudes, and the integration-normalization analysis for smaller amplitudes, resulting in great improvements in the grey scale quality and resolution of the ultimate diagnostic image. Non-uniformities and distortions, both of linearity and of gray scale that were associate with analog converters could be avoided. Small echo amplitudes emanating from



The DEC (Digital) PDP-11

interfaces between similar tissues bearing useful image information could be preserved. Multiple image storage capability was improved, and write and erase speed were also much enhanced to permit real time imaging (see below).

In the late 70s' DRAM memory boards (mainly from Intel®) and analog-digital converter electronics were expensive necessitating new and more targeted designs. New general-purpose microprocessors (see below) were starting to appear which gradually replaced the large **PDP-11** (<u>picture</u>). The early Searle design also incorporated digital differential analyzer, rate multiplier and peak value detector circuits. Image clarity was very substantially improved from the analog counterparts. The bulky size (kitchen cabinet) of the static machines with more advanced digital conversions seen at that time was partly contributed to by the computer electronics housed inside. A number of designs even up to the first few years of the 1980s still incorporated the entire <u>PDP-11</u> inside the console. By this time new and advanced microprocessors such as the **8086** from **Intel®** and the **6800** from **Motorola®** were coming into the market (see below).

Hitachi in Japan produced their scanner, the EUB-20Z in **1978** incorporating what they claimed to be the <u>world's first DSC</u> in an array ultrasound scanner. **David Robinson** and <u>George Kossoff</u> in Australia also described one of the earlier DSCs in **1978**, employing a 512 by 512 pixels digital memory. The images were stable and the simultaneous read and write capabilities allowed for versatility in processing the image. 4 bit (16 shades of gray) and <u>5 bit (32 shades)</u> machines had become available. **Aloka**® in Japan produced a similar digital device in the same year and was incoporated into their production models.

Subsequent developments in pulse-echo imaging and scan conversion was based on the recognition that ultrasonic echoes originate not only from major interfaces but also from the smallest mechanical structures of the human body. The advancements were therefore directed towards the detection of small echoes in the presence of noise and to display the subsequent information in the fullest dynamic range of **spatial details** and **echo amplitudes** and calling for **smaller spot size** and wider range of **brightness levels** in the display.



With early scanners made of vacumn tubes there was often problem of **drifts** and **numeric instability** which required **periodic re-calibration**. Another important consideration in the early days was the assumed velocity of ultrasounds in human tissues. Different centers adopted different values, ranging from 1540 m/s, 1580 m/s and 1600 m/s, and this had to be stated in their reports and papers. By about the early 1980s and after several meeting of the experts, the ultrasound community throughout the world settled for **1540 m/s**. This universal acceptance is important because with different velocity calibrations measurements like the biparietal diameter will end up with different normals.

Attenuation charcteristics, axial and lateral resolution, fluid enhancement characteristics



Tissue phantom imaging quality and accuracy of the scanner. Hospital physicists are often pre-occupied with such calibrations. With the later advent of array real-time scanners these tissue phantoms progressively became obsolete and by about the late-80s they have completely disappeared from the scene and few are bordered by the figure of 1540 m/s.

Also of interest to mention here is the **black-on-white** (b-o-w) display format that was commonly used in the United States prior to the mid-1980s. British, European and most other parts of the world had started with a **white-on-black** (w-o-b) display and had stuck to it since. It was initially thought that the b-o-w display format might visually allow the operator to



The black-on-white display

appreciate more detail on the screen. However, there were a number of reports in the early '80s which indicated excessive **eye-strain** to the operator who needed to look continously at the brighter b-o-w monitor screen in a dimmly lit examination room ^^. Eventually, there was a complete switch in the United States to the w-o-b display format , which was supposed to produce less eye-strain on the operator.

Real-time, the real revolution

The innovation which had soon **completely changed the practice of ultrasound scanning** was the advent of the **Real-time scanners**. The first real-time scanner, better known as fast B-scanners at that time, was developed by **Walter Krause and** <u>Richard Soldner</u> (with J Paetzold and **and Otto Kresse**) and manufactured as the <u>Vidoson</u>® by Siemens Medical Systems of Germany in 1965. D Hofmann, H Holländer and P Weiser published it's first use in Obstetrics and Gynecology in 1966 in the German language. Hofmann and Holländer's paper in 1968 on "<u>Intrauterine diagnosis of hydrops fetus universalis using ultrasound</u>" also in German, is probably the first paper in the medical literature describing formally the diagnosis of a fetal malformation using ultrasound.

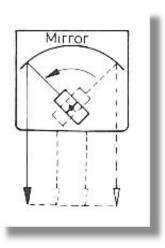


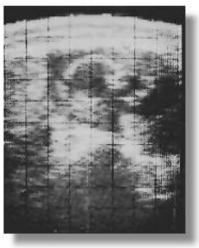
Richard Soldner

The <u>Vidoson</u> used 3 rotating transducers housed in front of a parabolic mirror in a water coupling system and produced 15 images per second. The image was made up of 120 lines and basic gray-

scaling was present. The use of fixed focus large face transducers produced a narrow beam to ensure good resolutions and image. Fetal life and motions could clearly be demonstrated.







The Vidoson*, its working mechanism and the resultant image of a fetal face and hand. The transducer housing is mounted on a mobile gantry and rigidly connected to the main console. The scanning frequency was 2.25 MHz. Scaling and caliper functions were not present.

Hans Holländer, in his paper in 1968 demonstrated the usefulness of a 'real-time' scanner in the diagnosis of ovarian tumors which were not spotted on pelvic examination. Malte Hinselmann, using the Vidoson, demonstrated in 1969 the universal visualization of fetal cardiac action from 12 weeks onwards. The Vidoson was popular in the ensuing 10 years or so and were used in many scientific work published from centers in Germany, France, Switzerland, Austria, Belgium, Italy and other European countries. The initial popularity was not based on its image resolution but rather its ability to allow the operator to display and study movements, such as fetal cardiac motion, gross body movements and fetal breathing movements (see also Part 3). In the International Symposium on Real-time ultrasound in Perinatal Medicine held in Charleroi, Belgium in 1978, most of the presentations were based on results from the Vidoson.

- Read a history of the development of ultrasonography at Siemens, Germany.
- Read also an article <u>40 Years of Realtime-Ultrasound Diagnostics</u> by Hans Hollander.

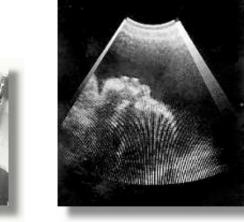
James Griffith and Walter Henry at the National Institute of Health produced a <u>mechanical</u> oscillating real-time scanning apparatus in 1973 which was capable of producing clear 30 degree sectoral real-time images of good resolution. The scanner was essentially a motordriven oscillating transducer coupled with a commercially available one-dimensional echocardiograph (the SmithKline Eckoline 20). The 2-D scanning device was hailed as one of the most significant milestones in the development of echocardiography, and indeed in the development of sonography in general. Other mechanical systems published included an oscillating design with membrane-oil coupling from <u>W N McDicken</u> in Edinburgh (1974, produced commercially as the EMI® Emisonic 4260), a continuously rotating wheel with radiallymounted transducers from <u>Hans Hendrik Holm</u> and Allen Northeved in Denmark (1975), and a single transducer direct-contact design from **Reginald Eggleton** in Indiana (1975). The design, which was supposed to have been modified from the mechanisms of an "electric toothbrush", was a commercial success. **Toshiba**®, in Japan produced their first prototype real-time



Carl Kretz

mechanical sector scanner in 1975, the <u>SSL-51H</u>. A number of others were available commercially soon afterwards and sold well such as the circular rotating system <u>Combison 100</u> from <u>Kretztechnik</u>® of Austria (1977), produced under the ingenuity of <u>Carl Kretz</u>.

Although these have relatively heavy probes they produced outstanding real time resolution in the near and far field (because of highly focused beams resulting from the relatively large curvatured transducers and the lens apparatus) and with much less image-degrading electronic noise that was associated with electronic scanners that soon became available at around the same time.



The large hand-held circular rotating transducer (Combison 100) from <u>KretzTechnik</u>® and the resultant sector image. The transducer is connected to the main console by a flexible cable.

The concept of the multi-element **linear electronic arrays** was first described by **Werner Buschmann** in an <u>ophthalmologic application</u> in **1964** in East Berlin. His probe, fabricated in collaboration with <u>Kretztechnik AG</u> consisted of 10 small transducers mounted on an arcshaped appartus to fit over the eye. Buschmann's transducer however, never became very functional in a clinical setting and did not go into serial production. A number of similar designs followed on the same concept. Jean Perilhou and her group in France, working under the auspices of the Philips® Company, described a <u>multi-element scanning array</u> in **1967**, although they do not produce images in a real-time fashion. The real-time array concept was further expanded by <u>Nicolaas Bom</u> at the Thoraxcenter, University Hospital, Erasmus University in Rotterdam, the Netherlands. His initial design in **1971**, which was described in his application for a Dutch patent, consisted of only **20 crystals** (each 4mm x 10mm). The probe face was 66 mm long and 10 mm wide and produced 20 scan lines. It operated at a frequency of 3.0 MHz



Nicolaas Born

sweeping at a frame rate of 150 frames/sec. The axial resolution was 1.25 mm while the beam width at 6 cm was 10 mm. This abeit simple and inadequate design at that time has evolved into the very sophisticated real-time scanners that are widely available today.



Multiscan system from Organon Teknika, 1972

In collaboration with cardiologist **Paul Hugenholtz** and local Dutch company **Organon Teknika**, they produced in **1972** the "<u>Multiscan system</u>", notably the **earliest** commercial linear array scanner in the world, mainly aimed at cardiac investigations.

The transducers operated at either 2.25 or 4.5 MHz, again with 20 crystals producing 20 scan lines. The lateral resolution of this improved version at a dynamic range of 10dB was 3.7 mm at 6 cm and 6 mm at 10 cm depth. It did not sell very well though because of its <u>relatively primitive resolution</u> and its inability to image abdominal structures adequately.

In Japan, **Rokuro Uchida** at **Aloka®** (see also **Part 1**) had similar research on the array technology in the late 1960s predating their European counterpart. In **1971** they published in Japanese (and presented at the Japan Society of Ultrasonics in Medicine) a system based on 200 closely interspaced transducers. Electronic

switching and use of overlapping groups of 20 small elements yielded 2-D images with a field depth of 20 cm at a rate of 17/frames per second. The company produced their <u>first</u> <u>prototype linear array scanner</u> in the same year. The model however, was not produced commercially or given a model number. The <u>first commercial linear array</u> <u>scanner</u> from **Aloka®** only debuted in **1976**. **Toshiba®** produced their first commercial real-time linear array counterpart in the same year, the <u>SSL-53H</u>, aimed at abdominal applications. Like the Aloka® this was a huge machine considering present day fabrication standards.



Subsequent to **Bom**'s work and the research in Rotterdam, <u>Leandre</u> **Pourcelot** and **Therese Planiol** in



Aloka's first linear array scanner, 1976

Tours, France was experimenting with more advanced segmented sequence transducer-array scanning possibilities in **1972**, in order to



Pourcelot's prototype real time scanner, 1972

enhance the lateral resolution of the devices. **Donald L King** at the **Columbia University** in New York described a 24-elements segmented sequence linear-array cardiac scanner in **1973**, in collaboration with the **Hoffrel Instruments Inc.**®, at Norwalk, Connecticut. In his design 3 crystal elements were fired simultaneously to produce a single pulse of ultrasound. The echoes returned from the 3 reflections were written into a single line on the scan. The crystals were stepped in a "1,2,3,... 2,3,4,... 3,4,5..." manner. There was however no delay lines for the implementation of <u>'focusing' techniques</u>.

Also concurrent to **King**'s work was the work in from the **Tony Whittingham** group at Newcastle-upon-tyne in England, where crystal stepping techniques were also being investigated.

"..... I tried various ideas, but the one that worked was to make an array of very narrow rectangular elements and to use a group of these to form a square aperture. This group of elements defining a composite transducer would scan the line in front of itself. Then you drop one element off from one end, put another element on at the other end, infront of the group, and advance the active group along the array in this way. When I was doing this I was totally unaware that it would work. I hope it would work, but I was worried that there would be cross-coupling from the end elements of the group into what should have been passive elements, so that you might not be able to get a well-defined active aperture. But it did work, and that proved to be the way forward, because you could make finer and finer elements and get more lines into the array" -- Tony



Tony Whittingham

Whittingham, describing his work in real-time imaging in the mid 1970s. ^

Images of the earlier models were nevertheless hampered by the problems of small crystal size, lobe artifacts, unwanted specular reflections, low dynamic range, unsatisfactory lateral resolution and image noise from electronic processing. There was an overwhelming need for the refinement of beam characteristics. Fredrick (Fritz) Thurstone, Olaf von Ramm and H Melton Jr at the Duke University published some of the earliest and most important work on electronic focusing using annular arrays ('71-'74), both on transmit and receive. Similar techniques were subsequently employed in the design of linear arrays transducers. Basing on these designs, a number of centers and private laboratories were starting to embark on making machines geared to examination of the abdomen. Albert Macovski at the Standford University filed a patent in 1974 for a circular array where the elements generate dynamically focused beams that could also be swept through space by adjusting the delays to the array. This was one of the more advanced designs in dynamic focusing techniques. Another important design - "signal processor for ultrasonic imaging" was described by the William Beaver group of the Varian Associates® in Palo Alto in California in 1975 where selection of scan angles and focusing distances were effected. George Kossoff in Australia also filed a patent in 1973 on a linear array system incoporating phased-focusing electronics. A summary of the advances in design can be found with M Maginness's article (at Stanford) "State-of-the-art in two-dimensional ultrasonic transducer array technology" in 1976.



The ADR realtime scanner ****

It was Martin H Wilcox, founder and engineer at the Advanced Diagnostic Research Corporation (ADR®, a company founded in 1972 in Tempe, Arizona), who designed and produced one of the earliest commercially available models of a linear-array real-time scanner in 1973 and very much set the standard for subsequent designs to follow. The array contained 64 crystals in a row (3 times the number in the earlier cardiac counterparts and 3 times as long and wide), fabricated with the best material available and in the best accoustic configurations and using 'stepping' crystals techniques. This was the first 'good-resolution' abdominal linear-array scanner that was in the commercial market.

Their second model <u>the 2130</u> marketed in **1975** had brought the linear-array principle and the

application of '<u>focusing techniques</u>' to commercial fruition. It was a big hit in the United States and had sold over 5000 units worldwide, including Germany and other European countries. The machine was marketed in Europe under the **Kranzbuehler** label. In 1980 a new 3.0 MHz variable focus transducer was added on to the 2130. The new transducer contained **506 crystal elements**, boasted both mechanical and phased focusing, improved gain and reduced noise, much quieter transducer operation, and switchable focal zones. The image had twice the number of data lines and probably the best real-time resolution in the industry at that time.



Marty Wilcox c.70s

" In Dallas, Texas, Ian was shown the first real-time scanning machine brought from Phoenix, Arizona, by some talented young men. Ian was of course, *wildly* excited. They wanted to carry him off to Phoenix to show him more, but sadly Ian couldn't change his next commitments. However, it wasn't too long before he had one of his own."

--- Alix Donald, wife of the late <u>lan Donald</u>, speaking in 1998 about their first encounter with the ADR real-time scanner in the early 1970s. ^{ref.}

ADR® merged with ATL® (Advanced Technology laboratories®, see below) in 1984. ADR® produced the 2150 in 1980 and the last model under the ADR label, the <u>ADR 4000</u> in 1982.

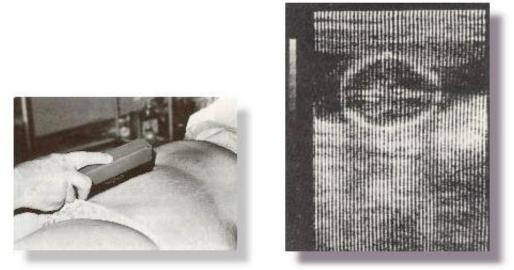


Toshiba SAL-20A marketed in 1978 Linear array and annular array technology had also been heavily investigated by the Japanese since the early 1970's. The country had been moving ahead very successfully with innovative electronic engineering in many domestic, commercial and professional sectors. Commercial linear array models from companies like Hitachi®, Toshiba®, and Aloka® soon began to dominate the world market. Hitachi® also produced their first linear array scanner the EUB-10 in 1976, followed by the EUB-20 in 1977 and the EUB-22 in 1979. The EUB-20Z produced in 1978 already incorporated the world's first digital scan converter.



Kazuhiro linuma

The Toshiba® SAL-10A and the more portable SAL-20A (pictured on the left) and <u>SAL-30A</u>, which were marketed in **1977**, **1978** and **1979** respectively, and the Aloka® <u>SSD-202</u> (1979), <u>SSD-203</u> (1980), SSD-240 (1981) and <u>SSD-256</u> (1982) were popular and had found their way into notable Institutions outside of Japan such as the King's College Hospital in London (Campbell), the Herlev (Gentofte) Hospital in Copenhagen (Holm), the Hospital Universitaire Brugman in Belgium (Levi) and were employed in many important early studies. The SAL-10A which was designed by acclaimed Japanese engineer Kazuhiro linuma, received many commendations. Other popular early choices included the Axiscan 5 (1976) and Abdoscan 5 (1979) from Roche Kontron®, the Sono R from Philips® (1978), the RA-1 (1980) and the Imager 2300 (1981) from Siemens®, and the LS 1500 (1981) from Picker/ Hitachi®. Aloka® scanners were marketed in the United States under the brand Narco Air-Shield®. Diagnostic Sonar® Ltd, a company founded in 1975 in Edinburgh, Scotland produced the first electronic real-time scanner, the System 85 in the United Kingdom in 1976.



Early scanner probe was bulky to fit on the abdomen *** Images from early real-time scanners had obstrusive scan lines, low dynamic range and resolution.

Many of the early models typically had very large probes housing an array of some 64 transducer (crystal) elements arranged in a linear row, and operating with sequential electronic switching or dynamic focusing. It was not until the early 1980s that probe size had gotten smaller and image resolution significantly improved.

Read the short History of the development of Medical Ultrasonics in Japan for a chronology of Japanese contributions to the development of ultrasound scanners.

At around the same time, steered-beam phased array transducers and annular array transducers with more complicated electronic circuitry were described, and had found their way into echocardiographic examinations because of the relatively small contact surface.



The phased-array scanning mechanism was first described by Jan C Somer at the University of Limberg in the Netherlands and in use from 1968, way





Jan C Somer

ahead of its time and several years before the appearance of linear-arrays systems. The principle of phased-arrays had probably been known much earlier where the technique was engaged in underwater submarine warfare and hence the technology was kept confidential. <u>Fredrick (Fritz) Thurstone</u> and <u>Olaf von Ramm</u> at the **Duke University** published one of the earliest and most significant phased-array designs in **1974**, which was incorporated into a number of commercial sector-array scanners. Very sharp focusing over an extended range was obtained from annular arrays using focusing methods on both transmit and receive. Other early significant contributors to the **beam-former techniques** included <u>Albert Macovski</u> at Standford University and



Frederick Thurstone

<u>Samuel Maslak</u> at Hewlett Packard®. Maslak later founded the <u>Acuson Corporation</u> (see also Part 3).

The <u>Kossoff</u> group in **Australia** had also made significant progress in the **annular phased array** transducer designs as early as **1973** and the technology was incoporated into their water-bath scanner, the **UI Octoson**. In England, **EMI**® produced the **Emisonic 4500**, a phased-array sector scanner which was nevertheless expensive, electronically noisy and had inferior resolution in the near fields. Early phased-arrays in the late 70s were all used in cardiac applications. Important manufacturers included **Varian**® and **Irex**®. In the first half of the 1980s, image quality in phased-arrays had continued to improve and some outstanding designs had come from **Irex**® and later on **Elscint**® (**Dynex**) and **Hewlett Packard**®. Despite the small probe size, phased-array sector scanners had never been popular with Obstetrical and Gynecological examinations.



The Digital Imager II from Picker, one of the newer static scanners in the early 1980s **Compound static scanners** continued it's tradition of being very huge bulky machines, probably influenced by the design norms of other imaging modalities such as tomographic x-ray machines and the bulky digital electronics housed in the console (see above on scan converters) before the impact of the micro-processors. <u>New static scanners</u> which were in great demand and produced excellent images were still on the drawing board and production line in the early 1980s. It was believed that real-time scanners would play only a **complimentary role** to static scanners in the assessment of moving structures. These static machines however were <u>starting to be</u> <u>replaced</u> or phased out at a rate that was **faster than expected**. There was apparently little practical, economical or clinical advantage of these costly machines over the more mobile and flexible electronic real-time scanners.

There were initially many who were so used to and skillful at operating the static machines that they were unhappy to switch over entirely to the real-time counterparts. They were also anxious about the latter's limited field of view, poorer resolution and allegedly 'less accurate' on-screen measuring system that they have only started to get used to not too long ago. Static scanners were not completely out of the scene until about **1985-86**. The switch-over

had serious financial implications to some companies who had a large inventory of static scanners.

Read a <u>commentary</u> by Royal J Bartrum, Dartmouth Medical School, in the book "Real-time ultrasonography" in 1982, on the switch-over to the real-time scanner.

From Digital's PDP-11 to Intel's 8080 and beyond

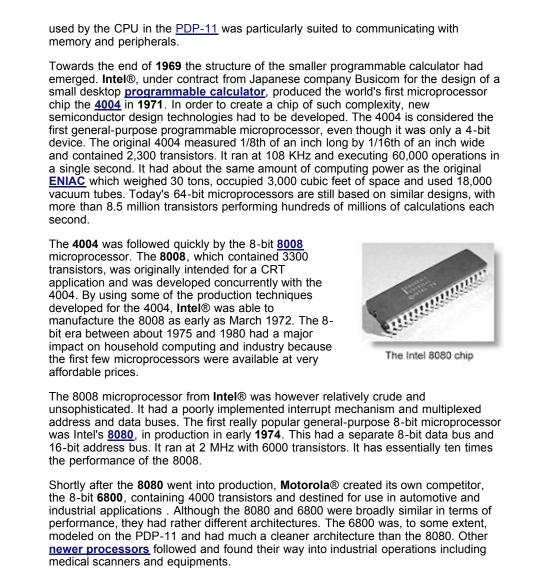
The rapid reduction in the physical size of the machine console in the later half of the 1970s (See Aloka® and Toshiba®'s early products above) was the direct result of the invention of the microprocessor and the evolution of the **minicomputer** into the **microcomputer**.



The DEC (Digital) PDP-11

By the late 1960's, computers built from discrete <u>transistors</u> and simple **integrated circuits** (IC) already existed. The first practical IC was fabricated in 1959 at Fairchild and Texas Instruments and Fairchild began its commercial manufacture in 1961. As manufacturing technology evolved, more and more transistors were put onto a single silicon chip with the maximum number of transistors per chip doubling every year between 1961 and 1971. They progressively became a device containing many circuits and was called a LSI (Large Scale

Integration). The <u>PDP-11 minicomputer</u> from DEC® containing many ICs and LSIs, was used in many scanner consoles up to the late 1970s. The UNIBUS architecture



Scanner engineering itself was soon in the hands of commercial companies rather than clinical personnel as advanced computer technologies were fiercely incoporated into each design to manipulate beam characteristics and signal processing to produce the best possible scan images. Apart from those mentioned above, other important early manufacturers of real-time equipments included Shimadsu® from Japan; EMI®, KretzTechnik AG, Bruel and Kjaer®, GEC® and Rohe® from Europe and Diasonics®, Dynex®, Ecoscan®, Elscint®, Hewlett-Packard®, Irex®, SKI®, Phosonic Searle®, Technicare® (acquired Unirad®) and Xonics® from the United States. The application of ultrasound in Obstetrics and Gynecology had since then undergone an explosive proliferation all over the world. By the early 1980s there were over 45 large and small ultrasonic scanner manufacturers worldwide.

Further Improvement in performance aside from focusing the ultrasound beam was acheived largely through an increase in the number of transducer crystals (or channels, from 64 to 128), improvements in transducer crystal technology (going into broad-band and high dynamic range), increasing array aperture (more crystals firing in a single time-frame), faster computational capabilities, improving technical agorithms for focusing on receive (increasing the number of focal zones along the beam), incoporating automatic time-gain controls and progressively replacing analog portions of the signal path to digital. It was perhaps regretable to see that **British manufacturers** have failed to keep up with developments made by other leaders in array technology, notably those from the **United States** and **Japan**. This was probably reflection of a similar trend in other arenas of electronic and micro-processor development in these countries. It is also of interest to note that the **Siemens Vidoson**® and the **Octoson**®from Australia both did not sell in North America at all. Both had the disadvantage of being **cumbersome** when scanners from other manufacturers were rapidly getting better in resolution and manuevability.



In the early 1980s (around 1980-1985), many agreed that mechanical sector scanners (be it rotating, oscillating or wobbling designs) which employed relatively large area transducers produced better and less noisy images than electronic linear-array scanners. Shown here are very good images from **SKI**® (left), **Diasonics**® (middle) and **ATL**® (right) taken in 1981. The market in Obstetrics and Gynecology was divided between the mechanical sector scanners and the linear-arrays until the second half of the 1980s where both were replaced by **convex** sector-arrays.



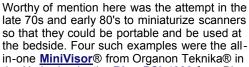
Hitachi's convex array in 1984

Because of its smaller convex contact surface, the **curvilinear** or **convex sector-array** fits much better on the abdomen and allows for a wider field of view than does the linear-array configuration. Work on the fabrication of an electronic **convex** array had started in the **late 1970s** in the larger Japanese companies such as **Hitachi®** (publishing their convex attachment to their **EUB-10A** scanner in 1978); **Aloka®** (filing their <u>patent</u> on the convex scanner in 1980), as well as in U. S. companies notably the **North American Philips®** and the <u>**Picker Corporation**®</u>, who had filed their <u>patents</u> for convex arrays and processors in **1979** and **1980** respectively. The first commercially available **convex array transducer** apparently only debuted in 1983 in a scanner from **Kontron Instruments®** in Europe, the <u>**Sigma 20**</u>, which was designed especially for use in Obstetrics and Gynecology. **Hitachi®** in Japan marketed their model <u>**EUB-40**</u> with their new convex array later on in the same year.

Toshiba® introduced a similar array in **1985**, in their new scanner model <u>SAL-</u><u>77A</u>. Interestingly, the design actually replaced an earlier model (by only about 9 months) the <u>SAL-90A</u> which boasted a new "trapezoid" linear array in which the face of the transducer was flat but a trapezoid-shaped image was produced from the 128 transducer elements using phased electronics. American machines were apparently still using linear arrays by 1985, although very shortly they were quickly replaced by the new convex configuration. By about 1987, convex arrays are standard on every new scanner, whether or not it is configured for use in Obstetrics and Gynecology.

Model number of some of the scanners made after 1980 from important manufacturers are listed <u>here</u> with the year in which they were marketed. Also <u>view pictures of some of the early scanners</u>.

Skin Coupling material for ultrasonic transmission has also switched from oil to a watersoluble (non cloth-staining) gel medium. One of the more well-known manufacturers was the <u>Parker Laboratories</u>® at New Jersey. Images are commonly recorded on "peelapart" <u>Polaroid® films</u> (the <u>Type 611</u> was most commonly used). **Multi-format** radiographic films (6-9 images on one film) using dedicated video imagers soon became mainstream with institutional users and **thermal paper printers** in the private practice market.





Polaroid type 611 "peel-apart" B&W film

the Netherlands, the <u>Bion PSI-4000</u> from Bion Coporation® in Denver, the <u>Shimasonic SDL-30</u> from Shimadzu Corporation®, Japan, and the <u>210DX</u> from Aloka®.

The <u>Minivisor</u>® (available from 1979) was a spin-off from <u>Bom</u>'s laboratory. It was battery operated, shaped like a mushroom, had no wires and used a 2-inches display with an on-screen caliper system and digital readout. The transducer is fused to the bottom of the device similar to a 'large' fetal pulse detector. <u>Juiry Wladimiroff</u> suggested in **1980** the device would be useful for routine BPD screening. Nevertheless the popularity of these machines were short-lived for several important reasons pertaining at that time: The resolution

was unsatisfactory because of the available electronics. The images of the 'standard' and larger devices, as well as their

The MiniVisor ****

overall 'portability' have seen rapid improvements round about the same time; and thirdly, real-time ultrasound has very rapidly established itself as a definitive diagnostic entity and the concern for good image information appeared to overide that of the extra portability.

The invention of the real-time scanner had enabled much more effective diagnosis of many fetal malformations and in particular cardiac anomalies which hitherto was impossible to diagnose accurately. (see Part 3). Fetal sonography and prenatal diagnosis (a term which was only coined in the 1970s) had emerged as the 'new' science in Obstetrics and fetal medicine.

<u>John C. Hobbins</u> at the Yale University, Connecticut and <u>Stuart Campbell</u> at the King's College Hospital in London were among others, the two most important forerunners on either side of the Atlantic. Their centers have also become two of the most important teaching centers in fetal sonography. Many of the research fellows and staff members that had come through Hobbins' department for example, have in time become celebrated names in the field of fetal sonography and prenatal diagnosis.



femur length measurements only became possible with the advent of real-time scanners

Other important early North American workers in fetal biometry included Peter Cooperberg, David Graham, Charles Hohler, Alfred Kurtz, Rudy Sabbagha and Roger Sanders. Their early work on the biparietal diameter was particularly notable, establishing charts for different populations, standardizing measuring methods and errors and comparing differences that



John C. Hobbins

may be present between measurements made on static and realtime equipments. (see also Part 3). The real-time scanner had soon enabled the accurate measurement of fetal limb bones that lead to the introduction of the important measurement of the fetal femur length by John Hobbins in 1979 for the evaluation of skeletal dysplasia followed by Gregory O'Brien and John Queenan who described it's use in fetal growth assessment. Phillipe Jeanty at Yale provided in 1984 measurement charts for all the fetal long bones.

Ultrasound guidance was started to be employed in procedures such as amniocentesis (Jens Bang and Allen Northeved 1972, Copenhagen), fetoscopy (John Hobbins and Maurice Mahoney, 1974) and transabdominal chorionic villus sampling (Steen Smidt-Jensen and N Hahnemann, 1984, Copenhagen).

Read here a short history of Amniocentesis. fetoscopy and chorionic villus sampling.

Transvaginal scanners



Vaginal A-scan from KretzTechnik circa 1968

Wild and Reid had invented and described the use of A-mode transvaginal and trans-rectal scanning transducers as early as 1955 (see Part 1). Using a proprietary A-mode vaginal scanner from KretzTechnik in the early 60s, Alfred Kratochwil in Austria had reported on fetal heart pulsation at slightly over 6 weeks menstrual age. He also developed a thimble attachment transducer to facilitate vaginal sonography with pelvic examination. Trans-vaginal sonograms were also reported by Lajos von Micsky in New York around the same time, using innovative prototype equipment that he devised. Other <u>A-mode vaginal transducers</u> were deviced by Japanese researchers. With the advent of B-mode equipments, A-mode vaginal scanners had slowly disappeared from the market.

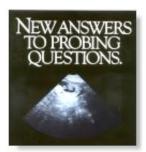
Shigemitsu Mizuno in Japan demonstrated in 1965 an endovaginal scanner for pelvic examination using a plan-position indication (PPI) B-mode format (the same format as an abdominal compound contact scanner). The device was mannually rotated and the resulting display was very similar to a circular military 'Radar" display. Used either transrectally or transvaginally, It was capable of producing some meaningful pictures of the pelvic organs. Salvator Levi at Brussels was one of the early advocates of Aloka® equipments and published on their transvaginal scanner as early as 1970. R A Khentov in the USSR and von Micsky in the United States also demonstrated vaginal and rectal scanning devices in the late 1960s.

Despite the advantages of transvaginal scans, its use had waited almost 20 years to become practical and popular in gynecological scanning. Athough the need and technology were there, the really practical real-time transvaginal scanning probe was not "invented" until **1985** when KretzTechnik® of Austria produced their first real-time mechanical vaginal sector scanner. The transducer had a scan angle of 240 degrees and was designed with the use for transvaginal ovum retrieval in mind, in collaborations with IVF pioneers Wilfried Feichtinger and Peter Kemeter in Austria. The coming of such mechanical devices was also in part the outcome of advances in microprocessor controller circuitry and ferro-magnetics resulting in much smaller and more efficient motors.



The first commercially available transvaginal transducer

IOTH MELZ LECTICIK IT 1900



advertisement of the vaginal probe from Philips

Dutch manufacturer **Philips**® followed on with <u>one of</u> <u>the earliest</u> mechanical vaginal scanners in the

second half of **1986**. The probe was in the shape of a microphone with a roundish elongated head housing a 5MHz 13mm wobbler transducer. It could be retro-fitted onto their real-time scanner <u>SDR 1550</u> which first debuted in 1985. Although they produced excellent images compared to their abdominal counterpart, mechanical endovaginal designs were not favored by many ultrasound manufacturers, partly because of the vibration that was generated.

Mechanical designs were rapidly followed by **electronic array versions** which are rather like a reduced-size abdominal **convex sector transducer** that has appeared around the same time from other manufacturers in Japan. **ALOKA**® produced an electronic sector version which could be retofitted onto their older model the <u>SSD-256</u>. **GE Medical Systems**® produced their first endovaginal probe to fit their <u>RT3200</u> in **mid 1987**. By **1988**

most manufacturers had endovaginal options installed in their scanners.



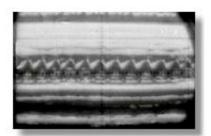
B-Joachim Hackeloer

The advent of tranvaginal scanning (at higher frequencies of 5 - 8 MHz and resulting in much finer resolution) had a significant impact on the diagnosis of **gynecological** and **early pregnancy** pathologies. In particular the accurate recognition of fetal cardiac pulsations in **missed abortions** was facilitated at an early gestational age of 6 weeks.

Non-palpable ovarian masses were often picked up by a vaginal scan. Transvaginal scanning has also progressively become standard practice in the management of **infertility** patients, in assessing follicular development and endometrial growth, as well as being an indispensible aid in the retrieval of oozytes from the ovaries. The assessment of **ovarian follicular development** had before that been based on static and real-time abdominal ultrasound, first popularised by **B-Joachim Hackelöer** and his group in Germany since 1977. (See also **Part 3**). And before that time, most clinicians did not envisage that cyclical changes in morphology and size of ovarian follices can be so closely followed with ultrasonography.

M-mode and Doppler

The M-mode (time-motion) display was first described by <u>Inge Edler</u> and <u>Hellmuth Hertz</u> in Lund, Sweden in 1954 using a modified metal-flaw detector from Siemens® of Germany. They demonstrated the feasibility of recording cardiac valvular motion ultrasonically. Sven Effert in Germany, who had been collaborating with Hertz in some of his work, futher demonstrated the usefulness of M-mode echocardiography, which had subsequently caught on as a mainstay investigation in cardiology. <u>Xin-Fang Wang</u> first described in China in 1964 the use of M-mode ultrasound in the study of <u>fetal</u> cardiac movements. Jens Bang and <u>Hans Henrik Holm</u> demonstrated fetal cardiac motion using M-mode from 10 weeks onward in 1968. In the same year <u>Alfred Kratochwil</u> described similar usefulness of detecting fetal cardiac motion by M-mode in patients with threatened abortion. These were nevetheless 'blind' procedures. <u>Hugh Robinson</u> in Glasgow described with great success in 1972 the detection



Early M-mode tracings are depicted on oscilloscope screens

of fetal cardiac motion at 7 weeks by A- and M-mode after locating the fetus with on B-mode ultrasound. In the same year **Fred Winsberg** in Canada described **M-mode diagnosis** of fetal cardiac anomaly.

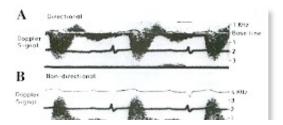


The <u>Doppler principle</u> was first described over 100 years ago by <u>Christian Andreas Doppler</u> in Austria in **1842**. Medical applications of the <u>Ultrasonic doppler techniques</u> were first implemented by <u>Shigeo Satomura</u> and Yasuhara Nimura at the Institute of Scientific and Industrial Research in Osaka, Japan in **1955** for the study of cardiac valvular motion and pulsations of peripheral blood vessels.

The **Satomura** team which also included <u>Ziro Kaneko</u>, pioneered <u>transcutaneous doppler flow</u> <u>measurements</u> in **1959**, several years ahead of the work at the **University of Washington** in Seattle.

ura *^* It has also become known from the work of Ziro kaneko and Kanemasa Kato in

Satomura's laboratory in **1962** that **blood flow** can be detected by the ultrasonic doppler method largely because blood itself consists of a suspension of "**uncorrelated point back-scatterers**" (the red cells) with varying density and compressibility, and not because of turbulence in the flow stream as was previously thought. In **1966**, **Kato** and **T Izumi** pioneered the <u>directional flow-meter</u> using the local oscillation method where **flow directions** were detected and



Base Line

displayed. This was a breakthrough in doppler instrumentation as **reverse flow** in blood vessels could then be documented.

Compared to the western world, throughout the history of doppler ultrasound, **Japanese** research had played a leading and major role in its development. Much of the ground-breaking research

Spectral and bi-directional flow demonstrated by Kato and Izumi in 1966. A: directional flow. B: non-directional

were presented in national engineering journals and at the annual scientific meeting of the Japan Society of Ultrasonics in Medicine (in japanese) and were thus unknown to the west until a later date.

A brief history of the pioneering work in **Doppler applications** in **Japan** is described <u>here</u>.



Donald Baker

following year.

In the United States, ground breaking work in Doppler instrumentation started at the University of Washington in Seattle from 1958 onwards led by <u>Robert Rushmer</u>, a pediatrician and physiologist, who was determined to characterise cardiovascular functions in intact unanaesthetised animals. He was soon joined by **Dean Franklin**, **Dick Ellis** and <u>Donald Baker</u>, engineers who were recruited into Rushmer's cardiovascular instrumentation development program. The team pioneered **transcutaneous continuous-wave** flow measurements and <u>spectral analysis</u> in 1963 and as a clinical application, surgeon <u>D Eugene Strandness</u> undertook the clinical testing. He published the first transcutaneous vascular spectral flow signals in the



Donald Baker infront of an early doppler device ****

Although continuous wave doppler instruments were quite in place by the mid 1960s, for the next 10 years or so it's use in **Obstetrics and Gynecology** was largely confined to that of a fetal pulse detector and continuous fetal heart rate monitoring (see below). Gynecologist **Wayne Johnson** working with <u>Rushmer</u> reported the detection of fetal life at 12 weeks gestation (from LMP) with **Strandness'** continuous wave instruments in **1965** (see also below). The continuous wave technology was transferred to **Smith Kline Instruments**® from the University of Washington and the company manufactured the first <u>Doptone</u> in 1965. Other early commercial suppliers of continuous wave doppler devices in vascular applications included **Parks® Electronics Inc.**, **Kay® Electric company** (the SonaGraph), and **Medasonics® Inc** (the Versatone). An audio transcript of "*Typical Doppler Sounds from peripheral arteries and veins*" prepared by D Eugene Strandness in the late 1970s can be heard <u>here</u>.

The continuous wave Doppler method did not provide explicit information about the distance between the ultrasonic transducer and the moving target. Further development led to the introduction of **pulsed-doppler** system by **Baker** in **1970**, a concept based on the repetitive propagation of short ultrasound bursts and analysis of the signal received at a preselected time delay with respect to emission (the sample volume). **Baker** also outlined a technique for determining the volume blood flow from Doppler velocity measurements. The team also recruited **John Reid** from **Pennsylvania** who complemented the flow designs with 2D and M-mode technology. New instruments basing on the ultrasound doppler principles were developed. The first **duplex pulsed-doppler** scanner, a milestone development in ultrasound instrumentations was designed and developed by **Frank Barber**, **Baker**, **Reid** and other colleagues in **1974**. The duplex scanner finally enabled 2D gray scale imaging to be used to guide the placement of the ultrasonic beam for doppler signal acquisition.



Third version of the duplex scanhead 1976. The plunger on the right moves in and out to adjust the range-gating

"...... The program was instrument and hardware orientated, because in those days, medical research was carried out in little the same way as wars are fought. Wars are fought according to the weapons you have, and the rules of the game will be according to the weapons you have. I think a lot of researches are in the same way. If you have the tools that nobody else has you can create a new game that nobody else can play...... People saw this as an opportunity. They saw an opportunity to get ahead of their colleagues and publish" --- Donald Baker, speaking on his early work in doppler instrumentations and his efforts in popularising them at ATL.

ATL® (Advanced Technology Laboratories) was founded in 1969 near Seattle, Washington, by a small group of engineers developing marine electronic systems. Starting form 1973, technology developed at the University of Washington Center for Bioengineering were <u>transferred to ATL</u> in the development of systems for diagnostic sonography. The first duplex pulsed-doppler scanner, the Mark1 (with pulsed-doppler module model 400) was released in 1974 for cardiac investigation. The later version 400B became part of the well-known ATL Mark V duplex scanner which debuted in 1978. Squibb Coporation® acquired ATL in 1980 and ADR in 1982. ADR was merged into ATL in 1984.



In England, pioneering work from <u>Peter Wells</u> in **Bristol** also demonstrated in **1969** that pulsed-doppler information may be obtained through the combination of range-gating methodogy and continuous wave and 2D techniques. In the late 1960s the **Okujima** and **Ohtsuki** group and the <u>Ziro Kaneko</u> group in Japan (after the death of **Shigeo Satomura**), and the **Pierre Peronneau** group in **Paris** were equally active



Early doppler device from England

in pioneering pulsed-doppler devices. The development of <u>pulsed-</u> <u>doppler instruments</u> by these groups enabled, for the first time, noninvasive localised measurements of blood velocity. In addition, their instrument could acquire velocity information at several positions along a vessel's diameter, thus enabling the velocity profile of the blood to be visualised. On the clinical side, various waveform **indices** were devised in doppler investigations to reflect flow inadequacies. <u>Leandre Pourcelot</u> introduced the '**resistance index'** in Tours, France in 1974. In the same year **R Gosling** and **D King** introduced the '**pulsatility index'**. The **J Drumm** group in Dublin, Ireland described the use of the **A/ B ratio** in 1980.

In the late 1970s, aside from the Seattle group, early duplex doppler devices were also researched and fabricated by the **T Ogawa** group in **Japan** (phased-array duplex sector, 1977), the **James Griffith** and **Walter Henry** group in the United States (phased-array duplex sector, 1978) and the <u>Bjørn A. Angelsen</u> group at **Trondheim**, Norway (combination duplex linear arrays, 1981).

Doppler development in Obstetrics and Gynecology



The Doptone from SKI

The use of spectral flow analysis on the **fetus and placenta** was a remarkable *late-comer*. The detection of fetal pulsations using doppler ultrasound was first reported in **1964** by **D A Callagan** who was working with ultrasonic devices at the United States Naval medical Research Institute at Bethesda, Maryland. In the following year (1965), Gynecologist **Wayne Johnson** working with the **Rushmer** team at the University of Washington (see above) reported 100% accuracy in the detection of fetal life in 25 patients at **12 weeks** (from LMP). **Smith Kline Instruments**® manufactured the first <u>Doptone</u> in the same year basing on their technology.

Edward Bishop at the University of Pennsylvania, using the SKI® <u>Doptone</u>, reported positive doppler

signals from **11 weeks** (LMP) pregnancies in **1966** and 65% success in locating the **site of the placenta** basing on audio doppler signals in the third trimester. The **Callagan** group in the same year reported doppler interrogation of the **fetal heart** and described "*the sound of horse's hoofs when running*' and oscilloscopic 'beats' of the **cardiac** doppler signals. **John Barton** at the Northwestern University in Chicago further reported in 1967 positive audio doppler signals at **10 weeks**. In 1968, the **Johnson** group in Seattle expanded on the usefulness of the doppler flowmeter (audio signals) to the **localization of the placenta**, demonstrating the characteristic '*whirlwind*' and '*rushing wind*' sound of placental blood flow.

In **1967**, the <u>Rushmer</u> group outlined the use of <u>doppler ultrasound in Obstetrics</u> in an article in JAMA, "*Clinical applications of a Transcutaneous Ultrasonic Flow Detector*", which was confined basically to the detection of fetal life, placental location, blood flow through the uterine vasculature and fetal movements. Aside from the **SKI®** <u>Doptone</u>, other similar devices marketed at that time included the **Ames® Ultradop** and the **Magnaflux® MD 500**. Other companies producing doppler fetal pulse detectors included Parks Medical Electronics, Imex, Medasonics, Sonicaid and Life Sciences. The **Doptone** was considered as one of the most important instruments that was ever invented in Obstetrics. Before that great difficulty was often encountered in detecting fetal life in both early and late gestations. This now seemingly trivial instrument has quite considerably changed the practice of Obstetrics since the 1960s.

While the use of vascular '<u>spectral</u> <u>flowmetry</u>' was quite in place by the mid-1960s (see above), the use of doppler flowmetry in pregnancy assessment was not followed up in the



American or European literature until about 1977 when 3 separate

Umbilical aterial doppler described by Takemura et al in 1968

groups of investigators were making important pioneering contributions.

The study of flow velocity waveforms in fetal and placental blood vessels have nevertheless been reported by Japanese reseachers as early as 1968. In that year H Takemura and Y Ashitaka described umbilical arterial and placental doppler spectral velocity waveforms at the 14th meeting of the Japan Society of Ultrasonics and Medicine. Although the clinical significance of these signals were not quite known, their devices had allowed them to publish very clear and remarkable signal traces resembling those produced on recent equipment (see above right).

In 1977, J E Drumm, a gynecologist and D E FitzGerald, then director of the Angiology Research Group of the Irish Foundation for Human Development in Dublin, Ireland, reported in the British Medical Journal the combined use of continuouswave doppler and 2-D static B-mode ultrasound in the study of flow velocity waveforms in the fetal umbilical artery, and described probably the first umbilical arterial velocity waveforms in the western literature. The shape and applications of these waveforms were not discussed, although they suggested that "... the shape of the blood-velocity waveforms will change with conditions affecting the efficiency of blood-supply, and the method should be useful in assessing conditons such as preeclampsia and intrauterine growth retardation ". The same group reported in 1980 in a much longer paper umbilical waveforms in relation to gestational age, their analysis in terms of systolic and diastolic components, and the use of various waveform ratios.



W D McCallum

Another pioneering group consisting of researchers from Stanford University, the University of Washington and the Varian associates®, described in 1978 fetal flow velocity waveforms basing on pulsed-wave (range-gated) doppler instruments to overcome the short-coming of the continuous wave counterpart. William McCallum, a gynecologist at Stanford (who immigrated from Belfast, Ireland in 1975) together with his co-workers devised sophisticated computer-based techniques to process the doppler signals through a series of fast Fourier transformations instead of the older and more primitive method of 'time-interval histogram'. Although they commented that no general conclusions could be reached

on the clinical significance of these waveforms from the small study, the group had firmly established the basic groundwork for further investigations.

Robert Gill, together with the Kossoff group in Sydney, Australia made quantitative measurements of human blood flow velocities since 1977 with the Octoson®. He determined that the flow in the fetal umbilical vein increased with gestational age but remains constant at around 103 ml per min per kg fetal weight. Accurate measurement of flow volumes and flow velocities in the fetal blood vessels was however affected by a diversity of factors such as operator skill, fetal position, blood vessel diameter and angle of insonation which made it an impractical investigation in the fetus. Sturla Eik-Nes, working with Karel Marsal in Norway, devised the first



Sturla Eik-Nes

1

hand-held linear-array real-time apparatus coupled with range-gated doppler in 1980. The group in that year documented blood flow velocities in the fetal aorta and reported in 1983 volume flow through the umbilical vein using the new apparatus.



Early doppler work also came from the Pentti Jouppila and Pertti Kirkinen group at Oulu, Finland who also worked with quantitative blood flow velocities in the umbilical vein and found significant reduction in flow in growth-retarded fetuses. Their work importantly demonstrated (in 1981) that guantitative umbilical venous flow became unrecordable in fetuses with severe growth-retardation and (in 1984) a significant negative correlation existed between umbilical venous flow and the cord haemoglobin.

Pentti Joupilla

Although a relationship could be established between low flow volumes and fetal compromise, such measurements were

difficult to be made accurately and thus had not become practical and popular as a fetal investigation. Measurement of volume and velocity flow in fetal blood vessels were not further pursued as a research and clinical tool after the mid-1980s.

Stuart Campbell and David Griffin at the King's College

Hospital in London suggested in **1983** that the shape of the arterial flow velocity waveforms would be more useful in fetal assessment. In the same year **Campbell** also reported on the usefullness of uterine and placental arcuate arterial waveforms, particularly in conditions such as pre-eclampsia.

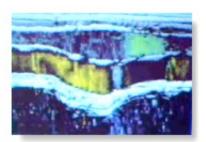
With the efforts of **WB Giles** and <u>Brian J Trudinger</u>, the Australian group made significant contributions to the study of **velocity waveforms** and had made popular the measurement of <u>waveform ratios</u> in the assessment of fetal well-being. Such ratios soon became standard in the assessment of fetal



Brian Trudinger

circulation in utero. The **Trudinger** group demonstrated in **1986** that abnormal doppler waveform tended to preceed abnormal fetal heart traces. The mid 1980s saw many other centers starting their investigations into doppler velocimetry, often on shared equipment with their cardiology collegues. The significance of a diminishing diastolic flow in the face of fetal compromise was most certainly established. It wasn't until the **late 1980s** that clinical applications of doppler arterial blood flow started to become an important and integral part in fetal assessment. (To continue in **Part 3**)

Further development led to **2D color flow imaging**. **Marco Brandestini** and his team at the University of Washington in **1975** obtained blood-flow images using a 128-point multi-gated pulsed doppler system, where velocity waveforms and flow images were encoded in color and superimposed on M-mode and gray scale 2-D anatomical images. The team included physician **Geoffrey Stevenson** and engineers **Mark Eyer** and **David Philips**, who brought in new technology on **scan-converters** from the **Thurstone** group at **Duke University**. They were able to demonstrate the value of color flow imaging in the diagnosis of various **cardiac defects**. <u>Leandre Pourcelot</u> in Tours, France also described their first color-coded doppler images in **1977**. Color doppler systems in the late 1970s and early 80s were however limited by the processing power of the equipment, the lack of good duplex arrays (as contrasted to the mechanical rotor systems) and the agorithm and technique with which doppler frequency estimation was performed.



2-D color flow image of the carotid and jugular vein (top) from the Brandestini group in 1975



The first real time color flow Doppler machine from Aloka, Japan (SSD-880)

It was not until the work of **Chihiro Kasai**, **Koroku Namekawa**, **Ryozo Omoto** and coworkers in **Tokyo**, **Japan**, which was published in the English language in **1985**, that led to the widespread realization that **real-time color flow imaging** could be a practical possibility. The group had already reported on the technical details and clinical (cardiac) applications in the Japanese language in **1983**. **Namekawa** and **Kasai** were bio-engineers working for **Aloka**® and **Omoto** was cardiologist at the **Saitama Medical School** in Tokyo which had a long history of collaboration with **Aloka**®. What is 'required' to produce a **flow image** of blood vessels is that the **amplitude**, **phase** and **frequency** contents of the returned echoes from a single linear array probe are captured and very rapidly analysed. The Japanese group used a phase detector based on an <u>autocorrelation technique</u> in which the changing phase of the received signal gave information about changing velocity along the ultrasonic beam. This provided a rapid means of frequency estimation to be performed in real-time. This approach to **color flow mapping** is still in use today.

Continuing advances in electronics have permitted the development of faster color Doppler instruments, which displayed two-dimensional blood velocity measurements at many frames per second. The successful introduction of color Doppler required fast and stable positioning of the ultrasound beam, which was provided by the development of the linear array and phased-array scanheads. A breakthrough filtering mechanism was also

deployed in which the high amplitude/ low velocity clutter signals generated by the movements of tissue structures and vessel wall are removed. Such filters were described by <u>Bjørn A.</u> <u>Angelsen</u> and K Kristofferson at the Department of Physiology and Biomedical engineering at Trondheim, Norway in 1979 on the analysis of moving targets in radar systems. In the early 80's the Trondheim group published important work on annular array color flow imaging technology. Color flow mapping not only allowed elucidation of blood flow but had also helped in the determination of pathology in tissues.



Bjorn Angelson

The Company Vingmed Sound AS® which manufactured some of the earliest doppler applications was formed in 1974 with technology transfers from Angelsen's Department. The company's doppler equipments performed very well and had a sizeable market in the United States, teaming up with companies like Irex®, Ausonics® and Interspec® in cardiac applications. Vingmed® was later acquired by GE Medical Systems®.

Advertisements of the first machine with real-time Color flow mapping capabilities from Aloka® (the SSD-880CW) made it's debut in medical journals in the middle of 1985 (It was produced in Japan in 1984). Toshiba® followed up with their

SSH-65A later on in the same year. Asim Kurjak in Croatia, using the Aloka® machine was the earliest pioneer to introduce the application of color flow doppler in fetal assessment, publishing his work in 1987. (See also Part 3).



Quantum's first ad in 1986

Quantum Medical Systems®, Issaquah, Washington, a new company formed by a group of engineers who left ATL, had started introducing the concept of real-time color doppler imaging at the AIUM meeting in the fall of 1983. The first color images were shown at the RSNA (Radiological Society of North America) meeting in December 1984. The prototype machine was tried out at several centers and one of the first papers was reported in the **October 1985** meeting of the RSNA by **Christopher Merritt**'s group at Tulane University, New Orleans. Quantum® marketed their first machine, the QAD-1 in 1986 which produced some very impressive real-time color flow images of the carotid and other arteries basing on the newer array technology. They called it "AngioDynography" although the term had not subsequently become popular. The transducer design and signal processing were described by Alfred Persson and Raymond Powis, one of Quantum's founders in their article "Recent advances in imaging and evaluation of blood flow using ultrasound" in 1986.

Color flow imaging made it's real impact in the United States in 1987 and in Europe in the

following year. <u>ATL</u>® after some re-organization, marketed its first color doppler machine, the <u>UltraMark 9</u> in **1988**. **Irex**® did not make its own color doppler but marketed **Aloka**®'s SSD-880 in the united States instead. Irex®, being acquired by the Johnson & Johnson Co®., and sold later to GE®, discontinued its line of duplex scanners. Quantum Medical Systems® was later acquired by Siemens® of Germany, who moved their main operations to Issaquah, Washington.

It was not until the early 1990's that the modality found it's way into the assessment of Gynecological and early pregnancy abnormalities.

"Power doppler" or "Color power imaging" continued to develop in the 1990s. "Tissue doppler imaging" developed further from a revived concept with the arrival of better computational electronics. These developments had important clinical impact on the diagnosis of malignant conditions where tissue vascularity is increased and on moving structures other then blood flow (see Part 3).

Tissue Characterization

The ability to develop methods for signal processing of ultrasonic echoes from tissues which allow correlation of acoustic parameters with tissue constituents had always been on the minds of early ultrasound inventors and reseachers such as John Wild, Tom Brown and Alfred Kratochwil. However, tissue characterization never became a fruitious area in ultrasonic applications and had never caught on particularly in the field of Obstetrics and Gynecology. Texture characterization of human body tissues has never been easy to perform. Quantitative methods used included fractal models, multichannel methods, analysis of first and second-order statistics of texture, Gaussian Markov random fields, mean scatterer spacing and multiresolution analysis. Invented algorithms are complicated and often inefficient.

The success of tissue characterization in medical ultrasound, and particularly in Obstetrics and Gynecolgy, is largely hampered by the presence of the inhomogeneous soft tissue medium and inconsistency of position between the tissue of interest and the insonifying ultrasound transducer (ie. abdominal wall etc.). As the acoustic wavefront propagates through the inhomogeneous tissue, it is modified in amplitude and phase which often has a large and unpredictable effect on the output voltage of the receiving transducer. In addition, the tissue introduces attenuation, and multiple transmission paths through the soft tissue may create replicated echoes. Back-scattered signals are also degraded by speckle noise, acoustic shadowing, and system distortions present in all instrumentation. Important early researchers in this area included Christopher Hill at Sutton, England; Frederick Kremkau at Yale, John Reid at Drexel and the Kossoff group in Sydney. Up to this day, no effective technique for tissue characterization is in use in Obstetrics and Gynecology.



sister chromatid exchanges in a normal chromosome spread

The safety of diagnostic ultrasound has received rigorous scrutiny since it's inception, particularly when high power ultrasound had been used in the 1940s for destructive and therapeutic purposes. Subsequent to a number of early reports which did not demonstrate any harmful effects of ultrasound insonation on human cells, Ian Donald, in co-operation with Malcolm Ferguson-Smith, director of the cytogenetic laboratory at the Queen Mother's Hospital conducted extensive experiments in 1967 to delineate possible harmful effects on high intensity ultrasound on interphase and mitotic chromosomes and did not find any. Studies in 1963 at the Juntendo Ultrasound Reasearch Center in Japan also did not reveal any harmful effects on pregnant rats exposed at the maximum power of diagnostic equipments for 3 days after fertilization. Bertil Sunden in Lund, Sweden found no teratogenic effects in his thesis research in 1964. Other studies conducted in England by El Kohorn in 1967 and John C. Hobbins in the United States also did not show any appreciable cytological effects.

Important early researchers included Wesley Nyborg at the Pennsylvania State University and later at the University of Vermont; Paul Carson at the University of Colorado; Raymond Gramiak at the University of Rochester; William and Francis

Fry, Floyd Dunn and William O'Brien Jr. at the University of Illinois; Christopher Hill at Sutton, England; Marvin Ziskin in Philadelphilia; Masao Ide and Kazuo Maeda in Toyko; and Stanley Barnett and George Kossoff in Australia, to name just a few. Much was done on the study of in-vitro effects of ultrasonic insonation at various intensities, their effects on

heating, cavitaion and bubbles; their mechanisms of action, standardization of intensity measurements and drawing up various Guidelines on Biological Safety.

Researchers were also looking at immuno-suppresive effects, platelet aggregation, cell phagocytosis, DNA damage, fetal weight alterations and increase in sister chromatid exchanges in the presence of ultrasound insonation. LM Hellman's study in 1970 on 400 newborns insonated before birth also reveal no increase in abnormalities in the infants. Many other studies followed. A Bioeffects committee was set up in the AIUM which reviews and monitors the world literature on ultrasound bioeffects on a regular basis. Similar watch groups were set up by the EFSUMB and the WFUMB. A definite harmful effect at diagnostic levels of ultrasound cannot be confirmed. Such findings give further impertus to the rapid development of ultrasonography in Obstetrics, where concerns for bioeffects safety was obvious.

In the 1980s and 90s, new studies emerged looking at the long term effects in children exposed prenatally to ultrasound. No definite adverse effects could be determined in the long term. Other activities in the United States are described here. The Bioeffects Committee of the AIUM made the following statement in 1982 and again in 1997:

" No confirmed biological effects on patients or instrument operators caused by exposure at intensities typical of present diagnostic ultrasound instruments have ever been reported. Although the possibility exists that such biological effects may be identified in the future, current data indicate that the benefits to patients of the prudent use of diagnostic ultrasound outweigh the risks, if any, that may be present.

It was widely held that the clinical use of ultrasound over the years has not established any adverse effect arising from its exposure. While randomized clinical studies are the most rigorous method for assessing potential adverse effects of diagnostic ultrasound, studies using this methodology show no evidence of any effect on birthweight in humans. Other epidemiologic studies have also shown no causal association of diagnostic ultrasound with any of the adverse fetal outcomes studies (AIUM statements). The World Federation, European Federation and the Australasian Society also held similar views. These activities and statements played an important role in the proliferation of ultrasound use in Obstetrics as any slightest harmful effect would likely have constituted a major deterrant.

Read the article: "Assessing the Risks for Modern Diagnostic Ultrasound Imaging" by Dr William O'Brien, Jr. ^

An important article on the Historical Development of Bioeffects Assessment can be found <u>here</u>.

Read also the article: "History of the American Institute of Ultrasound in Medicine's efforts to keep Ultrasound safe" by Wesley Nyborg

...... Image quality of real-time ultrasound scanners made steady improvements during the mid 1980's to early 90's secondary to the increasing versatility and affordability in microprocessor technology. Nev ertheless it was not until the early to mid 1990's that more substantial enhancements in image quality were seen (see Part 3)

...... The use of ultrasonography continued to boom into the 1980s. According to statistics from the Bureau of Radiological Health Surveys (FDA), in the United States, the percentage of hospitals using ultrasound for dating increased from 35% in 1976 to 97% in 1982

The World Federation of Ultrasound in Medicine and Biology (WFUMB) 1988 Meeting in Washington was preceeded by a two day Symposium on the History of Ultrasound. This event was the culmination of the sustained efforts of the AIUM Archives Committee, chaired by Barry Goldberg. Of the 500 or so delegates about 200 were recognised as pioneers. During the awards ceremony a further 40 pioneers all over the world were recognised In Memorium. The British Medical Association (BMUS) held a similar parallel event in Glasgow in the same year. In both meetings there were exhibition of many historical ultrasound instruments

...... Stuart Campbell with his committee of international luminaries soon started the International Society of Ultrasound in Obstetrics and Gynecology (ISUOG) in 1990 and held it's first world congress in the following year. He also became the founding editor of the Society's official journal: Utrasound in Obstetrics and Gynecology



Go to [Part 3]

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Horace Thompson's image courtesy of the AIUM.

George Kossoff's picture courtesy of Professor Salvator Levi.

* Courtesy of <u>KretzTechnik</u>®, Zipf, Austria.

^{***} Scottish machine, images reproduced with permission from Dr. RG Law, from his book 'Ultrasound in Clinical Obstetrics', John Wright and Sons Ltd, Bristol, 1980. **** Image courtesy of <u>Dr. Eric Blackwell</u>, reproduced with permission.

^ From "Medical Diagnostic Ultrasound: A Retrospective on its 40th Anniversary", reproduced with permission from Dr. Barry Goldberg. A-scope image courtesy of the Department of Ultrasonics, Polish Academy of Science.

^ quoted from "Looking at the Unborn: Historical sapects of Obstetrical Ultrasound" - Wellcome Witnesses to Twentieth Century Medicine Vol 5, An See Craig M. Sonography: an occupational health hazard. JDMS 1985;Vol. 1 No. 3:121-126.

 ^{Am} See Craig M. Sonography: an occupational nearth nazard. JDMS 1985;Vol. 1 No. 3:121-126.
 Image of Aloka's 1971 real-time scanner courtesy of Aloka Co.
 Dr. Kato and Dr. Takemura's spectral traces are from the article: "Introduction of the Ultrasonic Doppler Technique in medicine: A historical perspective" by Yasuhara Nimura which appeared in the journal of Medical Ultrasound 1998, 6:5-13.
 Prefessor Joupilla's image courtesy of Dr. Aydin Tekay at the Oulu University.
 It is not possible to include all the names who have contributed significantly to the advancement of Obstetrical and Gynecological sonography, some who may have been less well-known than the others and some who may not have published so extensively in the English language. Apologies are extended to those whose contribution has not been fully credited in this article.

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A short History of the development of Ultrasound in Obstetrics and Gynecology

Dr. Joseph Woo



Multi-channel



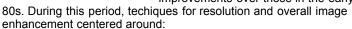
Technology Push and Consumer Pull



Ultrasound scanner technology

continued to develop and improve in the 1980s. Real-time scanners had rather standard appearance, sizes and fabrication. They are usually portable on 4 wheels with the monitor on the top of the console and rows of receptacles at the bottom to accomodate a variety of scanner probes. See some of these scanners here. By the mid 1980s curvilinear or convex abdominal transducers have come into the market which have a better fit to the Obstetric abdomen and have a wider field of view further from the transducer face. Curvilinear arrays have completely replaced the linear configuration by the late 1980s.

Prior to the 1990s, B-scan ultrasound images made steady progress in resolution and quality, but the improvements were not dramatic and except for a few really top-end brands, most had felt that images in the late 1980s did not have significant improvements over those in the early



small aperture

narrow band

the increase in the number of transducer crystals (or channels, from 64 to 128), improvements in transducer crystal technology (going into broad-band and high dynamic range), increasing array aperture (more crystals firing in a single time-frame), faster computational capabilities, improving technical agorithms for focusing on receive (increasing the number of focal zones along the beam), incoporating automatic time-gain controls and progressively replacing analog portions of the signal path to digital. See a brief discussion on the linear and phased-array principles.

Acuson Coporation®, a company founded in California in **1979**, marketed their first model <u>Acuson 128 System</u> in 1983, employing a 128-channel "<u>Computed Sonography platform</u>" based on a software-controlled image formation process. The machine shook the ultrasound community with its excellent resolution and clarity (and also the price). Many other companies followed on similar system designs. Other innovative breakthroughs were seen in designs from companies such as <u>ATL</u>® (Advanced Technology Laboratories), <u>GE</u>® (General Electric) and <u>Toshiba</u>®. The early to mid- 1980s was the time with the heaviest proliferation of standard-setting good quality machines. By the early 1980s there were over <u>45</u> <u>large and small diagnostic ultrasound equipment manufacturers</u> worldwide.

Image quality saw **real improvements in the early 1990s**. It is interesting to note that the availability of new and effective technologies to ultrasound scanners had also progressively stemmed from advances in technology in other areas of science such as **radar navigation**, **telecommunications** and **consumer electronics**. Such included the rapid developments in **cellular telephones**, **micro-computers**, **digital compact and versatile disk players**, and **high definition TVs**. The very high-speed digital electronics required for ultrasound application had become available at an affordable costs. The ultrasound imaging market alone would not have supported the development of these new technologies.

(Model number of scanners made after 1980 from important manufacturers are listed <u>here</u> with the year in which they were marketed).

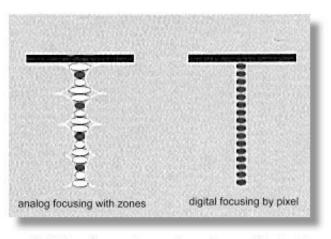
The new developments in the 1990s which has lead to some real enhancement in image quality and resolution include:

1. The entire signal processing chain becomes digital. The entire signal chain which includes:

[the transducer] --> [beamformer] --> [signal processor] --> [scan converter] --> [Monitor]

all operate under digital electronics.

Previously the beamformer (employing analog delay lines) and the signal processing stages are usually analog in their operation. The digital change-over was based on the very powerful computer platforms that were only available after the mid 1990s. The processor in the newer highend machines has the power equivalent of roughly 40 Pentium tm processors, executing some 20 to 30 billion operations per second. Most of the processing are also programmable software-based rather than hardwarebased and allow for much more versatility and finer adjustments in the manipulation of beam signals. Signals from and to the transducer elements are digitized before any signal processing, which is one of the most important advancement in



Digital beamformer reduces noise and increase focal points

ultrasound technology in the 90s. It opened the venue for dealing with some of more difficult areas in ultrasound physics.

Superfast digital beamformers allow for many times the number of focal points along the beam and produce microfine focal points on receive to the size of a screen pixel. Digital beamforming also reduces noise in the signal processing by several hundred folds producing a much cleaner picture.

2. Extensive use of refined broad-band wide aperture transducers, improving both definition of tissue textures and dynamic range. With wide aperture transducers, transmit and receive apodization also allowed for the electronic reduction of the lateral array elements (sidelobes). In the early 1990s there was much improvements in transducer material design and fabrication technology allowing for higher frequency transducers, improved sensitivity and contrast resolution. The number of channels in high-end systems went up to 256 and more recently to 512 and 1024 (2-D arrays) in several high-end systems allowing for extremely wide aperture on transmission and reception. In ultrasound physics, the lateral resolution is the product of the wavelength and the f-number. The f-number equals the depth of the returning echo divided by the aperture of the beam. (the aperture of the beam is the width of the number of simultaneous firing transducer elements in the array, that means the larger the aperture the more elements are fired simultaneously). Therefore lateral resolution will be best (smallest) if there is a large aperture and short wavelength (higher frequency).

Too large an aperture will slow the frame rate considerably and requires very fast computation and parallel processing. This has been made possible with the more recent digital electronics and the very powerful super-processors (see above). Many slightly older ultrasound systems are capable of using low f-numbers on reception at an affordable cost. However, they often employed large f-numbers on transmit in order to cover a large area. Significant improvement in lateral resolution requires low f-numbers both on transmit and receive. With the new 'very wide' aperture beamformer (often up to 128 channels), the transmit and receive f-numbers are lowered. The resulting improvements in lateral resolution can be as much as 4 times.

3. The phase data in returning ultrasound echoes, in addition to the amplitude data are processed in what is known as coherent image processing. The technique produced twice the amount of data from which to create ultrasound images of high resolution. The frame rate is also increased. The late 1990s has also seen transducer developing into 2D arrays which is made up of large number of elements arranged in rows and columns across the face of the transducer. Focusing occurs in two directions which produced a finer and clearer definition in



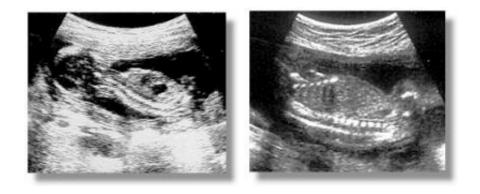
2-D arrays with focusing in two planes

both planes eliminating artifacts from adjacent tissue planes which may produce the partial volume effect.

4. <u>The advent of tissue harmonic imaging</u>. The technology, which has emerged as a major imaging trend in the last 4 years of the 1990s, made used of the generation of harmonic frequencies as an ultrasound wave propagates through tissue, dramatically reducing near field and side lobe artifacts. In a nutshell, tissue harmonic Imaging made use of lower frequency sound waves to improve penetration,

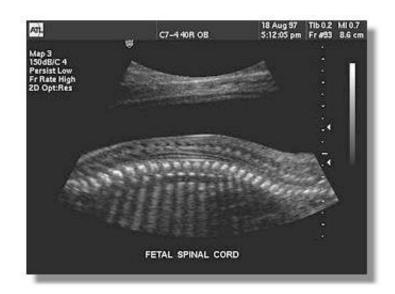
while receiving and processing only the higher frequency echoes produced by the body's inherent harmonic characteristics. This process can reduce clutter and improve image clarity significantly. As ultrasound waves propagate through tissue, there is non-linearities in sound propagation that gradually change the shape of the wave, a shape change that can only result from the development of harmonic frequencies within the wave. There are no harmonic frequencies present at the transducer face. They develop gradually as the wave propagates through tissue, and so in the near field there is very little harmonic energy available for reflection from tissue. Since the near field is a source of much of the artifact in the ultrasound image, selective display of harmonic energy will show dramatically less **near-field artifact**. The strength of the harmonic energy results from the strongest part of the beam, and weaker portions of the beam (side lobes, for example) generate relatively little harmonic energy. selective harmonic imaging will yield a **dramatically cleaner contrast** between adjacent tissue structures. It is a sonologist dream come true. The black is black and the white is white. Much of the fuzzy picture noise that have made diagnosis difficult are gone.

The development of harmonic imaging would not have been possible until the late 1990s as there must be excellent beam linearity on transmission and super sensitivity and dynamic range on receive to display the harmonic energy without an unacceptable amount of noise, as the harmonic signals are always much less in amplitude than the original fundamental signal. There must also be a very selective and fast digital filter within the receiver, to exclude the large percentage of the fundamental signal. Harmonic imaging is particularly useful in **obese patients**.





From left to right: Changes in image quality from <u>1985</u>, <u>1990</u> to <u>1995</u> respectively. There were improvements in spatial and contrast resolution, background noise reduction, dynamic range, and near and far field visualization.



More significant improvements came after the mid-1990's. This image from ATL ${f R} *$ demonstrating fetal spine and <u>cord</u>.

Ultrasound scanners came into <u>different categories</u> according to their performance and price. From the early 1980s, scanners have started to move into clinics and private offices and there is a trend to decentralise ultrasound services all over the world. Acceptance and demand from the lay public have also increased exponentially coupled with increased utilization by various medical specialties and sub-specialties. **Standards** and **quality of scans** became an emerging problem not seen in other areas of medical imaging, where Radiogists received the relevant training and underwent apppropriate examinations before running the service. **Obstetricians** were simply using the scanner probe as a torch to "look inside" the uterus. Standards varied and misdiagnosis was not uncommon. Obstetricians and Gynecologists took on the fact that they are the more suitable persons to do the scans as compared to their radiological colleagues. Special training centers and accreditation boards were gradually set up by the health authorities in the United States, Australia, Europe and other countries.





Joan Baker

was that medical insurance agencies such as "Blue Cross" had started reimbursing ultrasonographic examinations since **1966**. Medical sonographers also took up much of the responsibilities of the scans and sonography itself was recognised as a separate profession by the American Medical Association in 1974, the first sonography program being accredited in **1981** by the **Joint Review Committee for Education in Diagnostic Medical Sonography** (the **JRC-DMS**). The <u>American Registry of Diagnostic Medical Sonographers</u> was founded in **1974** by the <u>Society of Diagnostic Medical Sonographers</u> (SDMS). Joan Baker was the first Chairperson. The <u>AIUM</u> gradually went beyond developing guidelines and established an ultrasound practice accreditation program. Since 1996, this voluntary program has accredited over several hundred ultrasound practices. To achieve accreditation in obstetrics-gynecology, physicians and

sonographers must interpret a minimum of 170 ultrasound scans per year. In Australia, the Council of the Australian Society (ASUM) awards the Diploma of Diagnostic Ultrasound and the Diploma of Medical Ultrasound to medical practitioners and sonographers who successfully pass the examination.

In the United States, an impetus for the development of ultrasonography

And in both a '**Consumer pull**' and '**Technology Push'** situation the diagnostic application of ultrasound in the field of Obstetrics and Gynecology continued to expand into new horizons. In 1975, that is before the advent of real-time equipments, in the United States there were only 5 legitimate indications in obstetric sonography: • measurement of the biparietal diameter (and other dating purposes), • determining amniotic fluid volume, • diagnosis of early pregnancy failure, • evaluation of multiple gestations and • placental localisation. The indications have since the early 80's expanded into at least 2 dozen, including most notably the accurate evaluation of fetal growth and the diagnosis of fetal malformations.

Fetal biometry developed and 'flourished' in the 1980s as accurate fetal measurements do not require the prerequsite of very high resolution equipments. <u>At least two dozen measurements</u> were "invented" to assess gestational age and fetal







size, each claiming their unique usefulness. Nevertheless by the mid 1980s only a few parameters were considered as standard measurements and ones that had "stood the test of time". These include the crown-rump length (CRL), the <u>biparietal diameter</u> (BPD), the head circumference (HC), the femur length (FL) and the <u>abdominal circumference</u> (AC). Many other measurements were considered useful only in situations where fetal dysmorphology was in question.

Take for example the **BPD**, at least 200 charts are in used in the 1990s in different parts of the world (information supplied by commercial ultrasound scanner venders who set up the charts for their clients). Early workers in the United States who have published extensively on fetal biometry include <u>Rudy</u> <u>Sabbagha</u> at the Northwestern University, Chicago, <u>Alfred Kurtz</u> at the Thomas Jefferson University,







(top row) Rudy Sabbagha, John Hobbins, Peter Cooperberg. (bottom row) Roger Sanders, Alfred Kurtz, Charles Hohler. producing important work in biometry from the U.S.

John Hobbins at Yale, Charles Hohler in Phoenix, Arizona, Peter Cooperberg at the University of British Columbia, Canada, David Graham and Roger Sanders at Johns Hopkins, Baltimore and Frank Hadlock and Russell Deter at the Baylor College of Medicine, Houston, Texas. There are others from Britain and Europe like <u>Hansmann</u>, Jouppila, <u>Kurjak</u> and <u>Levi</u>. Fetal biometry was explored from many different perspectives and in different populations.

The <u>abdominal circumference</u> measurement which was described by **Campbell** and **Wilkin** in **1975** remained the mainstay measurement in the evaluation of fetal growth and nutrition. The assessment of **gestational age** and **intrauterine growth retardation** using ultrasonic parameters was the subject of a **huge number** of research papers. Fetal growth analysis and charting were also performed on desktop personal computers (PC) using commercial or home-made proprietory softwares.

 Stuart Campbell published the first BPD chart in 1971. Since then, Charts and Tables had become an important and integral part of Obstetric practice, at which Obstetricians and Gynecologists were slowly getting used to.

Estimation of in-utero fetal weight basing on the combination of the biparietal diameter and the thoracic circumference was first reported in 1965 by the <u>Thompson</u> group in Denver, Colorado. He reported an accuracy of within 300 grams in 66% of the weight estimates. This was followed by work from Garrett in Australia, <u>Hansmann</u> in Germany and <u>Campbell</u> in England. In 1977 the <u>Hobbins'</u> group at Yale published one of the most important papers in fetal biometry, "Estimation of fetal weight by computer-assisted analysis of fetal dimensions" which had started in the next 10 to 15 years, an almost non-stopping search all over the world for computer-generated models of fetal weight determination basing on multiple fetal parmeters.

Log₁₀ Weight = -1.7492 + 0.166 (BPD) + 0.046 (AC) - 0.002646 (AC) (BPD)

One of the several popular weight estimating equations, this one from MA Shepard and co-workers Normograms and weight estimations have become standard packages built into each ultrasound machine



Frank Hadlock

Similarily the craving to produce normograms for incremental growth of every 'measurable' part of the fetal body never stopped. One can find gestationalage normograms for parameters such as the Binocular diameter (Mayden K et al, 1982, Jeanty P et al, 1982), thigh circumference (Deter et al, 1983), clavicular length (Yarkoni et al, 1985), fractional spine length (Li DF et al, 1986), fetal foot length (Mercer BM et al, 1987) or fetal ear length (Birnholz JC et al, 1988); and charts for parameters such as cerebral ventricular width, cardiac chamber dimensions, chest circumference, limb lengths,

renal and **calyceal sizes** that were of great value in the assessment of specific fetal anomalies. In **1987 E Albert Reece** at Yale demonstrated the usefulness of the **trans-cerebellar diameter** as a growthindependent parameter to assess gestational age.



Software generated growth analysis and charts



Among many others, the comprehensive normograms of **Frank Hadlock** and <u>Russell Deter</u> from the Baylor College of Medicine, Houston, Texas were widely used.

The group had in particular incorporated the **femur length** measurement into the calculations and popularised the concept of limb length/ trunk circumference ratios in the assessment of fetal growth.



Russell Deter

In the last 40 years or so in the development of ultrasonic **fetal biometry**, there has been much effort among the obstetric and ultrasound community to devise and develop fetal measurements that would be able to distinguish between fetuses that are small because of **nutritional reasons** and those fetuses that are **"by nature"** small. **Head, limb versus abdomen ratios** were once thought of be of promise but as it was noted that the growth-retardation process also affected fetal head and limb growth in varying degrees, the value of these ratios did not stand up to their

initial expectations. Up to this day, there is still not one or several size measurement parameter in combination that can **unequivocally diagnose growth retardation** in the fetus when a woman is seen for the first time in the later part of pregnancy.

Visualization of the the **fetal yolk sac** with the real-time scan was first described by **Eric Sauerbrei** and **Peter Cooperberg** in Vancouver, Canada in **1980**. It is interesting to note that perhaps because of its size (and hence the difficulty to visualize with existing equipments at that time) its significance and usefulness in early pregnancy failures was not discussed until much later in the second half of the **1980**s.

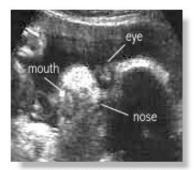
The diagnosis of fetal malformations obviously received the enormous attention that was deserved and findings of many abnormalities diagnosable by ultrasound have been described.

Ian Donald included a case of hydrocephaly in one of his early "introduction" ultrasound papers in 1961, which demonstrated "tissue interface within the body by ultrasonic echo sounding". In Bertil Sunden's thesis in 1964 there was description of the diagnosis of anencephaly in the third trimester using on the bi-stable Diasonograph. In 1968 D Hofmann and Hans Hollander in Germany reported on 9 cases of 'hydrops fetus universalis' diagnosed with the Vidoson (see Part 2) and William Garrett in Sydney reported the diagnosis by ultrasound of a fetus with polycystic kidneys using the CAL echoscope in 1970 (see also Part 1). These two papers were probably the two earliest papers describing formally the diagnosis of a congenital anomaly using ultrasound. Both reports were about cases in the third trimester and resulted in fetal death.



Large Exomphalos were considered as "straight forward" diagnosis

The diagnosis and management of a 17 weeks **anencephaly** was reported as early as **1972** by <u>Stuart Campbell</u> using statc B-mode equipment. This was followed by the diagnosis of **spina bifida** in **1975**. Both reports had appeared as landmark papers in the Lancet. They were the first cases of such conditions in which a correct diagnosis by ultrasound had effectively led to a termination of pregnancy. <u>Manfred Hansmann</u> in Bonn, Germany and John Hobbins at yale were among others, early pioneers in the ultrasonographic diagnosis of fetal malformations. With the advent of better real-time scanners, many more malformations were diagnosed, abeit in the late second trimester when fetal organs become more discernible on the scans. A review published in **1981** (Stephenson and weaver) reported that around <u>90 different fetal malformations</u> had been diagnosed by ultrasound.



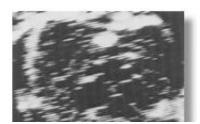
Looking at fetal face and lips

Common anomalies that were considered "straight forward" to diagnose at that time included **anencephaly**, **hydrocephaly**, **exomphalos**, **duodenal atresia**, **polycystic kidneys**, **hydrops fetalis** and **limb dysplasias**. More difficult areas for diagnosis of malformations were the **fetal face**, the **fetal extremities** and the **fetal heart**. The diagnostic accuracy progressively improved with more experience and better resolution machines. With the advent of the newer **high-resolution** scanners and the transvaginal transducer the diagnosis of these and other more subtle conditions were achieved, and particularly at an earlier gestation, moving from the third trimester of pregnancy to the second and later on to the **first trimester** in the latter half of the 1990's. **Fetal trisomies**, **spina bifida** and the more subtle **cardiac anomalies** were among the many examples. So-called **soft signs** and **sonographic markers for chromosomal anomalies** (**see below**) were started to be described.

 The ability to recognise and follow up in utero of all these malformations by ultrasound has opened up further the entire avenue of "Prenatal Diagnosis" and has markedly enhanced and pushed forward the study of congenital abnormalities among obstetricians, pediatricians, geneticist, pathologist and other allied specialties. All of a sudden, obstetricians started to learn about so many congenital malformations that they have not even heard of.



The diagnosis of fetal **cardiac malformations** gained foot in the early 1980s. Pioneers included the <u>Wladimiroff</u> group in Rotterdam, the Netherlands; the <u>Hobbins</u> group (<u>Charles Kleinman</u>, <u>Greggory Devore</u>, <u>Joshua Copel</u>, **Peter Grannum**) at Yale; <u>Lindsey Allan</u> at Guy's Hospital, London (now in New York); L W Lange, David Sahn at Portland, Oregon; <u>Kathryn Reed</u> at Tucson, Arizona and <u>Beryl Benacerraf</u> at Harvard.



The first real-time fetal heart images and quantitative data were published by the Lange, Sahn and Reed group in Tuscon, Arizona in 1980. Allan published her

echo/anatomical correlates in the same year. <u>Allan</u>, a pediatric cardiologist, described systematically real-time normal and abnormal ultrasonic anatomy of



A 4-chamber view from the early 80's

the fetal heart which laid the foundation for subsequent studies. Using ultrasonic equipment available in the early 1980s and much painstaking enthusiasm and skill, she had very importantly shown that real-time **cross-sectional study** and **diagnosis of fetal cardiac anomalies** in utero in the **second trimester** was a distinct possibility and would no doubt develop into a distinctive diagnostic science.



Lindsey Allan

Dev Maulik

The improvements in diagnostic capabilities that came with the 'new technology' scanners in the 90's had tremendous impact on fetal cardiac diagnosis. The usefulness of direct doppler interrogation of **fetal intracardiac flow** was first demonstarted in **1985** by **Dev Maulik** and **Navin Nanda** (Professor of Cardiology) at the University of Alabama. **Allan** and **Reed** followed up with more publications. The **Maulik** group further demonstrated the value of **color doppler** in fetal cardiac

studies in **1986**. **Greg Devore** soon popularised the use of <u>doppler</u> <u>color flow mapping</u> in the assessment of fetal cardiac malformations and particularly in a **screening situation** in **1987**. The use of color doppler has become indispensible in the diagnosis of more complicated cardiac malformations. By the late 1990s, the diagnostic accuracy of the nature of complex cardiac malformations in utero can be as high as 95 percent.



color doppler assessment of a VSD ***

Read also: the <u>American College of Cardiology Position Statement on Doppler Echocardiography in the Human</u> <u>Fetus prepared by Charles Kleinman</u>, James Huhta, and Norman Silverman which covers some of the historical aspects of fetal doppler echocardiography.

The accuracy of diagnosing fetal malformations in a "high-risk" populations have been variously reported in the early 1980s right up to the early 1990s. These included some classic reports from Kurjak in Croatia (1980), Campbell and Pearce in London (1983), Gembruch and Hansmann in Germany (1984), Sabbagha in the U.S. (1985) and a good number of other authors. Diagnostic accuracies in "targeted" scans progressively increased with improvement in instrumentations and accumulation of knowledge about the ultrasound characteristics of the various anomalies.

With improved resolution in the new equipment, diagnosis of fetal cardiac anomalies have moved into the late first trimester. All the improvements in machinery and earlier detection of abnormal structures in the fetus have nevertheless brought along with it "false positives" and difficult-to-be-sure-what-will-happen diagnosis that generate much undue anxiety in patients. Such has far-reaching effects on a woman's perception of child-bearing. Researchers try very hard to determine the course of events for a particularly abnormality or the implication and prognosis of a certain finding on ultrasound examination so that proper counselling can be done to alleviate anxiety and uncertainty from the diagnosis.



"Textbooks" in Obstetrical and Gynecological sonography had emerged out of necessity because of the large amount of information that had become available and as reference materials of those sitting for examinations. Books had gone from being just overviews and atlases to systematic discussions of ultrasound techniques and findings. Notably the books by <u>Peter Callen</u> at the University of California, San Francisco (1983) and **Sanders/James** at Johns Hopkins were popular and represented two of the earliest standard textbooks in the field. Multiple authorship contributed to the excellence of these texts. Many other texts followed, some devoted mainly to special topics such as fetal anomalies or doppler ultrasound.



Peter Callen



Karel Marsal

The advent of the real-time scanners also prompted research into **body movements** and **breathing movements** of the fetus The study of fetal breathing movements (FBM) was first suggested by **Geoffrey Dawes** and **K Boddy** at the Nuffield Institute of Medical Research, **Oxford Univeristy, England**, in the early '70s, in that the presence or absence of breathing movements, theiir amplitude and intervals will be indicative of fetal well-being. Much research into these areas came from the <u>Karel Marsal</u> group at the University Hospital at Malmo, Sweden, the **Tchobroutsky** group at the Maternite de Port-Royal, paris, the <u>Wladimiroff</u> group at Rotterdam, and the **Brian Trudinger** group in Australia. all of them having switched to the use of real-time apparatus in the early to mid '70s. <u>Wladimiroff</u> demonstrated in **1977** that maternal hyperventilation decreased fetal breathing. The quantitative documentations of fetal breathing movements however require elaborate equipments, and was very time-consuming, so much so it would be difficult to be

incoporated into clinical practice. The results also have wide overlap between positives and negatives. The advent of realtime had also raised hopes of being able to study **physiological responses**, <u>sensations</u> and **behavior** in the fetus. Again the assessment were time consuming and results were often equivocal, which made them unsuitable as clinical tests. <u>Jason Birnholz</u> at Harvard published several pioneering papers in these areas including the assessment of fetal movement patterns as a possible means of defining neurological **developmental milstones** in-utero and the development of **fetal eye movements** and possible 'dream states' in the feus.

The study of FBM and FM patterns did not gain further popularity although the semi-quantitative counting of FBMs and the documentation of 'adequate" fetal body movements were popularised further in **fetal biophysical profile** scores made popular by <u>Frank Manning</u> and <u>Lawrence Platt</u> in Winnipeg and Los Angeles, who had started studies into FBMs round about the same time as their European counterparts. In **1980**, **Manning** and **Platt** reported on the important finding that a **reactive cardiotocographic finding** was just as predictive as the presence of **FBMs** or total fetal movements.

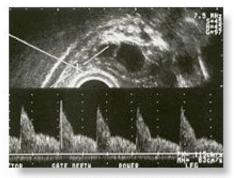


juiry Wladimiroff

Spectral doppler or **doppler velocimetry** (the study of doppler waveforms), had evolved to become a standard tool in the late 1980's in the assessment of fetal wellbeing and compromise. (See **Part 2** for more of its early development).

Its advent in Obstetrics in the mid 1980s had fascinated (or perhaps *misguided*) many as *the* new promise in fetal assessment. The **absence** and **reversal** of **end-diastolic flow** in the **umbilical arteries** in severely compromised fetuses were striking demonstration of fetal pathophysiology, so was the finding of clear and

unequivocal **increase in diastoic flow** in the **middle cerebral arterial** waveforms in mounting fetal hypoxia. It has also become clear that umbilical doppler velocimetry does not correlate with fetal weight in utero, nor is useful as a screening procedure.



Duplex doppler became standard in the 1990s



Torvid Kiserud

Stuart Campbell's group at the King's College Hospital in London reported in 1983 the evaluation of utero-placental flow velocity waveforms in compromised pregnancies with duplex doppler, and described the 'frequency index profile'. In 1984, P Reuwer in Utrecht, the Netherlands first discussed the ominous significance of absent end-diastolic flow in the umbilical artery. Further work from the Campbell group, including work from Gerald Hackett and T Cohen-Overbeek in 1986 and 1987 clearly demonstrated the prognostic significance of absent end-diastolic flow in the fetal descending aorta. In 1986, Brian Trudinger in Australia demonstrated that abnormal doppler waveform patterns tended to preceed abnormal cardiotocographic traces. In the same year the Wladimiroff group reported the value of middle cerebral artery waveforms in the assessment of severely compromised fetuses. Sanjay Vyas working at King's College Hospital in England described the use of renal artery waveforms in 1989. The value of fetal Venous blood flow in the assessment of fetal compromise was first suggested by Torvid Kiserud in Bergen, Norway in 1991. Giuseppe Rizzo at the Universita di Roma Tor Vergata in Italy furthered expounded the usefulness of the ductal venus velocimetry

in fetal acidemia and cardiac decompensation. In 1987, <u>Asim Kurjak</u> introduced the use of color flow doppler in fetal assessment.

Doppler ultrasound became a standard and indispensible tool in the evaluation of progressive **fetal anoxia** (the umbilical artery), **compensation** and **decompensation** (the middle cerebral artery), **acidosis** and progression to **cardiac failure** and **emminent fetal death** (the ductus venosus). It has also been employed in the assessment of women at risk of **pre-eclampsia** and **utero-placental arterial compromise**, leading to early and effective therapeutic intervention. It is of interest to note that historically, these velocimetric parmeters have appeared each a number of years apart with increasing sophistication of the apparatus. By the beginning of the 1990's, most mid- to highend ultrasonic equipments had incoporated **duplex doppler** as standard facility. In the mid 90's, **color flow mapping** had also found its way into most mid- and highend machines. Aside from aiding **cardiac diagnosis** (see above), impressive flow images were often popularly reported such as those found in **Vasa Previa** and the fetal **Circle of Willis**.

 Doppler velocimetry is not only a fascinating demonstration of fetal physiology that provides a vital assessment of fetal well-being, but will likely change the concept of routine antenatal care by picking up placental insufficiency at the earliest stage.



Amniocentesis with the Vidoson

As early as **1967**, and basing on the **Vidoson**, **Hofmann** and **Hollander** in Germany had discussed the importance of placental localization using ultrasound before amniocentesis. **Jens Bang** and **Allen Northeved** in Copenhagen described ultrasound-guided amniocentesis in **1972**. In the mid 1970s to early 80s, <u>genetic amniocentesis</u> was largely performed under static B-scan ultrasound guidance. An ultrasound scan was performed to locate a



Puncture adapter on a

feasible pocket of amniotic fluid before a

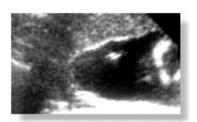
tap, which was done basically without actually seeing the needle tip going into the fluid pocket. With the advent of realtime scanners, a small number of centers had in the late 1970s started to perform amniocentesis by simultaneously visualizing the puncture needle tip on the scanner monitor. One such pioneer was the **Birnholz** group at **Harvard** who used an early **phased array** for the purpose. <u>Needle-guide adapters</u> soon became available from ultrasound manufacturers which could be coupled to the linear array or phased array sector probes where the needle passed through a fixed path either parallel or at an angle to the ultrasonic beam. These were **cumbersome** to use however, particularly in a busy setting. They also had serious problem of keeping the equipment **sterile**. The adapters may also increase the risk of traumatisation as it did not allow for the 'desired' and sensitive placement of needles.



Single operator 2-hands technique

Many centers started to do it <u>freehand</u> with an assistent holding onto the transducer probe that was commonly wrapped in a sterile adhesive drape. In 1984, <u>Wolfgang</u> <u>Holzgreve</u> in Basel, Switzerland described a large series of over 3000 'freehand' amniocentesis with low complication rate. Similar experience was also reported by Lawrence Platt in Los Angeles, who

emphasised on the need for the transducer probe to be manipulated by the same operator which resulted in better hand-eye co-ordination. In the following year, **Roberto Romero** at **Yale** formally described the single operator two-hands



Siemens prope

Visualization of the needle and tip through the placeta into the fluid

technique in amniocentesis and the reduction in the number of multiple taps and bloody taps associated with the procedure. Most centers soon adopted this single operator technique, which had become popular because of it convenience and effectiveness. Chorionic Villus sampling (CVS) also relied heavily on sonographic guidance. Z Kazy and his group in the USSR reported in 1980 fetal sexing and enzyme assay on chorion biopsies taken at 6 -12 weeks' gestation, using either an endoscopic or ultrasound-guided approach. RH Ward in London and Bruno Brambati in Milan both reported transcervical CVS under ultrasonic guidance in 1983. Brambati reported the success rate of obtaining chorionic villi rose from 75% without ultrasound to 96% with ultrasound guidance. Danish investigators Steen Smidt-Jensen and N Hahnemann first described the ultrasound-guided transabdominal approach in 1984. Other adjunctive ultrasonic techniques were reported by the Brambati group and the Golbus group in San Francisco in 1985.

The <u>Beryl Benacerraf</u> group at Harvard (se below) reported the feasibility of **early amniocentesis** (11-14 weeks) in **1988**. In 1990, the same group reported an early fetal loss rate of **over 2.3%**. Several important reviews in the mid-90s confirmed this **high incidence of fetal loss**. The practice has for this reason not gained general acceptance.

Read here <u>a short history of Amniocentesis</u>, fetoscopy and chorionic villus sampling.



Fernand Daffos

Other interventional intra-uterine diagnostic and therapeutic procedures also started to catch on. After fetoscopy, **ultrasound-guided** pure fetal blood **cordocentesis** was pioneered in France in **1983** by **Fernand Daffos**. Pure fetal blood was aspirated in-utero at around 18 weeks from the umbilical vein near the placental insertion of the cord using a twenty gauge needle under ultrasound guidance. Their group reported the first case of Haemophilia A diagnosed in-utero by this method. The procedure was also popularised around the same time in England by the <u>Stuart Campbell</u> and **Charles Rodeck** group at King's College Hospital. The <u>Hobbins</u> group at **Yale** described their technique in **1985** and called the procedure **percutaneous umbilical blood sampling** (**PUBS**). This replaced blood sampling via **fetoscopy** which the group had pioneered in **1974**. **Kypros**



Kypros Nicolaides

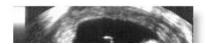
Nicolaides at King's developed the single operator two-hands method and became a leading figure in cordocentesis exploring many important aspects of fetal physiology and pathophysiology. With the advent of color flow mapping, the technique has become even more accessible. In **1988**, **Umberto Nicolini**, working with **Charles Rodeck** at the Queen Charlotte's Maternity Hospital in London, first described fetal blood sampling from the **intrahepatic** portion of the **umbilical vein** in the fetus, as an alternative procedure in cases where cord needling was unsuccessful. In the **late 1980s** fetoscopy has mainly been reserved for **tissue** or **organ sampling**, and fetal blood sampling has evolved to become one of quick **confirmation of abnormal karyotype** in the 22-24 weeks fetus, when a chromosomal abnormality has been suggested on ultrasound scan.

Various in-utero drainage procedures under ultrasound guidance were described in the late 1980s such as the drainage of **Chylothorax** and **hydronephrosis**, and the treating of fetuses by **transfusion** into the umbilcal vessels. **Selective reduction** of the number of fetuses in cases of triplet or quadruplet pregnancies was first described by **Richard Berkowitz** in New York in **1988**, using intra-cardiac potassium chloride injections given under ultrasonic control. **Vesico-amniotic shunt placement** was described by the <u>Mitchell Golbus</u> group (with <u>Michael Harrison</u>, <u>Roy Filly</u>, <u>Peter Callen</u>) in **San Francisco** in **1982**. The group became one of the most important forerunners in **fetal surgery** and continued to make many new fetal surgical innovations. In that year, they published their multicenter classic paper "**Fetal Surgery 1982**" in the New England Journal of Medicine.

Read here <u>a short history of Amniocentesis</u>, fetoscopy and chorionic villus sampling.



With improved resolution from transvaginal sonography, **Mark Cullen** at Yale first reported in **1990** a large series of congenital anomalies detected in the **first trimester**

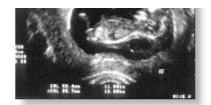




Ilan E. Timor-Tritsch

using **transvaginal ultrasound**, and pointed out the importance of a good understanding of normal embryonic development in such diagnosis. Many studies followed around the same time reporting on the usefulness of the first trimester **transvaginal scan** in the evaluation of fetal anomalies. **Moshe Bronshtein**, working in **Haifa, Israel**, described extensively since the early 1990s results of transvaginal sonography in the first trimester. A similar

report to Cullen's also appeared in **1990**. <u>Ilan E. Timor-Tritsch</u>, working in Israel and later on at New York University, followed on with many reports on fetal anatomy and anomalies systematically studied using high resolution transvaginal



Cystic hygroma diagnosed at 12 weeks with vaginal scan

transducers in the first trimester, opening up convincingly a new area in fetal ultrasound diagnosis, that of "sonoembryology". Timor-Tritsch was also credited for organizing the first three transvaginal ultrasound courses in the United States.

As ultrasound became a widely available and popular investigation, it contributed heavily to several **population screening programs** that took place between the late 70's and the 90's. The first was the **Maternal serum alpha-feto protein (MSAFP) screening** programs for the detection of neurotube defects (NTD). It started in the United Kingdom in the late 1970s, notably in parts of **Scotland**, where the prevalence of NTDs was high.

In 1972, David Brock and R Sutcliffe measured the AFP values in the amniotic fluid of 31 pregnancies with anencephaly and 6 pregnancies with spina bifida. All of the cases of anencephaly and most of those with spina bifida demonstrated markedly elevated AFP levels. This was a landmark 'discovery' in the history of prenatal diagnosis. In 1974, <u>Nick Wald</u> and co-workers at the University of London reported in the Lancet maternal serum AFP levels in 7 pregnancies with open neural tube defects which were significantly higher than that in 14 other controls. This led to the idea of measuring MSAFP in screening for NTDs. The 19-centers U.K. Collaborative study in 1977 subsequently demonstrated the utility of this test for prospective open neural tube defect screening. Similar results were arrived at in large-scale studies in the United States in 1979. By 1984, MSAFP screening had also become part of standard antenatal care in the U.S.

As scanner resolution and sonography skill improved, **ultrasound** gradually replaced **amniocentesis** in the diagnosis of screened-positive cases. By the mid-1990s amniocentesis is often not performed in patients with elevated MSAFP levels. Either a positive or negative diagnosis is made **basing solely on ultrasound findings**.



Salvator Levi

The second was the **routine fetal scan** at 20 weeks which had progressively become an integral part of antenatal care in the early **1990's**.

In the late 1970s a number of **large scale population ultrasound screening studies** have been described variously from London, Germany, Brussels, Sweden, Norway, Finland and other countries in Europe and in the United States. At least 20 similar large-scales studies were reported up to the year 1990.

There is apparently a distinct difference in the attitude towards routine ultrasound screening between the Europeans and the Americans. **Routine screening scans** were introduced in **Germany** in 1980, in **Norway** in 1986 and in **Icelend** in 1987. The scans basically try to date the pregnancy, exclude twins and detect any fetal malformations that may be present. In the U.S., routine scans in pregnancy has however been looked upon with much controversy and their **cost-effectiveness** and validity in improving **'quantifiable' perinatal outcome** has not been

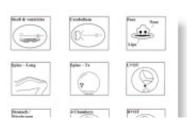
firmly established. Nevertheless, those who argue for a routine scan would claim that parents have a natural desire to know if any fetal congenital anomaly is present or if there is any health compromise in the fetus. Being able to reassure the parents is a natural part of prenatal care.

Two important large scale studies emerged to address the issue: the **RADIUS study** (Routine Antenatal Diagnostic Imaging with Ultrasound) with a cohort of 15,000 low-risk pregnancies in the United States in **1993** and the <u>Eurofetus</u> <u>Study</u> in Europe in **1997** in which 200,000 low-risk pregnant women in 60 hospitals had obstetric ultrasound examinations performed in centers proficient in prenatal diagnosis. <u>Salvator Levi</u> was the **Project leader** of the **Eurofetus Study Group** and a strong proponent for routine screening. A **61**% overall detection rate of structural anomalies in the Eurofetus study contrasted sharply with the **35%** overall detection rate in the RADIUS study.

In order to address the conflicting data and conflicting opinions on this topic, a conference was held at The Rockefeller University in **New York City**, in June **1997**, sponsored by the New York Academy of Sciences. Over **150** scientists and clinicians participated in the meeting, with highly informative presentations and discussions.

The following conclusion was made:

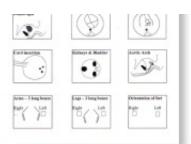
" In summary we have tried to put together comprehensive stateof-the-art information on the routine obstetric ultrasound controversy. Our conclusion is that routine obstetric ultrasound is warranted for all pregnancies, but only if it is performed in a quality manner. Although there is still scientific and economic controversy about our conclusion, we would argue that, at a minimum, there is an ethical obligation to present the option of an 18-22-week routine obstetric ultrasound examination in



clinical centers in which quality ultrasound is available. We hope that our efforts will move public policy in this direction and encourage further discourse on this most important topic in contemporary obstetrics." -Preface, Ann NY Acad Sci 1998 847: 1-ix.

A discussion on the findings of these two important studies can also be found here.

In any case, by the late 1990s and the turn of the century, an ultrasound examination for each and every pregnancy at 20 weeks has become quite standard practice worldwide, as long as facilities are available. As the skill of ultrasonographers and equipment improved the diagnostic accuracies of the examinations also improved. This improvement is also brought on by the large number of *ultrasound training courses* throughout the world. Some, like the fetal echocardiography course at Guy's Hospital in London has been shown to



scanning guidelines formulated for 20 weeks screening. Click to enlarge

effectively improve the pick-up rate of congenital heart defects in certain parts of the United Kingdom. Many other centers in the United States, Europe and the United Kingdom, also run well-attended and effective courses in the prenatal ultrasound diagnosis of congenital anomalies. Guidelines formulated by accreditation bodies also have a positive effect on ensuring quality of the examinations.

> The standard 20 weeks fetal examination could have been the single most important development in the practice of Obstetric ultrasound in the last 40 years, ever since the day of the biparietal diameter. It was a development that has culminated from developments in skill, training, practice, attitudes, machinery and administration. •



Beryl Benacerraf

The third was the screening for chromosomal abnormalites, notably Down syndrome, which became popular in the mid **1990's**. Screening basing on multiple biochemical parameters in the low-risk population had started around 1990 in some centers and because of its relative convenience had soon caught on and become standard antenatal care in many parts of the world.

Screening basing on ultrasonic parameters had not become popular until the mid-1990s. Measurement of the nuchal translucency, which was first described by Beryl Benacerraf and her group at the Harvard Medical School in 1985 formed the basis of such screening. Working with fetuses between 15 and 20 weeks of gestation her group discovered good correlation between a thickened nuchal skin fold that was above 5mm and the presence of Down syndrome in the fetus. Benacerraf had also earlier on published biometric parameters (shorter femurs and humerus, and

decreased femur length/ biparietal diameter ratios) as markers for the diagnosis of Down syndrome which formed one of the earliest observations and endeavours for an "indirect" diagnosis of chromosomal anomalies. Endocardial cushion defect, atrial and ventricular septal defects, omphaloceole, pyelectasis, choroid plexus cyst, echogenic cardiac foci, echogenic bowels, hypoplasia of the middle phalanx of the fifth digit and simian creases were considered as other 'indirect' or 'soft' signs of fetuses with Down syndrome.

Benacerraf had on top of other things brought to the attention of researchers and clinicians the necessity and feasibility of visualising 'small' abnomalites in the fetus and had in so doing pushed scanning skills, machine resolution and operator patience to the limit. Anomalies such as club foot, early hydrocephalus, phalangeal abnormalities, facial clefts are some of these examples. In a contribution to a book on prenatal diagnosis Benacerraf wrote in 1989^:

"..... Although many fetal anomalies -- such as anencephaly, hydrocephalus, and anterior abdominal wall abnormalities -- can now be diagnosed sonographically even in the second trimester, more refined diagnosis involve examination of the face and extremities. It is not sufficient, however, to diagnose isolated cleft lip and palate or a clubfoot. Rather, the presence of these lesions should stimulate the ultrasonographer to seek a pattern and fit the pieces of the puzzles together in order to recognize the syndromes associated with chromosomal anomalies

The reason for the seemingly 'late' popularity of 'ultrasonic' screening was probably twofold. Much time was spend initially on making the measurement at 16-19 weeks (following important work from Benacerraf, James Crane, Hélène Grandjean and others) and not at 11-14 weeks as practised now (see below). Measurement of the nuchal fold at this latter gestation is demanding on operator skill and machine resolution and is also error prone.

With improved resolution of ultrasound scanners, better understanding of fetal pathophysiology and more emerging data, ultrasonic screening for nuchal fold thickness has moved from between 16 and 19 weeks (using a cutoff level of around 6mm) to between 11 and 14 weeks (using a cutoff of around 3mm) in the first trimester. Kypros Nicolaides and his group at King's published the landmark paper in 1992 in the British Medical Journal, where the measurement of nuchal translucency between 11 and 14 weeks was used to screen for Down syndrome. He



measurement of the nuchal skinfold

demonstrated the importance of likelihood ratios in the detection. The group later on turned out some of the most important data regarding the appication of nuchal translucency measurements including risk estimates and the quantitization of the measurement into gestational-age related multiples of the median (MoM).

Read also: <u>A discussion on the Nuchal Translucency</u> from an online book by <u>Kypros Nicolaides</u>, NJ Sebire and RJM Snijders.



Ovarian follicles on vaginal scan The imaging modality quite dramatically altered the management of infertility patients

In **Gynecology**, ultrasound has started as a diagnostic tool in the differentiation and assessment of solid, cystic or mixed masses in the pelvis. Even in the late 1970s (still very much a static-B era), it has already become a well-established and indispensibe tool in the evaluation of a variety of pelvic pathologies. In **1976**, the <u>**B-J Hackelöer**</u> and <u>**Hansmann**</u> group in Germany, basing on the static Bscan, reported on the successful monitoring of **follicular size** and **number** in patients undergoing ovulation induction. A 'very-full' bladder was a pre-requisite for good visualization of the ovaries. Follicular growth was noted to be linear and is around 1-4mm/day in the pre-ovulatory time period and follicles ovulate when they reach 15 to 28 mm in diameter. Other morphological parameters were also described. Ultrasound monitoring was 'formally' introduced into ovulation induction programs in **1979**. By about 1980-82 there were a number of important reports attesting to the usefulness of abdominal ultrasound in the assessment of follicular development and ovulation. In **1982**, the **Colm O'Herlihy**, <u>Lachlan de</u> <u>**Crespigny** and <u>**Hugh Robinson**</u> group at the **Royal Women's Hospital** in Melbourne, Australia, published on important follicular size criterion and protocols</u>

for ovulation inductions. Other important early work had also come from the <u>Joupilla</u> group in Finland, the Lopata group in Melbourne, the Queenan and O'Brien group in England and the <u>Fleischer</u> group in Tennessee. Transvaginal scanners replaced the abdominal counterparts after they became available in the mid 1980's. The addition of endometrial evaluation using transvaginal scanning enhanced diagnostic accuracies in the mangement of ovulation induction cycles. Follicular and edometrial sonography, although tremendously useful when used in combination with estrogen assays was unable to ro predict ovulation and avoid multiple pregnancies.

Vaginal sonography had also become indispensible in the evaluation of **non-palpable masses**, **ascites**, **uterine and cervical leisions**, **early pregnancies** and the **localization of IUCDs**. It's value as a tool in the diagnosis of **ectopic pregnancies** and **ovarian and edometrial cancers** was extensively re-evaluated in the late 1980's and later on in the early 90's with addition of **transvaginal color flow imaging** (see below). As mentioned the greatest development of transvaginal imaging in the late 1980s has been in **assisted reproduction**, where all aspects of diagnosis and management are incomplete without a vaginal scan. From initial assessment for pelvic pathologies to surveillance of ovarian follicles and edometrial responses with or without medications, to ovum retrieval in In-vitro fertilization/ embryo transfer cycles, vaginal sonography had become essential and indispensible. The diagnosis of **ectopic pregnancies** continued to be a challenge, despite better machines and the transvaginal approach. A number of authors such as **Roberto Romero** at **Yale** devised diagnostic criterion for making a diagnosis, combining the use of sonography, HCG levels and color doppler assessment, which allowed a vast majority of the diagnosis to be made.



Susan Lenz

Interventional sonography in gynecology dated back to the early 1970s when <u>Hans Henrik</u> <u>Holm</u> described percutaneous puncture of ovarian tumours in **1972**. They performed over 500 procedures with very few serious complications. It was not until **1982** that **David Graham** and <u>Roger C Sanders</u> at the Johns Hopkins Hospital, Baltimore, revisited the idea of transvaginal aspiration of pelvic masses under transabdominal ultrasound guidance. There was a necessity to develop similar techniques for the retrieval of follicles in IVF programmes which has hitherto been achieved only through laparoscopy. <u>Susan Lenz</u> and JG Lauritsen at the University Hospital Rigshospitalet in **Copenhagen** described percutaneous transabdominal - transvesical aspiration of ovarian follicles in **1981** and **1982** which showed for the first time that ovum retrieval can be performed as an ultrasound-guided and out-patient procedure.

Transvaginal ovum retrieval under abdominal ultrasound guidance was further described by <u>Norbert Gleicher</u> in Chicago in **1983** in a letter to the **Lancet** and several months later by the **P Dellenbach** group in Shiltigheim, France in **1984**. They reported for the first time successful pregnancies (5 out of 30 patients) using this technique. They further reported on favorable results in **1985** in more than 100 cases of oozyte retreival using this 'transabdominal scan - transvaginal puncture' method.





W Feichtinger

The advantage of this technique is that the ovaries are more accessible and the procedure is safer and relatively pain free. More importantly, the procedure is

Kretztechnik's mechanical rotary vaginal scanner in 1985 with puncture attachment

repeatable on an out-patient basis, and dramatically cuts down the cost of the IVF procedure. The true impact on ovum pickup came with the appearance of the mechanical **transvaginal sector scanner** from <u>Kretztechnik</u> in **1985** when <u>Wilfried Feichtinger</u> and <u>Peter Kemeter</u> in Austria described its use in transvaginal aspiration of ovarian follicles for IVF. Since then, ovum retrieval had steadfastly become an outpatient routine compared to just a few years ago when it was done as a laparoscopic procedure under general anaesthesia. The technique has also found its way into many ultrasound-guided interventional procedures in gynaecology (refer to **Part 2**).

 In another consumer-pull and technology-push situation, ovum retrieval has interestingly gone from a transabdominal scan - transabdominal puncture approach to a transabdominal - transvaginal approach and further onto the universally accepted transvaginal - transvaginal approach.

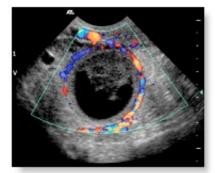
Campbell's group at Kings (with John Parsons) was among one of the earliest pioneering groups to have set up an outpatient oocyte retrieval service in IVF. In **1989** Campbell's department was the first to publish on a large scale **screening project for ovarian cancer** using abdominal ultrasound over a span of 5 years. Their results showed a low positive predictive value. Subsequent to this Paul DePriest's group in Kentucky published in 1993 results in ovarian cancer screening using the vaginal approach which arrived at similarily low postitive predictive values. Morphological scoring systems to improve the diagnosis of ovarian cancer in ultrasound-detected pelvic masses were described by several workers, notably Ilan E. Timor-Tritsch and AM Sassone in 1991. Results of studies appeared to indicate that many women would undergo surgical procedures to diagnose relatively few cancers. It has so far not been convincingly demonstrated that screening will reduce morbidity or mortality from ovarian cancer or for that matter improve the health status of women. Bengt Karlsson and Seth Granberg in Helsinki, Finland reported in 1994 the use of endometrial thickness measurement (over 5mm) to predict endometrial cancer.



Asim Kurjak

Gynecology.

The application of **doppler ultrasound** in **gynecology** did not appear until the mid 1980's when Kenneth Taylor at Yale described blood flow in the ovarian and uterine arteries in 1985 and Asim Kurjak in Croatia pioneered the use of transvaginal color doppler in the assessment of the pelvic circulation in 1989. Kurjak was the founder of the lan Donald Inter-University School of Medical Ultrasound in Dubronik, one of the largest and most important Ultrasound schools in the world. The Coatian group continued to contribute heavily to the applications of trans-vaginal color doppler in Obstetrics and



Corpus Luteum cvst diagnosed with the help of color flow doppler

It should be mentioned here that the use of transvaginal 2-D, doppler and color doppler ultrasound almost started around the same time in the late

1980s, and one finds the introduction of vaginal doppler studies almost coincided with the first reports on vaginal sonography. Work also came out from the United States from the Arthur Fleischer group in Nashville, Tennessee in the early 1990s on ovarian tumour vascularity using transvaginal color doppler. The group had around the same time published other important work on transvaginal ultrasound in gynecology. Color flow imaging of **Intra-follicular blood flow** and **impedance in the uterine arteries** during assisted reproductive cycles further added to the diagnostic capabilities of vaginal doppler ultrasound.



Arthur Fleischer

The work of Tom Bourne and the group at Kings also confirmed the usefulness of color doppler in refining the prediction of ovarian cancer in ovarian masses, and in a screening setting. Their group had also made exhaustive investigations into the use of transvaginal color doppler in the diagnosis of a variety of gynecological conditions. They have also documented the usefullness of periovulatory blood flow in ovarian and uterine arteries in the management of assisted reproductive cycles.

In Color power imaging (Power doppler), red or blue luminosity is used to indicate the power or amplitude of the blood flow signal. The process is more sensitive than color velocity imaging. The display of color from image areas with low amplitude echoes can by adjustment be inhibited and only high amplitude echoes are displayed and color coded according to their power or velocity. The

process has been called "Tissue doppler imaging" by W N McDicken in England in 1992. This was expounded in 1994 by K Miyatake and M Yamagishi in Japan in the evaluation of left ventricular wall motion. The development had come about with the availability of more powerful electronics. Further developments had also led to the degree of tumor vascularization being quantitatively estimated. The approach has received much attention from the gynecology sector in the investigation of pelvic malignancies.

Three-dimensional ultrasound comes of age. Visualization of the fetus in 3-D has always been on the minds of many investigators, including Tom Brown in Glasgow in the early 1970s, who had developed an elaborate Multiplanar scanner in 1973, under the Sonicaid Ltd®. With improvements in ultrasonic and computer technology, work on threedimensional visualization began to appear in the early 1980's. Some basic computer algorithms came from the group at Stanford (JF Brinkley, WD McCallum and others) and also from the Holm group at Gentofte, Denmark. Other work came from the domain of cardiologists where initial efforts were directed to acertaining the volume of cardiac chambers. Real-time scanner probes mounted on articulated arms were often employed where positions of the probe can be accurately determined. The principle has always been to stack successive parallel image sections together with their positional information into a computer.



Kazunori Baba at the Institute of Medical Electronics, University of Tokyo, Japan, first reported on a 3-D ultrasound system in 1984 and succeeded in obtaining 3-D fetal images by processing the raw 2-D images on a minicomputer in 1986. Their setup was reported in the Acta Obstetrica et Gynaecologica Japonica. Baba, with Kazuo





Kazunori Baba

Satoh and Shoichi Sakamoto at the Saitama Medical Center described the improved equipments in 1989 in which they used a traditional real-time convex array probe from an Aloka SSD280 scanner mounted on the positionsensing arm of a static compound scanner (Aloka M8U-10C). The images obtained were processed on elaborate computer systems (see picture with description below).



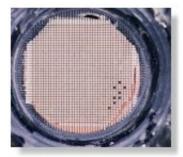
Baba's Early 3-D image of a 19 weeks fetus

This approach successfully produced 3-D images of the fetus which were nevertheless inferior to that produced on convenional 2-D scanners. At the same time, to generate each 3-D image it took on an average some 10 minutes for data input and reconstruction making the setup impractical for routine clinical use. **Baba** published in **1992** in the Japanese language the first book on ultrasonography in Obstetrics and Gynecology which contained <u>chapters on 3-D ultrasound</u>. In the mid 1990s, **Baba** collaborated with <u>ALOKA</u>® with technology developed at the Biomedical Engineering Department of the **Tokyo University**, and was a driving force in the development of commercial 3-D ultrasound technology in Japan.



Kazunori Baba's 3-D setup in the mid 1980s. A linear array probe was mounted on an articulated arm for position sensing. On the right is the computer setup for making the calculations. (click on the picture for larger view and description)

Another group at the **Columbia University** led by **Donald King** described in **1990** other approaches and computer algorithms for 3-D spatial registration and display of position and orientation of real-time ultrasound images. **HC Kuo**, **FM Chang** and **CH Wu** at the National Cheng Kung University Hospital in Taiwan, Republic of China, reported in **1992** 3-D visualization of the fetal face, cerebellum, and cervical vertebrate using a the **Combison 330** from <u>Kretztechnik</u>®, Zipf, Austria. The **Combison 330** which appeared in **1989**, was the first commercial 3-D scanner in the market. The Taiwanese group were also the first to describe 3-D visualisation of the fetal heart in the same year although at that time they were only able to image static parts in 3-D.



2-D matrix-array at Duke

In **1987**, the Center for Emerging Cardiovascular Technologies at **Duke University** started a project to develop a <u>real-time volumetric scanner</u> for imaging the heart. In **1991** they produced a <u>matrix array scanner</u> that could image cardiac structures in real-time and 3-D. In **1994**, <u>Olaf von Ramm</u>, <u>Stephen Smith</u> and their team produced an improved scanner that could provide good resolution down to 20 centimeters. The team developed state-of-the-art "Medical Ultrasound imaging" integrated circuits (**MUSIC**) which were capable of processing signals from multiple real-time phased-array images. The microprocessors were developed in collaboration with the **Volumetric Medical Imaging Inc.** at Durham, North Carolina. The MUSIC 3.2, a 40MHz



Olaf T. von Ramm

1.2µ chip completed in **1994**, was the basis for the beam-former in the world's first **electronically steered** matrix-array 3-D ultrasound imager. This became available commercially from Volumetric Medical Imaging, Inc. in **1997**.

The matrix-array transducer, which steered the ultrasound beam in three dimensions, contained 2,000 elements in which 512 were used for image formation. The beam-former produced **4,096 lines** running at **30 frames per second**. This required as much ultrasound signal processing power as eight top-end 2-D systems, running on microprocessors that execute instructions 30 times the speed of a typical 2 GHz Pentium tm. Due to the relatively small size of the 2-D matrix array probe, it is more suited to **cardiac examination** rather than for the abdomen. The apparatus is also costly to produce and poses problem in manufacturing and in image quality due to the large amount of crystals and interconnections.



Other pioneering investigators included **Ian Kelly** and **John Gardener** at the **Middlessex Hospital** in London and the <u>Sturla Eik-Nes</u> group at **Tronheim**, Norway, using equipments from **Vingmed**® which were partly developed at the University's bio-engineering department (see **Part 2**). They were able to demonstrate early gestational age fetuses with their apparatus. **Wilfried Feichtinger** at the University of Vienna, Austria reported images of 10



Professor Kratochwil from A-scan to 3-D

weeks embryos imaged with **3-D transvaginal transducers** in **1993**. Kretztechnik®. had in this year marketed their <u>2nd generation 3-D scanner</u> the <u>Voluson 530D</u>. <u>Alfred Kratochwil</u> had continued his support in the development of 3-D technology at Kretztechnic® and was active in the teaching of 3-D sonography after his retirement. The <u>Ulrike Hamper</u> group at **Johns Hopkins** reported images of various **congenital malformations** with a prototype 3-D scanner. Computation was based on a 486 computer together with a RISC processor (860/240 mhz).

Thomas Nelson and Dolores Pretorius at the University of California, San Diego, approached the carotid arteries with their prototype 3-D system in **1992** and produced very

successful images. The signal chain consisted of a transducer-array moving along the patient's neck producing sequentially sampled images which were digitised, acquired and surface-rendered on the connecting workstation. They collaborated with <u>development</u> by <u>Donal Downey</u> and <u>Aaron Fenster</u> at the <u>Imaging Research Laboratories</u> of the John Robarts Research Institute at the **University of Western Ontario**, Canada.

Their group continued to make refinements to the instrumentation and started to publish on **fetal visualization** in the following years and continuing on to become one of the **most important** research teams in the field of 3-D ultrasound in Obstetrics and Gynecology. In 1996, **Nelson's** group and the <u>Medical Imaging group</u> at the **university College Hospital in London** published independent researches on **4-D** (motion 3-D) **fetal echocardiography**, using sonographic cardiac gating methods to remove motion artefacts, which are present with conventional (static) 3-D methods. A useful feature of 3-D display is the **cine loop**, in which the rendered 3-D volumes are viewed as they rotate. This capability enhances depth perception and gives a true 3-D perspective of both normal and abnormal structures.



Dolores Pretorius



Eberhard Merz

In **1995**, <u>Eberhard Merz</u> at the Center for Diagnostic Ultrasound and Prenatal Therapy, University of Mainz, Germany, demonstrated the usefulness of multiplanar orthogonal imaging as well as as **surface**

views and **transparent views** in the diagnosis and confirmation of fetal **surface** and **skeletal** anomalies such as cleft lips and complex multiple malformations. He and his co-workers reported a large series of over 600 cases of fetal diagnosis using 3-D ultrasound. In **1997**, his team reported on the diagnosis of facial anomalies using **trans-vaginal** 3-D scans.

In Obstetrical and Gynecological 3-D imaging, mechanical

designs appeared to be the only popular choice. Twodimensional arrays are mechanically moved to provide the third dimension by sweeping or rotating, using either <u>constrained free-hand adapters</u> or an existing probe alongside with an external <u>motion-sensing system</u>. The most sucessfully deployed transducer design is the mechanically-driven arrays that is built-in into the probe housing from the Austrian manufacturer <u>Kretztechnik</u>®. Their technique was described in the paper "**3D ultrasound - the Kretztechnik Voluson approach**" in the European Journal of Ultrasound in **1994**.

The process of acquisition is microprocessor-controlled and automatic. In the display of the acquired data, the degree of transparency is first chosen which involves applying a mixture of ray-traced, volume-rendered illumination and maximum intensity or summed voxel projection. Perception in 3-D surface is acheived by a combination of depth shading, color-mapping, texture mapping and



Hand-held 3-D probe from Kretztechnik

ray-traced volume rendering. The introduction of **Multiplanar reformatting** has allowed the generation of any arbituary slice within the data aquired. In obstetrics this is valuable for measurement, and for obtaining re-constructed critical views (such as the 4-chamber view) or scans orthogonal to the face and soft palate. All these are heavily dependent of the **software algorithms** and processing power of the computers within the machines.

Volume rendering in medical imaging has in fact much of its roots in computer graphics engineering. Volume rendering developed as a separate body of techniques, mainly within the computer graphics literature, before and independent of its application to medical data. One of the earliest pioneers in volume rendering is <u>Marc Levoy</u> at the University of North Carolina (now at Stanford). Volumes are rendered directly from sampled data without first creating an intermediate surface representation. This creates images that represent the underlying data very accurately and can reveal fine details that might be obscured with surface methods. Following his initial paper in **1988: "Display of Surfaces from Volume Data"** where he described the classic volume ray tracing algorithm and has been the basis (directly or indirectly) for most commercial 3D ultrasound systems, Levoy has published a number of papers that have continued to break new ground. Volume rendering has made a major impact on the many scientific, engineering and medical disciplines that create and display large multi-dimensional datasets.

Many of the volume rendering agorithms and technology had actually originated

from computer scientists at the **filmmaking** company <u>Pixar Animation Studios</u>, famous for its 3D computer animated films! Initial volume rendering techniques and agorithms weres "invented" by company founders **Robert Drebin**, <u>Loren</u> <u>Carpenter</u>, and <u>Pat Hanrahan</u>. The algorithm embodied three key ideas: Directional shading based on the gradient in a volume, digital compositing to combine the slices of a volume, and Image warping, also applied to a volume.

Gradient shading of volumes first appeared in a 1986 paper by <u>Karl-Heinz</u> <u>Hoehne</u>, hamburg, who called it graylevel gradient shading. Compositing can be traced through **Thomas Porter** and **Tom Duff**'s 1984 paper to <u>Edwin Catmull</u> and <u>Alvy Ray Smith</u>'s invention of the alpha channel in the mid-70's. Image warping is a special case of texture mapping, which dates back to Edwin Catmull's 1974 PhD thesis at the University of Utah.

Volume rendering approximates the passage of light through a participating media. In this respect, **James Blinn**'s 1982 paper on clouds and dusty surfaces, **Jim Kajiya**'s 1984 paper on volume densities, and **Holly Rushmeier**'s 1987 paper with **Ken Torrance** on zonal radiosity must also be regarded as formative. This relationship between volume rendering and light transport was pointed out to the volume rendering community by **Wolfgang Krueger**. Another key pioneer was <u>Gabor Herman</u> at the City University of New York, who in a 1979 paper with **Ksun Kao Liu** proposed using cube-shaped voxels to display computed tomography data.^{^^}

(From citations by Dr. Marc Leroy, Stanford University. Excerpted with permission).



Cleft lip in 3-D **

<u>Medison</u>®, which had acquired <u>Kretztechnik</u>® in 1996 continued to produce more advanced versions of the <u>Voluson</u> series of scanners that produced some of the best 3-D images in the market. <u>Bernard Benoit</u> in Nice, France working in collaboration with <u>Kretztechnik</u>®, published some of the earliest, most stunning and convincing <u>3-D images</u> in the mid- 1990s using prototype **Voluson** scanners (see picture of cleft lip on the left). His pictures had been responsible for drawing the attention of many to this new scanning modality.

In the second half of the 1990's at least <u>twenty other important centers</u> worldwide were embarking on distinctive laboratory and clinical research into 3-D ultrasound, usually backed by work done at their own university's medical physics and bio-engineering departments, or commercial enterprises. Many of the innovations relied heavily on software programming. **3-D ultrasound technology** would not have been a viable commercial proposition if not for the fact that computer technology was in the mid to late 1990s at a stage where the computations can be done with **staggering speed and at an acceptable cost**.

In November **1996**, with technical assistance from Takashi Okai and Shiro Kozuma from **ALOKA®**, <u>Kazunori Baba</u> published in the Lancet their initial experience with <u>real-time processable 3-D</u>, which used a simpler algorithm compared to conventional 3-D rendering. <u>Fetal surfaces</u> are demonstrated in near real-time imaging basing on simple 'accoustic impedance thresholding' to identify fetal surfaces in the amniotic fluid. Image quality was very high and required less expensive computers to make the calculations. However the viewing direction is limited to that of the probe so that a desirable direction may not always be possible. Sufficient amniotic fluid is also prerequisite to a good scan. Similarily image clarity depends on the difference between the accoustic impedance of the fetal skin and that of amniotic fluid which made good images difficult to obtain before 20 weeks. **Aloka®** adapted the technology into their commercial scanners.



3-D image of the fetus from real-time processable 3-D

1 600



Other early manufacturers of 3-D systems included ATL®, <u>Tom-Tec Imaging Systems</u>®, GE-Vingmed®, 3D-

EchoTech®, and **Life Imaging Systems Inc.**®. The <u>first English textbook</u> on 3-D ultrasound in Obstetrics and Gynecology came out in **1996** which was edited by **Kazunori Baba** and **Davor Jurkovic**, King's College Hospital, London.

Eberhard Merz hosted the **First World Congress on 3-D Ultrasound in Obstetrics and Gynecology** in Mainz in 1997. Many important teething issues surrounding the new practice of 3-D sonography were discussed. In 1999 the <u>3D Focus group</u> was formed by the **ISUOG** to look after matters concerning the practice and education of 3-D ultrasound in Obstetrics and Gynecology.

The increasing availability of **3-D ultrasound** has resulted largely from the rapid advancement in computer technology and the decreasing

cost of micro-processor electronics. The benefits that 3-D has bought to ultrasound diagnosis has quickly become a matter for debate. The important advantages of 3-D over conventional 2-D ultrasound as it is at the turn of the millenium are its ability to enhance <u>maternal-fetal</u> <u>bonding</u>, improved comprehension of certain fetal anomalies by parents, Improved recognition and better confirmation of certain anomalies such as cleft lips, polydactyl, micrognathia, malformed ears, club foot, vertebral malformations and other anomalies appearing on the 'exterior' of the fetus, consequent to the benefit of volume and **surface rendering**. The development of **transvaginal 3-D probes** have further enhanced its value in the early diagnosis of congenital malformations.

In a recent article by <u>Asim Kurjak</u> and his team, "*Three-dimensional* sonography in prenatal diagnosis: a luxury or a necessity?" (Journal of Perinatology, issue 3, 2000), he concluded,



Reports of the use of 3D ultrasound appearing in health and parenting magazines

".... the main advantages of three-dimensional ultrasound in perinatal medicine and antenatal diagnosis include scanning in the coronal plane, improved assessment of complex anatomic structures, surface analysis of minor defects, volumetric measuring of organs, "plastic" transparent imaging of fetal skeleton, spatial presentation of blood flow arborization and, finally, storage of scanned volumes and images. It is our decided opinion that three-dimensional sonography has gained a valuable place in prenatal diagnosis, becoming a necessity for every modern perinatal unit ".



Stuart Campbell at a 3-D scan

Stuart Campbell at the St. Georges Hospital in London was one of the early proponents for the 3-D scan to be an important catalyst for mothers to bond to their babies. What are known as 're-assurance scans' and the perhaps misnamed 'entertainment scans' have started to develop. The attraction of being able to look at the face of your baby before birth was enthusiastically reported in lay parenting and health magazines. Manufacturers had adopted an unprecedented "profit marketing" strategy to advertise to providers and "reverse marketing" strategy to advertise to consumers, particularly after the arrival of the 4-D (dynamic or motion 3-D) machines.

Barbara Maier and his group (Horst Steiner, Alf Staudach etc.) in a study in Salzburg in 1996 reported that mothers are more incentive to endure pregnancy-

related difficulties, reduced anxiety, and improved capacity to cope. **Pretorius** reported in the same year that **improved bonding** between the mother and fetus could motivate mothers to refrain from smoking and other harmful behaviors during pregnancy.

Usefulness has been reported for calculating volumes of the

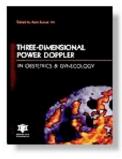




Harm-Gerd Blass

gestational sac, fetal lungs and heart from second trimester to term, placental volume, liver volume, and thigh and abdominal volume for the estimation of fetal weight. Andreas Lee with the Kratochwil group evaluated the accuracy of fetal weight estimation using 3-D abdominal and thigh volumes and reported in 1994 similar mean errors as compared to standard 2-D evaluations. Fong-Ming Chang's group at the National Cheng Kung University Hospital in Taiwan, Republic of China reported in 1997 the feasibility of predicting birth weight by upper arm volume measured in 3-D. In 1998, <u>Harm-Gerd Blaas</u> at Tronheim, Norway published <u>3-D</u> studies of embryos that were less than 10mm and further expanded the usefulness and importance of 3-D sonography as an in-vivo research tool in fetal embryology. Transvaginal 3-D work on early fetal anatomy and malformations had also come out of <u>Ilan Timor-Tritsch</u>'s center in New York. 3-D power doppler in the study of placental blood circulation was first described in

1996 by **C J Ritchie** in a proprietary setup at the Bioengineering Department of the University of Washington, Seattle. **Dolores Pretorius** published on its usefulness and techniques in **1998**. The Croatian group led by <u>Asim Kurjak</u> and **Sanja Kupesic** expounded this new diagnostic entity further. Their book "*Three-Dimensional Power Doppler in Obstetrics and Gynecology*" was published in **2000**.



In gynecological applications, **Davor Jurkovic** at Kings convincingly demonstrated in **1995** the usefulness of 3-D ultrasound in accurately differentiating uterine anomalies such as <u>bicornuate uterus</u> and septate uteri. Similarily the assessment of the endometrial cavity with **3-D sonohysterography** and characterization of endometrial masses, adhesions, tubo-ovarian masses, hydrosalpinges, ovarian cysts, small intraovarian tumors and mullerian anomalies have all been quickly and convincingly demonstrated. Diagnostic accuracy of malignant ovarian tumors can be up to 100 percent as reported by **Bonilla-Musoles** in 1995, who also demonstrated the value of 3-D examination over the convention 2-D transvaginal scans and 2-D sonohysterography in the diagnosis of **endometrial lesions**. **3-D color power doppler** is valuable



Davor Jurkovic

for visualization of intra-tumor flow and thus is useful in evaluating in particular cervical carcinomas and ovarian carcinomas. It is envisaged that the investigation will lead to greater appreciation of <u>tumor</u> <u>angiogenesis</u>. Another potential benefit of 3-D ultrasound lies in data documentation, storage, and networking. Digitally

saved volumes of patient data can be readily transferred to a remote site for interpretation or second-opinion consultation. How much these all add up to make 3-D ultrasound <u>cost-effective</u> and an indispensible tool in Obstetrics and Gynecology will remain to be seen.

Epilogue

The evolution of diagnostic ultrasonography has been the combined efforts of physicists, mechanical, electrical and bio-medical engineers, computer technologists, clincians, sonographers, researchers, university and government administrators as well as adventurous and perceptive commercial enterprises.

I particularly salute the **ingenious engineers** and **physicists** throughout the history of the development as they are the true heroes behind the entire 'scientific' advance in ultrasonic imaging.



Without them the innovative ideas of the brightest clinicians cannot be put into action. Developments in echocardiography, neurosonography, ophthalmology and breast echography have all supplemented the advancement in ultrasound instrumentations and methodologies in Obstetrical and Gynecological sonography. The first linear-arrays for example were invented for the purpose of ophthalmologic and cardiac investigations. Ultrasonography has very quickly become the single most important diagnostic investigation in the field of Obstetrics and the **healthcare for women**.

Interestingly, diagnostic medical ultrasound had evolved from technology used in mapping waves through liquid (the **sonar**), through air (the **radar**) and through solids (the **metal-flaw detector**). The **A-scan** which had evolved from the sonar and early metal-flaw detectors would not have a lasting impact on clinical medicine without evolving into the **B-scan** which had it's origin in the military radar. The A-scan did not provide sufficiently accurate, reproducible and interpretable information to allow a firm diagnosis to be made, particularly in Gynecology. The bistable B-scan would not have advanced to become a respectable diagnostic tool as it is now, without the development of the **scan-converter** and **gray-scaling**. The gray scale **compound static scanner**, with the incorporation of progressive electronic and computer technology available in the late 1970s had establish itself as a genuine stand-alone clinical diagnostic tool, providing hitherto unavailable information to the clinician regarding a particular disease condition. **Howry's** original concept of deriving clear outline anatomical pictures by selectively recording larger echoes from major interfaces and suppressing any other small echoes was completely reversed in later developments, where attempts are made to detect the smallest echoes in the presence of noise and displaying them in finer spatial detail and echo amplitudes.

The arrival of the **real-time scanners** have added further impetus to ultrasound techniques and had established ultrasonography as **the most important imaging modality** in Obstetrics and Gynecology.

The concept of the **transvaginal scanner** was in situ in the early 1950's but was unable to make any real headway until the appearance of sophisticated mechanical and electronic sectoral real-time vaginal scanners in the mid 1980's. Ovum retrieval has for example, interestingly gone from a transabdominal scan - transabdominal puncture approach to a transabdominal - transvaginal approach and further onto the universally accepted transvaginal - transvaginal approach.

A **'technology push'** situation further evolved when enhancement in diagnostic capabilities of scanners was propelled by the almost explosive advancements in electronic and microprocessor technology, occurring most significantly in the 1980s and 90s. The advent of ultrasonography in Obstetrics has also 'created' the new specialty called **Prenatal diagnosis** that has developed by leaps and bounds since it's early conception. Ultrasound has markedly enhanced and pushed forward the study of congenital abnormalities among obstetricians, pediatricians, geneticist, pathologist and other allied specialties. All of a sudden, obstetricians started to learn about so many congenital malformations that they have not even heard of.

On the other hand, every single measurable parts of the fetus has been measured and their changes throughout gestation documented. It is of interest to note that historically the 4 basic fetal measurements, namely the BPD, the CRL, the AC and the FL had evolved successively at different time periods (' 62/' 68, ' 73, ' 75, ' 80 respectively) each being brought on by technical developments in ultrasound instrumentations at that time (B-scan, gray-scale, real-time). Stuart Campbell published the first BPD chart in 1971. Since then, charts and tables had become an important and integral part of Obstetric practice, at which Obstetricians and Gynecologists were slowly getting used to.

Fetal malformations were diagnosed with increasing accuracy and at an earlier gestation. Bold, daring and visionary clinicians and researchers invented new interventional techniques that work under the guidance of ultrasound to diagnose fetal disease and gynecological conditions. Doppler devices moved progressively from depicting flow velocity waveforms to color flow mapping, power doppler and doppler tissue imaging. Velocimetric parameters of the umbilical artery, the middle cerebral artery and the ductus venosus had made their appearance one after another again subsequent to progressive developments in the imaging apparatus. 3-D ultrasound made the scene in the late 1980's and further revolutionarised sonography in obstetrics and gynecology.

It is regretable that John Wild's original conception of precise quantitative detection of cancer echoes with ultrasound also had not materialize to the initial expectations. So was the application of tissue characterization in the specialty. On the other hand, all the improvements in machinery and earlier detection of abnormal structures in the fetus have nevertheless brought along with it "false positives" and difficult-to-be-sure diagnosis that could generate much undue anxiety in patients. Such could only have far-reaching effects on a woman's perception of child-bearing.

I am still finding it awsome to imagine the "wire-frame" images of Douglass Howry and lan Donald could have now become "photo-realistic", and between all these are the unfolding magnificence of the invention and genius of science and man's endeavour to find-out and to perfect. From the detection of life to the measuring of fetal sizes; from the determination of morphological normality to the evaluation of circulatory and growth dynamics, all have been making profound changes to the entire concept of routine antenatal care and Obstetric practice.

J. W.

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^A In Diagnosis and Therapy of Fetal Anomalies. John C Hobbins and Beryl R Benacerraf (eds), Churchill Livingstone, 1989.
 ^A Excepted with permission from the homepage of <u>Dr. Marc Leroy</u>, Stanford University: <u>Citation; SIGGRAPH 1996 Computer Graphics</u>

Achievement Award - It is not possible to include all the names who have contributed significantly to the advancement of Obstetrical and Gynecological sonography, some who may have been less well-known than the others and some who may not have published so extensively in the English language. Apologies are extended to those whose contribution has not been fully credited in this article.

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