AUTOMATED ANALYSIS OF EEG PATTERNS IN SUBJECTS UNDER ABUSIVE LEVELS OF SEDATIVE-HYPNOTICS

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by

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CHAPTER I

INTRODUCTION AND REVIEW OF THE LITERATURE

1.1 INTRODUCTION

The pionering work of Aserinsky and Kleitman (1953, 1955) has provided sleep researchers with better understanding of the fundamentals of nocturnal sleep. They observed that periodically throughout the night, rapid eye movements (REM) occured in association with low voltage mixed frequency EEG activity. The REM periods have been shown to occur about every 90 minutes in normal individuals and may last for 15-30 minutes. These periods of REM alternate with the other four stages (1,2,3 and 4) of sleep, that have been collectively called non-REM (NREM) sleep (Rechtschaffen and Kales, 1968).

Most of the studies on the effects of drugs on human sleep have been evaluated using the system of sleep staging (Rechtschaffen and Kales, 1968). One whole night's sleep in the sleep laboratory results on the average in eight hours of continuous EEG-EOG recording. This data is divided into hundreds of one minute epochs. An Electroencephalographer has to score each of these epochs in one of six possible stages (0,1,2,3,4 and REM) of sleep (Rechtschaffen and Kales, 1968). In the case of drug studies, if a drug decreases one sleep stage, it must unavoidably increase one or more other stages. The issue then revolves around whether a given drug effect on the sleep patterns is a direct drug effect on the EEGs or whether it is an "artifact" due to this system of scoring.. It should be kept in mind that accepted definitions of sleep stages have been derived from drug-free sleep, and that the drugs can produce patterns of effects on the sleep EEG that do not fit the normal definitions (Kay et al, 1972).

Some researchers have suggested alternative solutions to this problem. They have proposed attempts at quantification of drug effects on the EEG signals themselves instead of looking for changes conventionally measured by sleep researchers (Kay et al, 1972; Karacan et al, 1973; Smith and Karacan, 1973; Saito, 1974; Karacan et al, 1975; Kay, 1976).

The data utilized in this work was originally obtained by the sleep research group of the Department of Psychiatry of the Baylor College of Medicine in Houston, Texas, for a project designed to study sleep-related growth hormone release patterns in: (a) patients being withdrawn from chronic and abusive use of secobarbital and methaqualone; (b) non-drug users that were given acute toxic dosages of these The main reason why these two drugs were chosen is drugs. because Houston is one of the top four areas in the nation where barbiturates and methaqualone are widely used. In the present study we try to quantify the effects of these two drugs on the sleep EEG of the above subjects by analyzing the sleep EEG waveforms with statistical and spectral analysis techniques. To date, we have not found any other studies that use these techniques to study the effects of

abusive or therapeutic levels of these two agents on sleep EEG patterns.

1.2 EFFECT OF SECOBARBITAL AND METHAQUALONE ON HUMAN SLEEP

Secobarbital and methaqualone can be classified as sedative-hypnotics. The common characteristic these sedative hypnotics have is their depressant effects on the CNS, with sedation at lower doses and hypnosis at higher doses (Kay, 1976).

Secobarbital belongs to that group of barbiturates which are derivatives of malonylurea or barbituric acid. The first hypnotic barbiturate, barbital, was introduced into medicine by Fisher and von Mering in 1903 (Kay, 1972). More than a billion hypnotic doses of barbiturates are produced each year in the United States alone, and numerous cases of addiction and death due to acute poisoning have been reported (Maynert, 1971).

The barbiturates are general depressants. They depress the activity of nerve, skeletal muscle, smooth muscle and cardiac muscle. These effects require large concentrations, generally higher than those necessary to produce CNS depression. However, it should be emphasized that the CNS is exquisitely sensitive to barbiturates, so that, when the drugs are given in sedative or hypnotic doses, direct action on peripheral structures is absent or negligible (Sharpless, 1970).

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Secobarbital is marketed by Lilly Laboratories under the trade name of Seconal. The recommended hypnotic dosage for adults is 100 mg administered at bedtime (Sharpless, 1970). Secobarbital, as well as most of the barbiturates, is known to reduce REM sleep. (Lester, 1960; Lester et al, 1968; Allnutt and O'Conner, 1971; Feinberg et al, 1974; Kales and Bixler, 1975). Sleep spindles are increased by secobarbital (Allnutt and O'Conner, 1971; Feinberg et al, 1974), however, Lester et al (1968) found an opposite effect. Delta sleep is not significantly affected by barbiturates (Kay, 1972, 1973, 1976). Lester et al (1968) reported an increase in beta activity. All the above investigations were short-term studies, and the amount of drug given to the subjects was either 100 or 200 mg of secobarbital. The amount of barbiturates required to produce physical dependence in man has been carefully studied (Fraser et al, 1958). They reported that in adult males consumption of up to 400 mg of secobarbital daily does not usually create a significant degree of physical dependence. Thus, the minimal addictive dose of secobarbital would be more than 400 mg daily.

Methaqualone is a sedative and hypnotic agent, chemically unrelated to other sedative-hypnotics. Its hypnotic activity was demonstrated by Gujral and his coworkers in 1955; however, the exact mode of its sedative-hypnotic action is not known yet (Sharpless, 1970). The recommended hypnotic dosage is 150-300 mg. Methaqualone is marketed under the trade name of Quaalude by Rorer Laboratories.

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Physical dependence in methaqualone has reached alarming proportions and some authorities on drug abuse believe that the potential for addiction and abuse for this drug are as serious as those for barbiturates in general, and for heroin (Swinyard, 1975).

The reported effects of methaqualone on the human sleep are comparable to those of the barbiturates (Kay et al, 1973). A decrease in REM sleep was found by Kales et al (1970), Rechtschaffen et al (1970) and Risberg et al (1975). However, Goldstein et al (1970) found that methaqualone did not affect REM sleep. An increase in spindle activity was reported by Goldstein et al(1970) and Risberg et al (1975). However, Kales et al (1970) did not find any significant change in the spindle activity. A decrease in stage 4 sleep was observed by Goldstein et al (1970). Itil et al (1974) found that 300 mg of methaqualone decreased spindle sleep and stage four sleep. All the above works were short-term studies and the amount of methaqualone administered to the subjects was either 150, 250 or 300 mg.

1.3 USE OF SPECTRAL ANALYSIS TO STUDY THE EEG SIGNALS DURING SLEEP

The spectrum of single EEG records has already become familiar in EEG work through the various types of analog frequency analyzers (Grass and Gibbs, 1938; Knott et al, 1942; Walter, 1943; and others).

In the last two decades, the methods of statistical

time series analysis have become of considerable importance in processing large amounts of random data (Blackman and Tukey, 1958; Parzen, 1964; Jones, 1965; Bendat and Piersol, 1966,1971; Jenkins and Watts, 1968; Otnes and Enochson, 1972). Of these methods, spectral analysis has found several applications in electroencephalography. The power spectrum is a representation of the mean square value, i.e. the variance of zero-mean EEG, as a function of frequency. Walter et al (1963) applied the methods suggested by Blackman and Tukey (1958) to compute the power spectrum of single EEG records. Other researchers applied similar methods (Walter and Adey, 1965; Hord et al, 1965; Dummermuth and Flühler, 1967; Dummermuth et al, 1970; Joy et al, 1971). The work of Hord et al (1965) was primarily concerned with the resolution and stability of the EEG power spectrum in the four stages of sleep. Johnson et al (1969) used spectral analysis to detect frequencies that are common to all stages as well as those unique to a specific stage of sleep. Dummermuth et al (1972) made a study of the different sleep stages in normal adults using spectral analysis techniques. Hagne et al (1973) used spectral analysis to study the waking EEG in normal infants. Bronzino et al (1973) investigated the possibility of using spectral "fingerprints" to distinguish between waking, slow wave sleep (SWS) and REM stages in the cat.

The use of the spectral components of the EEG to automatically classify the stages of sleep has been pursued by Lubin et al (1969), Larsen and Walter (1969, 1970), Martin et al (1972) and Larsen (1975). Evaluation of these automatic techniques revealed the delta frequency band as the most useful discriminant with alpha and sigma bands next in importance. Lubin et al (1969) divided the EEG power spectra in five bands and a linear-discriminator for all sleep stages was computed using stepwise multiple regression techniques. The overall percentage of agreement with the human scorer was very poor. However, better results in classifying some stages were obtained by Larsen and Walter (1969) who divided the EEG power spectra in bands and then used a multiple discriminant analysis for the classification of the different sleep stages. Martin et al (1972) used, in addition to the spectral components, EEG amplitude information and REM presence in order to discriminate between the sleep stages. Lubin et al (1969) conclude that detectors for such phasic events as eye movement and k-complexes will have to be added to the frequency analysis to bring the performance of any system close to that of human scorers.

1.4 USE OF SPECTRAL ANALYSIS TO STUDY THE EFFECTS OF DRUGS ON THE EEG SIGNALS

Very few researchers have used spectral analysis techniques to evaluate the effects of drugs upon the EEG signals. Most of the studies have been done by using the classical sleep staging type of approach. An excellent review of the works applying this classical approach is

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presented by Kay (1973, 1976).

Schalleck et al (1968) used power spectral analysis to study the effects of some drugs (chlorpromazine, diazepam, pentobarbital, and methyprylon) on the electrocorticogram (ECoG) of the cat. They made comparisons between the power spectra before and after the administration of a particular drug. Itil et al (1968) used a combination of period analysis and analog power spectral analysis to study the effects of three psychotropic compounds (chlorpromazine, imipramine and chlordiazepoxide) on human subjects.

Joy et al (1971a) applied spectral analysis to study the effects of diazepam and atropine on the ECoG of the They presented a three dimensional representation monkey. of the dynamic behavior of the EEG spetra during control and after the administration of one of the drugs. This type of representation of the EEG spectra was later on called Compressed Spectral Array (CSA) by Bickford et al (1972). Furthermore, this kind of plotting was used by Singleton and Poulter (1967) to plot the spectra of the sound of the male killer whale call. Joy et al (1971b), utilizing their previous type of approach, studied the effects of several types of benzodiazepines on the ECoG of primates. Schneider (1974) used the CSA to represent the dynamic behavior of the EEG spectra during halothane anesthesia in the dog. Killam and Killam (1975) used the same spectral analysis techniques of Joy et al (1971) to study opiate dependence in primates. Kay (1975) applied spectral and period analysis to study

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human EEG through a cycle of methadone dependence. Analog frequency analysis has been used by Saito (1974) to study the effects of psychotropic drugs on the human EEG.

Caille (1974) studied the effects of psychotropic drugs on the human sleep, using power spectral analysis. A special purpose computer was developed in order to analyze the data in real time. The emphasis of this study was in the REM and NREM periodicity.

1.5 USE OF OTHER AUTOMATED TECHNIQUES TO STUDY THE EFFECTS OF DRUGS ON THE EEG SIGNALS

Another time series analysis method that has been used in electroencephalography is period analysis (Saltzberg and Burch, 1959; Burch and Childers, 1961; Burch et al, 1964).

Fink et al (1967) applied period analysis to study the changes in the human EEG due to the administration of pentothal at different rates. Itil et al (1974) classified psychotropic drugs based on digital computer sleep prints. This method uses parameters derived from period analysis for the classification of the EEG sleep stages.

An automatic hybrid system used for EEG sleep staging (Smith and Karacan, 1971) was applied to the quantification of the effects of a hypnotic-like drug on slow-wave sleep (Smith and Karacan, 1973). In this study the effects of the drug on the amplitude, frequency and amount of delta waves (stages 3 and 4) were quantified. The various digital computer techniques that have been applied to the quantification of the EEG and to the automatic recognition of the sleep stages have been reviewed by Cox et al (1972) and by Gevins et al (1975).

1.6 SCOPE OF THIS STUDY

In the present work we use statistical techniques, including spectral analysis, which are more sensitive to the variations of the EEG waveforms, than the classical approach of sleep staging, in order to test their applicability to detect the possible effects of the two drugs (seconal and quaalude) on the human EEG during an entire night's sleep. The reasons we chose spectral analysis instead of other time series techniques are: (1) the noise nature of the signals in consideration which make this approach plausible; (2) its easy implementation with the facilities at hand; (3) the possibility of using the optics facilities in our Department to compute the EEG power spectra, in which case the process will be much faster and cheaper.

Our techniques were tested in two groups of six subjects each. The first group is formed of the drug addicts that were withdrawn from secobarbital and methaqualone (three subjects in each drug). The second group is composed of non-drug users that were given toxic doses of these drugs (again there are three subjects for each drug). As we mentioned in Section 1.1, these are the subjects that have participated in an early sleep-growth hormone release study conducted by the sleep research group of the Department of Psychiatry of the Baylor College of Medicine. For each subject four nights of sleep EEG were processed.

Due to the large amount of data to be processed, several algorithms were developed in order to handle the problem efficiently. To reduce computational time, the power spectrum of a 30 sec epoch was computed at intervals of 2 1/2 minutes throughout the night. This approach showed to be more than satisfactory for the purpose of this study. In order to get meaningful power spectra, the epochs must be completely or nearly artifact free. To accomplish this, most of the researchers using spectral analysis usually select visually artifact free epochs (Hord et al, 1965; Johnson et al, 1969; Larsen and Walter, 1969; Lubin et al, 1969; Dummermuth et al, 1972, Bronzino et al, 1973; Hagne et al, 1973). We solved this problem by developing a special purpose artifact detection algorithm based on a chi-square goodness-of-fit test to a Gaussian distribution (see Chapter II).

The algorithm that computes the power spectrum of each of the epochs, was extensively evaluated using band limited white noise as a test signal to check the statistical stability of the resultant spectral estimates. After the power spectrum of each of the chosen epochs is computed, they are plotted in a form of a Compressed Spectral Array, which emphasizes the dynamic shift of the activity in the frequency bands during the night. The results of the spectral analysis are further processed by dividing the power spectrum of each epoch in seven frequency bands. For each of these bands, it is possible to plot the percentage of power (expressed as the power in the band divided by the total power for the epoch), the spectral peaks, and its corresponding frequency as a function of the time of the night. Besides this, the mean frequency in each epoch and the results of the chisquare test can be plotted for the entire night. The above described data reduction techniques seem to be quite efficient in showing drug related patterns in the sleep EEG. These techniques form a statistical package that can be used efficiently on a dedicated general purpose digital computer for further studies on EEG patterns quantification.

With the implemented system, about 45 minutes are required to process an entire night of sleep EEG (based on an average of eight hours of EEG data), which is much faster than the two or three hours required by an experienced Electroencephalographer to score the same night.

This dissertation is divided in five chapters. The first serves as an introduction and gives a historical background on the techniques used to quantify the effects of drugs on the EEG. In Chapter II the subjects and the methods are discussed. Also an algorithm for data reduction and detection of EEG artifacts is developed. In Chapter III the basic theory of spectral analysis is reviewed, and the approach used to compute the power spectrum is presented. An algorithm to extract the main features from the power spectrum is also presented. The results for each of the subjects and the comparisons between them are the objective of Chapter IV. In Chapter V some conclusions are drawn and some suggestions are made for further research on the subject.

CHAPTER II

DATA ACQUISITION AND PREPROCESSING

2.1 SUBJECTS

Two groups of subjects were utilized in this study. One consisted of hypnotic drug abusers and was called Group 1. The other group was composed of non-drug users (Group 2).

The subjects in Group 1 were five males, 16-35 year of age, and one 25-year-old female. They were patients in a local Poly Drug Treatment Unit at the Center Pavilion Hospital in Houston. Three of the patients were taking secobarbital and the other three were taking methaqualone.

The subjects in Group 2 were six males, 22-26 years of age. They were medical students and staff members of the Center Pavilion Hospital in Houston. All subjects were in good health.

2.2 METHODS

2.2.1 Group 1

On admission to the Poly Drug Treatment Unit, the patients gave a history of multiple drug abuse and were evaluated for signs of drug dependence. If such signs appeared, the patients underwent a titration procedure to establish their drug tolerance levels. Beginning in the morning, each patient was administered hourly therapeutic doses of the primary drug of abuse until neurological and vital signs of toxicity appeared. The total amount of drug administered during this titration procedure was taken as the starting dosage for the withdrawal process. On the following days, the total drug intake was reduced by approximately one therapeutic dose per day until zero drug intake levels were reached. Two factors could modify this scheduled step-by-step reduction in the daily drug intake: (1) occasionally the patient's clinical status necessitated a slower or faster than scheduled reduction or even some backtracking on the schedule; (2) some patients declined to take one or more scheduled doses and thus accelerated the withdrawal process. The length of the withdrawal process differed from patient to patient due to these schedule changes and because the patients began the withdrawal process with different initial doses.

Tables 2.1 and 2.2 show the age and sex of each patient, their withdrawal drug, their daily drug intake throughout the treatment, and the nights for which the EEG was automatically processed for this study. The tables also show the nights when the EEG was recorded after the subject had left the hospital and the nights when either the EEG was not recorded on magnetic tape or the tapes were not suitable for playback. The night records used in this study were recorded on various occasions: (1) either on the titration night or on the night following it; (2) on one or two nights while the subject was taking the drug; (3) on the night following the first day of zero drug intake; (4) on a night

TABLE 2.1

GROUP 1 - SECOBARBITAL

·····							
	Subject SDl	Sex Age M 21	Subject SD2	Sex Age M 16	Subject SD3	Sex Age M 17	
Nights	Record Number	Daily Dosage (mg)	Record Number	Daily Dosage (mg)	Record Number	Daily Dosage (mg)	
lst	15305	100	15012 ^N	500	15030*	350	
2nd	15310	500	15017*	300	15036	150	
3rd	15315	400	15021	250	15044*	200	
4th	15320*	500	15025	200	15051	100	
5th	15322*	1000	15029*	150	15058	50	
6th	15326	400	15037	0	15066	0	
7th	15329	500	15043*	0	15071 ^N	0	
8th	15332	400	15052	0	15079*	0	
9th	15337	300	15060	0	15087	0	
10th	15340*	200	15065	0	15092	0	
llth	15344	100	15072	0	15097	0	
12th	15352	0	15080	0	15101	0	
13th	15358*	0	15086*	0	15105*	0	
14th	15364	0 15091 0		-	-		
15th	15370 0		-	-	-		
l6th	15375*	75* 0		-	-		
17th	15383	0	-	-	-	-	
18th	15390	0	-	-	-	-	
						1	

*Night records used in this study.

^NEither the record was not on magnetic tape or the tape was bad.

TABLE 2.2

GROUP 1 - METHAQUALONE

	·			ŧ					
	SUBJECT SD4	SEX F	AGE 25	SUBJECT SD5	SEX M	AGE 35	SUBJECT SD6	SEX M	AGE 16
	Record	Daily		Record	Daily		Record	Daily	
Nights	Number	Dosage (mg)		Number	Dosage (mg)		Number	Dosage (mg)	
lst	14497*	2100		14498*	3000		15336	0	
2nd	15002 ^N	2400		15003	2100		15341*	600	
3rd	15006*	1500		15007*	1875		15351*	300	
4th	15010	1500		15011	1800		15359*	0	
5th	15015	1200		15016*	1650		15363 ^N	0	
6th	15019	900		15020	1350		15368	0	
7th	15024*	600		15026	1350		15376*	0	
8th	15032	300		15031	1200		15382 ^N	0	
9th	15038	150		15039	1050		15389 ^N	0	
10th	15045	0		15046	600		15391 ^N	0	
llth	15050*	0		15053	600		-	-	
12th	15059 ^N	0		15061	300		-	-	
13th '	15064 ^N	0		15063*	0		-	-	
14th	15073 ^N	0		15074 ^N	0		-	·	
15th	15078+	0		15081 ^N	0		-	_ [
l6th	15085+	0		15142 ⁺ 0				-	
17th	-	-		15148+	0		-	-	
18th	-	-		15155+	0		-		
									1

*Night records used in this study.

⁺Recorded after the patient left the hospital.

^NEither the record was not on magnetic tape or the tape was bad.
close to the patient's discharge from the hospital.

The recordings of the sleep EEG were made with a GRASS Model 78 polygraph. Three pair of electrodes were placed in accordance with the 10-20 international system, F_1-F_7 , P_1-T_5 , $0_3-0_2P_2$ (between 0_2 and P_2). In addition, two pairs of electrodes were used for recording EOG signals and one pair for monitoring the heart rate. Electrodes placement were as described by Williams et al (1964 and 1974). The amplifiers were set to cutoff frequencies (half amplitudes) at 1 and 100 Hz. All EEG channels plus a timer channel were recorded on paper at a speed of 15mm/sec and on magnetic tape using a SANGAMO Model 35100 14 track FM tape recorder, at a speed of 15/16 ips. Each night the polygraph was calibrated to make a 5mm pen deflection corresponding to 50 μ V. In addition to this, a 10 Hz sine wave with an amplitude of 50 µV peak-to-peak was also applied to each channel as a calibration signal.

2.2.2 Group 2

In this group the study consisted of two periods: the first period was to investiagte the effects of acute toxic levels of the drugs on the sleep EEG; the second period was designed to get baseline data under normal circumstances.

In the first period, the subjects were admitted to the hospital and stayed there for at least three consecutive nights. All of them were given careful physical examina-

	SUBJECT SC3	SEX M	AGE 24	SUBJECT SC2	SEX M	AGE 23	SUBJECT SCl	SEX M	AGE 22
Sleep	Record	Daily Dosage		Record	Dail Dosa (mg	y ge)	Record	Daily Dosage (mg)	
		(9	/	Nullaser	(1119	/	Number	1110	
lst	15968	none		15943	none		15978	none	
Nap	15971*	1000 ¹		15945*	600 ¹		15983*	600 ¹	
2nd	15971*	-		15952*	-		15989*	-	
3rd	159 7 9*	none		15956*	none		15994*	none	
Baseline Night	line 16326* none		e	16328*	none		16093*	none	

TABLE 2.3 GROUP 2 - SECOBARBITAL

*Records that were automatically processed for this study. ¹The drug was given before the nap.

TABLE 2.4 GROUP 2 - METHAQUALONE

	SUBJECT SC4	SEX M	AGE 25	SUBJECT SC5	SEX M	AGE 23	SUBJECT SC6	SEX M	AGE 22
Sleep	Record	Daily Dosage		Record	Daily Dosage		Record	Daily Dosage	
Sessions	Sessions Number (mg)		Number	(mg)		Number	(mg)		
lst	15908	none		15923	none		15982	none	
Nap	15909*	4200 ¹		15928*	3000 ¹		15984*	1800 ¹	
2nd	15912*	-		15928*	-		15990*	-	
3rd	15917*	none		15935 ^N	none		15995*	none	
Baseline Night	eline 16070* none ht		e	16090*	none		16339*	none	

*Records that were automatically processed for this study. ^NRecords not available on magnetic tape. ¹The drug was given before tha nap. tions prior to the beginning of the experiment. In the first day they ate regular food, retired to bed at their usual bedtime, were awakened the next morning at their usual aris-This first night was an adjustment night for the ing time. subjects. In the morning of the second day they were fed liquid food, and after the application of the EEG electrodes they were given an initial dosage of 600mg of either secobarbital or methaqualone. Each subject was then checked for neurological and vital signs of toxicity. If these signals were absent, and additional dosage of 200mg of the drug was administered hourly, until signs of toxicity appeared. After this titration procedure, if the subject desired they could take a nap. Following the nap, they retired to bed between 0:00 and 1:00 AM and were awakened, after sleeping for their normal length of time. Two of the subjects slept straight from the beginning of the nap until the next morning. On the third night the subjects retired to bed at their usual time and were awakened at their usual arising time.

Approximately one month after the first period, the subjects reported to the sleep laboratory on four consecutive nights for the recording of the baseline EEG data. Their retiring and arising time were adjusted to their usual schedule. The EEG from the third night of sleep was used in this study as the baseline for each subject.

Tables 2.3 and 2.4 show the age and sex of each subject, the drug and dosage they were given and the nights

for which the EEG's were automatically processed for this study. The tables also show the nights where the sleep EEG's were not recorded on magnetic tape.

The recording procedures for the sleep EEG's were the same as those used in the Group 1, described in Section 2.2.1.

2.3 DATA ACQUISITION

2.3.1 Data Preparation

The recorded raw EEG data were played off-line from a SANGAMO Model 35100, 14 track FM tape recorder into an SS-100 Analog Computer through a bandpass filter. The playback speed was 30 ips which is 32 times faster than the speed used in the recording stage. This procedure results in a time-compression of the same order, which means that an eight hours record can be digitized in 15 minutes. For simplicity, filter bandwidths, sampling frequency and signal durations are given in terms of the equivalent real time signals.

In order to reduce the computational time needed, this study was based on data from the central derivation (P_1-T_5) only. In the central channel spindle activity is more predominant than in the frontal and occipital channels. The central channel also shows activity that is more predominant in the other two channels such as: delta in the frontal and alpha in the occipital. (Smith et al, 1976)

The reproduced raw EEG signal from (P1-T5) was passed through a KROHNHITE Bandpass Filter with an adjustable bandwidth of 0.2-40 Hz and 18 dB/octave roll-off. The upper cutoff frequency was chosen, so that we could investigate any activity above 30 Hz that could be present due to the effects of the durgs. The filtered signal was fed into a hybrid system comprised of a SS-100 Analog Computer (Hybrid Systems Inc.) and an IBM 360/44 Digital Computer. In the Analog Computer the signal was amplified by two operational amplifiers. The first amplifier had a fixed gain of 10 and the second one was of a variable gain. With this set-up the total gain was adjusted, for each record, so that a $50\mu V$ calibration signal was equivalent to a 30 volts deflection at the input of the A/D multiplexer. Since the SS-100 Analog Computer permits output swings of ±100V we can have EEG signals up to 300µV peak-to-peak with no distortion through the system.

2.3.2 Analog-to-Digital Conversion

Conversion of a continuous analog signal into an equivalent digitized form involves sampling in the time domain and quantization in the amplitude domain.

The sampling rate was chosen to be 128 samples/sec which means that we can represent signals up to 64 Hz without any aliasing (Bergland, 1969). In this way 60 Hz from the power line, which is a common source of noise in EEG signals, will not be aliased into lower frequency bands.

In the quantization process the amplitude of each data sample must be expressed by some fixed number of digits. In this way, no matter how many quantization levels are available, there will always be an error associated with this process, called quantization error. The quantization error is related to the value of a quantization level q and can be given as the ratio of the exact magnitude of the sampled value x, and its quantized approximation expressed as a power of 2,

$$e_{q} = \pm (x/2) \cdot (1/2^{N}) = \pm q/2$$

where N is the number of bits that make up the digital number. For ideal conversion ε_q will have a uniform probability distribution with a standard deviation of 0.29q (Beauchamp, 1973). We can think of this quantization error as a white noise added to the signal. The SS-100 Analog Computer can quantize a signal with 8192 level units (i.e. 2^{13} or 13 bits), then the peak signal to noise ratio is given by,

$$(S/N) = 20\log(8192q)/(0.29q) = 89 \text{ dB}$$

which is more than satisfactory for the present application.

2.3.3 Data Conversion and Storage

A FORTRAN program called CONVERSION was written in order to perform the following functions:

(1) to signal the operator when the HybridSystem is ready to start the A/D conversionprocess;

(2) to permit the operator to start the A/D conversion at any desired segment of the analog tape, by pressing a switch, which starts the clock of the analog computer. This clock controls the sampling rate, and thus starts the A/D conversion; (3) to convert the analog sampled data into binary integer numbers, by the use of the A/D system subroutines; (4) to write those integer numbers on a 9-track magnetic tape (SCRATCH) for tem-

porary storage.

A detailed listing of CONVERSION appears in Appendix B.

The digitized data from SCRATCH is copied into a standard magnetic tape labeled Master Tape for permanent storage.

The integer data is written on the magnetic tape in blocks of 7680 bytes, corresponding to a 30 seconds epoch. Since the tape driver can write at a density of 800 bytes/in, the length l_r of each record on the tape is,

 $\ell_r = 7680/800 = 9.6$ "

Since the interecord gap (IRG) is 0.6", the total length of each record l_+ is

$$l_t = 9.6"+0.6" = 10.2"$$

For a night of sleep of eight hours we will have 8h/30sec = 960 records which will occupy a total tape length of

On the average then we can expect to accomodate two nights of sampled sleep EEG in each 2400 ft digital tape.

2.4 DATA PREPROCESSING

2.4.1 Artifacts and Data Reduction

The first big problem that we faced when we tried to automatically process the all-night sleep EEG signals was that of artifacts in the EEG record. Artifacts can be caused by several sources such as: loose electrodes, body movement, contamination of scalp electrodes by eye movements, etc. Most of the reported work using spectral techniques to analyze EEG signals has been conducted using artifact free epochs selected visually. Since in the present study we processed about 50 nights of sleep EEGs, the task of selecting artifact free epochs by visual inspection would have been practically impossible. At the beginning of this study, we tried to detect artifacts using an amplitude level criterion, due to the fact that artifacts usually have high amplitudes. With this method we ended up rejecting epochs with high amplitude delta activity and accepting other epochs with low amplitude artifacts.

One approach that worked much better than the amplitude criterion was a chi-square goodness-of-fit test to a Gaussian distribution (CSQ) to check for artifacts. This test is very sensitive to nonstationarities in the epochs and it will produce a large chi-square coefficient, X^2 , when an artifact is present. This artifact detection method will be explained in more details in Section 2.4.3.

Another big problem we faced was the huge amount of data to be processed. In order to minimize this problem, we decided not to process the epochs continuously during the night, but instead to pick one 30 second epoch each 2 1/2 minutes. Since our main interest was not in the automatic scoring of the sleep stages, but in studying possible patterns caused by the effects of the hypnotic drugs on the sleep EEGs during the night, this approach was justified. Furthermore, the amount of data was reduced to 20% of the original total.

The above preprocessing was implemented by a computer program, to be described in the next section.

2.4.2 The Main Algorithm

A FORTRAN program called REDUCE was written to perform the following tasks: (1) to reduce the amount of data to be processed by selecting one epoch in each set of five epochs from the original Master Tape (this means that at each 2 1/2 minutes one 30 seconds epoch containing 3840 data points was selected);(2) to check the first epoch of each set for artifacts; (3) to start checking the next three epochs of the set for an artifact free or nearly free epoch, in case an artifact had been already detected in the first epoch; (4) to print a message each time an artifact is detected, stating the cause of rejection and the number of the epoch where it was detected; (5) to print another message if all the epochs in a given set were rejected due to artifacts; (6) to compute statistical parameters such as maximum and minimum amplitude values, mean, variance, kurtosis, skewness, and chi-square coefficient for each chosen epoch; (7) to write on an Auxiliary Tape all epochs that were selected from the Master Tape and their respective statistical parameters.

Figure 2.1 shows a flowchart diagram of REDUCE. Detailed listings of REDUCE and its subroutines are given in Appendix B. We should mention two of the subroutines: RDTAPE and TWRITE, both coded in Assembler Language. They were used to handle input and output operations (I/O) between the computer and the tape units. Due to the length and large number of tape records (7680 bytes), the use of these subroutines reduced the amount of computer time spent in I/O operations by a factor of ten when compared to the same operations executed in FORTRAN.

Four input parameters are required by REDUCE: the starting record (epoch) on the Master Tape for each night; the maximum amplitude level for the amplitude artifact detection test; the upper (UB) and lower (LB) bound for the CSQ test.

The number of the starting epoch on the Master Tape was obtained after the completion of the A/D conversion procedure, when the sampled EEG data was still on the SCRATCH tape. The power spectra for the first ten epochs were computed and then plotted on the IBM 360 computer





printer. By visual inspection, the first plot that showed neither a power spectrum of the calibration signal (sine wave), nor one corresponding to obvious artifacts, was marked, and its number saved as being the number of the starting epoch on the Master Tape. The choice of the remaining input parameters of REDUCE is explained in the next section.

Figure 2.2 shows a typical output printout page generated by REDUCE.

2.4.3 Detection of Artifacts

During the preprocessing phase we used two tests for the detection of artifacts. One was based on amplitude criteria and the other used results of the chi-square test for goodness-of-fit to a Normal Distribution.

To check for very large amplitude artifacts each sampled data point was compared with maximum amplitude levels. In this study the maximum amplitude levels were set at 99 volts ($\pm 300 \mu V$), respectively. Since the Analog Computer allows a maximum swing of ± 100 volts at the output of its amplifiers, any saturation of the amplifiers due to large signals at the input can thus be detected.

If no artifacts were detected by the amplitude test, then the epoch was submitted to a CSQ test. The implementation of this test is explained in detail in Appendix A. The acceptance region for the test was

 $x^2 \le 66.2$

	ARTIFACT	DETECTED	AT	EPOCH	ND.	15	XSQR=1918.97
	ARTIFACT	DETECTED	AT	EPOCH	ND.	20	XSQR=4612.00
	ARTIFACT	DETECTED	AT	EPOCH	ND.	25	XSQR= 301.01
	ARTIFACT	DETECTED	AT	EPOCH	NO.	30	XSQR=2548.85
	ARTIFACT	DETECTED	AT	EPOCH	NO.	31	XSQR=2995.70
	ARTIFACT	DETECTED	AT	EPOCH	ND.	32	XSQR=3855.78
	ARTIFACT	DETECTED	AT	EPOCH	N0.	33	XSQR= 407.23
¢	A SET OF	EPOCHS WAS	SKI	IPPED .	DUE	TO AR	TIFACTS
	ARTIFACT	DETECTED	AT	EPOCH	NO.	45	XSQR=3231.46
	ARTIFACT	DETECTED	AT	EPOCH	ND.	46	XSQR= 844.14
	ARTIFACT	DETECTED	AT	EPOCH	ND.	55	XSQR= 654.64
	ARTIFACT	DETECTED	AT	EPOCH	ND.	65	XSQR=1878.64
	ARTIFACT	DETECTED	A T	EPOCH	ND.	70	XSQR= 486.10
	AMPLITUDE	EXCEEDED	AT	EPOCH	NO.	71	X=-393.27
	ARTIFACT	DETECTED	AT	EPOCH	ND.	85	XSQR= 316.07
	ARTIFACT	DETECTED	AT	EPOCH	NO.	115	XSQR= 517.79
	ARTIFACT	DETECTED	AT	EPOCH	NO.	116	XSQR= 300.20
	ARTIFACT	DETECTED	AT	EPOCH	NO.	117	XSQR= 266.59
	ARTIFACT	DETECTED	AT	EPOCH	ND.	118	XSQR= 350.20
	ARTIFACT	DETECTED	AT	EPOCH	NO.	135	XSQR= 476.75
	ARTIFACT	DETECTED	AT	EPOCH	NO.	136	XSQR=2256.10
	ARTIFACT	DETECTED	AT	EPOCH	NO.	140	XSQR=1803.87
	ARTIFACT	DETECTED	AT	EPOCH	NO.	141	XSQR=1022.24
	ARTIFACT	DETECTED	AT	EPOCH	NO.	150	XSQR= 704.93

FIGURE 2.2 Partial Output Printout from REDUCE
(Subject SD4, 11th night, record #15050)

What this means is that, if the computed x^2 coefficient for a given epoch were within the acceptance region above, the hypothesis that the amplitude distribution function for the epoch follows a Gaussian Distribution could be accepted at the 5% level of significance. The test, by its own formulation, assumes that the set of samples to be tested represents a set of independent random observations. However, in practice, for a bandlimited EEG sequence this assumption does not hold due to the interdependence of successive samples, and the efficacy of the statistical hypothesis is consequently weakened. The problem of interdependence between samples will become even worse when the sampling rate is increased above the Nyquist rate (McEwen, 1975). Due to this fact, and because, furthermore, the EEG is not a purely random process, we would expect that the values of x^2 would not fall necessarily in the acceptance region of the test. For the purpose of detecting artifacts these limitations are not important with the data at hand. After examining ten nights of three subjects in which no artifact detection algorithm was used, we found the following results: artifacts were present in all the epochs in which $x^2 > 280$; artifacts were also present in about 75% of the epochs in which $160 < x^2 < 280$ and that 95% of the epochs with $x^2 < 160$ were artifact free or nearly free. Therefore, there is some uncertainty in the interval 160 < x^2 < 280 as far as the presence of artifacts is concerned. Based on this, we designed the following detection test: (1) if $x^2 < 160$ the epoch is accepted as

artifact free or nearly free; (2) if $x^2 > 280$ the epoch is rejected; (3) if $160 < x^2 < 280$ the epoch is saved by storing its sampled data points in the computer core memory and the algorithm (REDUCE) proceeds to test the next epoch in the set of 5; (4) if none of the remaining epochs in the set has a $x^2 < 160$, then the epoch with the smallest x^2 in the interval 160 < x^2 < 280 will be selected. The above detection algorithm was designed having in mind that no more than four epochs were to be rejected, by any combination of the two tests in any given set of five 30 seconds epochs. The values 160 and 280 correspond to what we called the upper and lower bounds for the CSQ test in Section 2.4.2, and they are input parameters to REDUCE. Later in this study, we found that a lower bound of 180 could be used instead of 160 as was first suggested. The reason we kept using the former value was that several night records had been already processed with the lower bound set at 160. By looking at the printed outputs of REDUCE, we concluded that the change of this parameter to 180 would affect less than 5% of the total number of epochs for each night. Furthermore, an extra amount of computer time would be required to reprocess all the data.

Allowing the selections of epochs with X^2 value in the range 160 < $X^2 \leq 280$ could lead to only two possibilities: (1) the epoch does not have any artifacts; (2) the epoch contains artifacts indeed. In this case, the algorithm will misclassify it as artifact free. In the later case most of the time, we detected the artifact by visual inspection of the Compressed Spectral Array (Section 3.4), and, therefore, we could minimize the changes of misclas-sification.

In order to better understand the operation of the CSQ artifact detection test we will present below six examples derived from the actual data. The epochs in the examples were chosen from the printed results of REDUCE, as being representatives of the test performance under various conditions. The plots were obtained by Digital-to-Analog (D/A) conversion of the sampled EEG, previously stored on the Master Tape, using the same Hybrid System as described in Section 2.3.1 plus a six channel BECKMAN RD DYNOGRAPH.

A FORTRAN program called DTOA was written to perform the D/A conversion and to plot three epochs simultaneously on the polygraph. A listing of this program appears on Appendix B. The polygraph speed was set to 15 mm/sec and calibrated so that 50 μ V would correspond to 5 mm deflection.

Example 1:

Figures 2.4 a, b, c, and d show four successive 30 seconds epochs starting at minute 14 of the 10th night of subject SD1 (record #15340). In the first three epochs, Figures 2.4a, b, and c, there exist high amplitude very low frequency artifacts against a low amplitude and higher frequency background. These three epochs were rejected by the algorithm, because their x^2 values were 507.9, 724.3, and



(b)

FIGURE 2.4 EEGs from Subject SD1 - record #15340

(a) Epoch #28, $X^2 = 507.9$ (b) Epoch #29, $X^2 = 724.3$



(d)

FIGURE 2.4 (continued)

(c) Epoch #30, $X^2 = 377.8$ (d) Epoch #31, $X^2 = 65.7$



FIGURE 2.5 Power Spectrum (Subject SDL, 10th night, record #15340)

(a) Epoch #28 (b) Epoch #29 (c) Epoch #30 (d) Epoch #31

377.8, respectively. The fourth epoch, Figure 2.4d, was chosen to represent the set since its X^2 value was 65.7. In Figures 2.5a, b, c, and d the corresponding power spectrum plots for each of the four epochs are shown. From these it can be seen that the presence of the artifact and of the slow wave activity that followed it resulted in a much larger peak in the Delta Band than the one produced by the entire alpha activity at 9.5 Hz. So by looking at the power spectrum plot the presence of the artifact plus the slow wave activity gave the false impression of a predominant delta activity. In order to see how much of this activity is due to the artifact itself, we recomputed the power spectrum of the epoch shown in Figure 2.4b, with the artifact removed. This operation can be easily performed in the digital computer, by removing only the sampled data points that are comprised of the artifact. Since the time of occurence of the artifact in the epoch, and its duration can be estimated from Figure 2.4b, to locate the sampled data points to be removed becomes an easy task. After the removal of these data points, an equal number of zeros are inserted at the end of the modified epoch in order to keep its duration at 30 seconds. Figure 2.12a, shows the power spectrum corresponding to the epoch in Figure 2.4b with the artifact removed. Comparing the plots in Figures 2.5b and 2.12a, we can see that the artifact alone is responsible for about 70% of the total peak intensity in the Delta Band.

Example 2:

Figures 2.6a and b illustrate two epochs that were correctly selected by the algorithm and their respective X^2 's. Figure 2.6a shows an epoch with slow wave sleep from the 13th night of subject SD2. Spindle activity is clearly seen in the epoch of Figure 2.6b which belongs to the 10th night of subject SD1.

Example 3:

In Figures 2.7a, b, c and d we plotted four successive 30 seconds epochs belonging to the 13th night of subject SD1. The computed X^2 value for the first epoch was 246.3 which fell in the interval $160 < x^2 < 280$. According to the rules of the artifact detection algorithm, this epoch was saved in the computer core memory and the algorithm proceeded to test the next epoch. The X^2 value for the second epoch was 175.7, still inside the above interval. This x^2 value was smaller than the one for the first epoch (175.7 < 246.3)and, therefore, the algorithm substituted the second epoch for the first and saved it in core memory. The third epoch that was tested showed a x^2 value of 207.3 that was in the interval 160 < x^2 < 280. Since the x^2 value for the third epoch was greater than the x^2 value of the second epoch (207.3 > 175.7), the algorithm kept the second epoch previously stored. The fourth and last epoch of the set had a x^2 value of 96.5, and, therefore, this epoch was chosen by the system to represent the set, since its X^2 value was



(b)

FIGURE 2.6 (a) EEG from Subject SD2 - record #15086 Epoch #63, $X^2 = 92.4$

> (b) EEG from Subject SD1 - record #15340 Epoch #69, $X^2 = 106.3$



(b)

FIGURE 2.7 EEGs from Subject SD1 - record #15358

(a) Epoch #50, $x^2 = 246.3$ (b) Epoch #51, $x^2 = 175.7$



FIGURE 2.7 (continued)

(c) Epoch #52, $x_2^2 = 207.3$ (d) Epoch #53, $x^2 = 96.5$





(a) Epoch #50
(b) Epoch #51
(c) Epoch #52
(d) Epoch #53

below the 160 threshold. This example illustrates the case where no visible artifacts can be seen, but three of the x^2 values fall into the region of uncertainty for artifact detection. The decision by the algorithm to reject the first three epochs was correct, according to the rules set before, even though they showed no visible artifacts. The large values of x^2 in the first three epochs are probable due to the high amplitude alpha bursts that appeared against a background of low amplitude activity. Figure 2.8a, b, c and d show the power spectrum plots of the four epochs studied in this example. These plots show that the power spectra for the epochs are essentially the same, except for differences in peak intensities in the alpha band.

Example 4:

Figures 2.9a and b show two consecutive 30 seconds epochs from the 13th night of subject SD1. For the first epoch the computed X^2 value was 204.7 which was in the uncertainty region of the test. Therefore, the algorithm saved this epoch and proceeded to test the second one. Since the X^2 value for this epoch was 96.0, the algorithm selected it as being artifact free, and, therefore, rejected the first epoch. The decision of rejecting the first epoch was a correct one, since a low frequency artifact can be seen in Figure 2.9a. The power spectrum plots corresponding to these two epochs are shown in Figure 2.10a and b. The presence of the low frequency artifacts in epoch #45,



(b)

FIGURE 2.9 EEGs from Subject SD1 - record #15358

(a) Epoch #45, $X^2 = 204.7$ (b) Epoch #46, $X^2 = 96.0$



FIGURE 2.10 Power Spectrum (Subject SD1, 13th night, record #15358)

- (a) Epoch #45
- (b) Epoch #46

Figure 2.9a, resulted in a low frequency spectral peak of intensity comparable to the one corresponding to the alpha activity in the epoch and located at 9.5 Hz. Using the same procedure described in Example 1, we removed the artifact from the epoch and recomputed its power spectrum which is shown in Figure 2.12b. Comparing this plot with the one already shown in Figure 2.10a, we can see the change introduced in the power spectrum as a result of the artifact.

By comparing the results of Examples 3 and 1, we can see the need of having an uncertainty region $(160 < x^2 < 280)$ in the artifact detection algorithm.

Example 5:

In this example we illustrate a case where the CSQ test failed to detect an artifact. Figure 2.11a shows epoch #83 from the 13th night of subject SD2. The computed X^2 value for this epoch was 70.1 and, therefore, it was selected by the algorithm as being artifact free. As we can see from Figure 2.11a an artifact with high frequency activity, probably caused by muscle twitch, is present at the beginning of the epoch. The presence of this artifact resulted in a broadband power spectrum as illustrated in Figure 2.13a.It is this particular characteristic of the resultant power spectrum that enables us to detect this kind of misclassification by the CSQ test, when we examine visually the CSA plots.



(b)

FIGURE 2.11 EEGs from Subject SD2 - record #15086

(a) Epoch #83,
$$x^2 = 70.1$$

(b) Epoch #88, $x^2 = 562.3$



(d)

FIGURE 2.11 (continued)

(c) Epoch #89,
$$x^2 = 171.2$$

(d) Epoch #90, $x^2 = 76.5$





(a) Epoch #29, Figure 2.46, with artifact removed
(b) Epoch #45, Figure 2.9a, with artifact removed



FIGURE 2.13 Power Spectrum (Subject SD2, 13th night, record #15086)

(a) Epoch #83 (b) Epoch #88 (c) Epoch #89 (d) Epoch #90

Example 6:

In Figures 2.11b, c and d three consecutive epochs from the 13th night of subject SD2 are shown. Starting at the first epoch and continuing up to the beginning of the second epoch, there are high amplitude artifacts mixed with high frequency activity, probably due to body movements with muscle twitches. (Figures 2.11b and c). The computed x^2 coefficients for the three epochs in this example were 562.3, 171.2 and 76.5 respectively. According to the algorithm rules the last epoch was chosen to represent the set.

Figures 2.13b, c and d show the power spectrum plots corresponding to the three epochs discussed in this example. The first plot, Figure 2.13b, shows a very high intensity peak at 1 Hz. The second one, Figure 2.13c, shows a peak at 2 Hz of much lower intensity than the one before. In both plots there is a relatively high power content in the upper parts of the spectrum, probably due to muscle activity, as we had already mentioned. The main difference between the plot shown in Figure 2.13d and the other two is the reduction in the power contents in the upper parts of the spectrum.

2.4.4 Performance of the Artifact Detection Algorithm

In order to evaluate the performance of the present artifact detection algorithm, we compared the percentage of epochs rejected due to artifacts, by visual inspection of

the CSA plots, with the same quantity obtained before when no artifact detection test was used. Table 2.5 presents the results of this comparison for nine different nights of three subjects. The record number corresponding to each night appears in the first column; the second and third columns show the percentages of epochs that have to be removed from the CSA plots under the two conditions specified above. The last column shows the percentage of change in the results from the second and third columns. From these results we may conclude that with the use of the present artifact detection algorithm, the number of epochs with artifacts that were present in the CSA plots has been reduced from 58.9% to 94.8% when compared to the results when no detection algorithm was used.
TABLE 2.5

Performance of the Artifact Detection Algorithm

Percentage of Rejected Epochs									
Visual Inspection	n of the CSA's								
With No Artifact Detection	With Artifact Detection	Improvement							
7.3%	3.0%	58.9%							
10.0%	2.6%	74.0%							
14.5%	1.6	90.0%							
6.9%	2.7%	61.0%							
7.2%	2.6%	63.8%							
16.7%	1.5%	91.0%							
20.7%	3.1%	85.0%							
13.3%	2.0%	84.9%							
9.7%	0.5%	94.8%							
	Percentage of Re by <u>Visual Inspection</u> With No Artifact <u>Detection</u> 7.3% 10.0% 14.5% 6.9% 7.2% 16.7% 20.7% 13.3% 9.7%	Percentage of Rejected Epochs by Visual Inspection of the CSA's With No Artifact Detection Detection Detection 7.3% 3.0% 10.0% 2.6% 14.5% 1.6 6.9% 2.7% 7.2% 2.6% 16.7% 1.5% 20.7% 3.1% 13.3% 2.0% 9.7% 0.5%							

CHAPTER III

DATA ANALYSIS

3.1 SPECTRAL ANALYSIS

3.1.1 The Fast Fourier Transform

An infinite-range Fourier transform of a real-valued or a complex-valued record x(t) is defined by the complex-valued quantity

$$X(f) = \int_{-\infty}^{\infty} x(t) \exp[-j2\pi ft] dt \qquad (3.1)$$

For a finite time interval of x(t), say in the range (0,T), the finite-range Fourier transform is defined by

$$X(f,T) = \int_0^T x(t) \exp[-j2\pi ft] dt$$

If x(t) is sampled at N equally spaced points a distance Δt apart, where Δt has been selected such that $\Delta t = 1/(2f_c) = 1/f_s$, where f_c is the highest frequency to be represented and f_s is the sampling frequency of interest (Nyquist rate), then, for an arbitrary frequency f, the discrete version of (3.1) (Discrete Fourier Transform, DFT) is given by: (Bendat and Piersol, 1971)

$$X(f,T) = \Delta t \sum_{n=0}^{N-1} x \exp[-j2\pi f n \Delta t]$$

where $x_n = x(n\Delta t)$ $n = 0, 1, 2, \dots, N-1$

The usual selection of discrete frequency values for the computation of X(f,T) is

$$f_k = k\Delta f = k/T = k/N\Delta t$$

At these discrete frequencies the Fourier coefficients are given by

$$X_{k} = \frac{X(f_{k},T)}{\Delta t} = \sum_{n=0}^{N-1} x_{n} \exp[-j2\pi kn/N] \quad k = 0,1,2,..N-1$$

The above results are unique only out to k = N/2-1 since the Nyquist cutoff frequency occurs at this point. Fast Fourier transform (FFT) methods are utilized to compute these quantities.

The FFT is simply an efficient method for computing the DFT. The FFT can be used in place of the continuous Fourier transform only to the same extent as the DFT, but with a substantial reduction in computer time. The Cooley-Tukey FFT algorithm (Cooley and Tukey, 1965) results in a procedure requiring a number of operations proportional to $Nlog_2N$ rather than N^2 which is required in the DFT algorithms (Brigham and Morrow, 1967; Bergland, 1969; Brigham, 1974).

3.1.2 Leakage Reduction

The problem of leakage is inherent in the Fourier analysis of any finite record of data. The finite record is formed by observing the infinite duration signal for a finite period of time, disregarding what happened to the signal before and after this period of time. This is equivalent to multiplying the actual signal by a rectangular data window. Now, multiplication in the time domain is equivalent to convolution in the frequency domain (Simpson and Houts,1971). The Fourier transform of a rectangular data window is a function with amplitude of the form (sinx)/x. Thus, the Fourier transform of a finite record of data will have additional spectral estimates due to the sidelobes of the sinx/x shape of the Fourier Transformed data window. These sidelobes will "leak" power into adjacent spectral estimates (Bergland, 1969).

The usual approach to reduce leakage is to apply to the actual signal a data window which has lower sidelobes in the frequency domain. A cosine tapered data window was suggested by Tukey (1967) and Bingham et al (1967) to replace the rectangular window in order to reduce the leakage. In this window, a cosine taper is applied to the first and last ten percent of the data, and a unit weight is applied to the data in between. Since only 20 percent of the data points are given weight other than the unit, the computation required to apply this window in the time domain is relatively small. The cosine tapering window W(t) has the form:

$$W(t) = \begin{cases} 0.5(1-\cos\pi t/0.1T), & 0 \le t \le 0.1T \\ 1, & .1T \le t < 0.9T \\ 0.5(1-\cos\pi (T-t)/0.1T) & .9T \le t \le T \end{cases}$$

where T is the data epoch length.

3.1.3 Spectral Resolution and Stability

The need to balance the bandwidth (frequency resolution) of a power spectral estimate against its statistical stability, when dealing with Gaussian data, has been expressed in the form of an uncertainty relation (Bendat and Piersol, 1971).

The frequency resolution of a power spectral estimate is directly related to the time length of the epoch $(B_e = 1/T)$. In other words, the greater the time epoch the better the resolution will be.

The power spectral density function of a stationary random process x(t), when the sample record is of unlimited length T, is given by (Bendat and Piersol, 1971)

$$G_{X}(f) = 2\lim_{T \to \infty} (1/T) E[|X(f,T)|^{2}]$$
(3.2)

where X(f,T) is the finite Fourier transform of x(t), and E denotes expectation. Now an estimate of $G_x(f)$ can be obtained by simply omitting the limiting and expectation operations in (3.2). This will yield the estimate

$$\widetilde{G}_{x}(f) = (2/T) |X(f,T)|^{2}$$
 (3.3)

with the narrowest possible resolution $B_e = 1/T$. Since X(f,T) is a complex number $(X(f,T) = X_R(f,T)+jX_I(f,T))$, it can be shown, assuming Gaussian data, that the real and imaginary parts are uncorrelated random variables with zero mean and equal variances. Furthermore, it can be shown that by squaring them and summing they produce a χ^2 variable

(Bendat and Piersol, 1971; Otnes and Enochson, 1972). The mean and variance of the chi-square variable are n and 2n, respectively, and the normalized standard error, which defines the random portion of the estimator error, is (Bendat and Piersol, 1971)

$$\varepsilon_{r} = \sigma[\tilde{G}_{x}(f)]/G_{x}(f) = (\sqrt{2n})/n = \sqrt{2/n}$$
 (3.4)

where n=2 (number of degrees of freedom). Therefore, $\varepsilon_r = 1$, which means that the standard deviation of the estimate is as great as the quantity being estimated. This random error would be unnacceptable for most applications.

The estimates given by (3.3) are now estimates of the power spectral density function where $B_{\rho} = 1/T$ so that each estimate is governed by a chi-square distribution with two degrees of freedom. In order to get useful estimates we must reduce the standard error ε_r , which is equivalent to increasing the spectral stability. Essentially there are two ways of achieving this. The first way is by averaging over an ensemble of q estimates. The second way is to average together the results for *l* contiguous spectral components of a single sample record. These two methods are also known as smoothing techniques, since, in order to decrease the statistical variability of the spectral estimates, they introduce a smoothing in $G_{v}(f)$, a reduction of spectral resolution. We can also think of these two operations as approximating the expectation operation in Eq. (3.2), and, therefore, making the estimates consistent.

A variation to the first method above was proposed by Welch (1967). This variation involves sectioning a record in segments with variable overlapping intervals, computing the modified periodiograms of these segments and averaging. Principal advantages of this method are:(a) reduction in the required core storage; (b) convenient **a**pplication in test for nonstationarity.

Another way of expressing the normalized standard error ε_r of Eq. (3.4) is in terms of the resolution bandwidth B_{and} the time length of the record T:

$$\varepsilon_r = \sqrt{2/n} = \sqrt{1/B_e T}$$
(3.5)

where $n = 2B_e^T$ is the number of degrees of freedom (Blackman and Tukey, 1958). Of course these results depend on the signal (noise) being exactly Gaussian. Real noises and especially real signals need not be exactly Gaussian. Thus exact results in Gaussian theory would be approximations in practice (Blackman and Tukey, 1958; Richards, 1967).

3.1.4 Spectral Averaging

As we have seen before, one way of gaining statistical stability in G(f) is by averaging together the raw estimates of *l* contiguous spectral components. This operation is achieved by using what is called "spectral windows". Several types of spectral windows have been suggested in the literature.

Two classical spectral windows that have enjoyed a great deal of popularity are the Hann and Hamming Windows,

respectively. (Blackman and Tukey, 1958; Otnes and Enochson, 1972).

The Hann Window is the most widely used mainly due to its easy implementation. It is a triangular window with span of three and has the following weights:

$$\hat{G}_{k} = 0.25\tilde{G}_{k-1} + 0.5\tilde{G}_{k} + 0.25\tilde{G}_{k+1}$$

where \tilde{G}_k is the raw estimate at harmonic k and \hat{G}_k is the corresponding smoothed estimate. An alternative weighting, due to Hamming, achieves a lower amplitude for the immediately adjacent side-lobes, but a slower falling off for subsequent lobes. The weighting is:

$$\hat{G}_{k} = 0.23\tilde{G}_{k-1} + 0.54\tilde{G}_{k} + 0.23\tilde{G}_{k+1}$$

Another convenient method of smoothing the power spectrum is to use symmetric triangular weighting (Singleton and Poulter, 1967) of the form:

$$\hat{G}_{k} = (1/m^{2}) \sum_{j=m+1}^{m-1} (m-|j|) \tilde{G}_{k+j}$$

of span 2m-1, with index of the center value used as the index of the smoothed value. A triangular smoothing of the power spectral density function estimate with span 2m-1 is equivalent to multiplying the autocorrelation function estimate by

$$\left[\frac{\sin(m\pi j/N)}{m\sin(\pi j/N)}\right]^2 \quad \text{for } j = 0, 1, 2, \dots N/2$$

3.1.5 The Present Approach

The 30 seconds epoch selected by REDUCE (Section 2.4.2) were processed as a unit. Each of these epochs contained 3840 sampled data points. Before the data was passed through a cosine tapering window, as described in Section 3.1.2, the estimated mean was subtracted from each data point. Since the FFT algorithm used required a number of data points porportional to a power of two, and the data set had only 3840 points, 296 zeros were added to the original data after the tapering resulting in a new set with 4096 points (2^{12}). Due to the nature of the FFT computational formulas, the effect of adding zeros to the actual data is that of decreasing the space between the spectral estimates, thereby increasing their resolution.

The raw power spectrum for each epoch was calculated with a resolution of 0.03125 Hz ($B_e = 1/T = 1/32$ sec). The raw power spectrum was then smoothed, first by a triangular window with a span of 15, and then by a Hanning window, resulting in a modified power spectrum with a final resolution of $B'_e = 0.5$ Hz. Assuming that the data under consideration is from a Gaussian random process, the standard error ε_r computed from Eq. (3.5), by substituting in $B_e =$ 0.5 Hz and T = 32 sec, will be $\varepsilon_r = 0.25$. The equivalent number of degrees of freedom is from Eq. (3.4),

n =
$$2/(\epsilon_r^2)$$
 = $2/(0.25)^2$ = 32

The $(1-\alpha)$ confidence interval for a spectral density function G(f) based upon an estimate $\hat{G}(f)$ with n degrees of freedom is given by: (Bendat and Piersol, 1971)

$$\hat{nG}(f)/\chi^2_{n;\alpha/2} \leq G(f) < \hat{nG}(f)/\chi^2_{n;1-\alpha/2}$$

In the present work, for a 95% confidence interval for G(f) we have,

$$32\hat{G}(f)/\chi^2_{32;0.025} \leq G(f) < 32\hat{G}(f)/\chi^2_{32;0.975}$$

From a table of percentage of points of a chi-square distribution (Meyer, 1975) it is found, for $\alpha = 0.05$, that $\chi^2_{32;0.025} = 46.98$ and $\chi^2_{32;0.975} = 16.79$, so the interval reduces to

$$0.63\hat{G}(f) < G(f) < 1.79\hat{G}(f)$$

Expressing the above equation in decibels (dB),

$$\hat{H}(f) - 2.0 dB < H(f) < \hat{H}(f) + 2.52 dB$$
 (3.6)

where $H(f) = 10\log G(f)$ and $\hat{H}(f) = 10\log \hat{G}(f)$.

In order to check the performance of the present approach, a bandlimited Gaussian signal with an upper cutoff frequency of B=15 Hz, from a noise generator (Hewlett Packard 3722A) was digitized at 128 samples/sec in epochs of 30 sec duration. The noise generator had an internal digital filter with a roll-off of more than 25 dB/octave above the cutoff frequency B. At the cutoff frequency B the specified attenuation was -3dB. A 30 seconds epoch was frequency analyzed, and the resulting power spectrum is shown in Figure 3.1. The dots are the actual spectral estimates values and they fell within the confidence interval given by Equation (3.6).

3.1.6 Computing the Power Spectral Estimates for the All Night Sleep Data

A FORTRAN program called PSPEC has been written in order to compute the power spectral density function of each epoch for the entire night. A flowchart diagram of PSPEC is shown in Figure 3.2. A detailed list of PSPEC and its subroutines is given in Appendix B.

The input data to PSPEC was the preprocessed data previously selected by REDUCE and stored on the Auxiliary Tapes (Section 2.4.2). Besides the 3840 sampled data points of each epoch, seven additional statistical parameters (maximum and minimum amplitude values, mean, variance, kurtosis, skewness and X^2 coefficient) computed by REDUCE, were stored for each 30 sec record of the tape. After each of these records had been read in, the mean was subtracted from each data point, and the data was tapered by a cosine tapering window (subroutine TAPER1). After zeros had been added to the tapered data, as described in Section 3.1.5, the data was ready to be Fourier transformed.

In the beginning of this work we used a FFT computer subroutined called FOURL to compute the Fourier coef-





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ficients. This subroutine is a FORTRAN version of the Cooley-Tukey FFT algorithm for use upon one-dimensional arrays of complex data whose length is a power of two, and it was written by Brenner (1969). Since our data was real we had to fill with zeros all the imaginary entries of the complex array. The resulting complex Fourier coefficients were symmetrical about N/2. In this way, besides using twice as much core space as we needed, we were also computing twice as many complex Fourier coefficients. The execution time of this subroutine for 4096 real data points was 5.3 seconds. In order to minimize the execution time of PSPEC, we replaced the above FFT algorithm by another one suggested by Singleton (1967) and coded by Aird and Kamer (1975) for the IMSL Library that is available at the Engineering Systems Simulation Laboratory of the Cullen College of Engineering. This New FFT algorithm is called FFTR and it computes the FFT of a real vector of length N, where N is any positive even integer, and the resulting complex Fourier coefficients are computed up to N/2. The execution time of this subroutine for 4096 real data points was 2.74 seconds. As we can see, this algorithm, besides using half of the core space required by the first one, is almost twice as fast.

After the complex Fourier coefficients were obtained, the raw power spectrum was computed. This raw power spectrum was smoothed, as described in Section 3.2.5, using two subroutines: SMT15 and SMOOTH. The maximum and minimum values of the smoothed power spectrum were then computed. Finally, 81 spectral estimates, corresponding to frequencies from 0 to 40 Hz, the seven statistical parameters previously read in, and the maximum and minimum values of the power spectrum were written on a tape (Output Tape). The statistical parameters, plus maximum and minimum values of the power spectrum were also printed in paper as shown in Figure 3.3. The maximum and minimum values of the spectral estimates are used in the scaling of the CSA plots.

3.2 COMPRESSED SPECTRAL ARRAY (CSA)

Once all the power spectral estimates for the entire night were on a digital tape, as described in Section 3.1.6, we then proceeded to obtain the Compressed Spectral Array (CSA). The CSA's were generated at the Image Processing Laboratory of the Electrical Engineering Department, whose facilities include a HP-2100A minicomputer, a HP-7900A Disc Drive, a HP-7970B Digital Tape Unit, a BEEHIVE CRT Display with console, a Teletype Unit and a BENSON-LEHNER incremental plotter.

A FORTRAN program called ARA has been written in order to generate the CSA. A detailed list of ARA and its subroutines is given in Appendix B. A flowchart diagram of ARA is given in Figure 3.4. The teletype or the CRT display with the console were used for input-output communication with the minicomputer.

The CSA is generated by the hidden-line suppression method, and it consists simply of point by point comparison

EPOCH	XMAX	XMIN	VAR I ANC E	SKEWNESS	KURTOSIS	XSQR	MEAN	PSPECMX	PSPECMN
1234567890123456789012345678901234567890123456789012345678901234567890	$\begin{array}{c} 2520888117345384536879991620482680046630244999982166737957\\ 567456486937736455486737921286783142455577495368201794885112522121245557749536820127948851125221212455577495368201279488512957\\ 5674564869377366937736453845720220799916220482680046630244999982166737957\\ 5674564869377366873795766874221212427204344957368851122186737957\\ 5674576687379991622042121242722124245557736874495736875720188512221212455577368757209882512212212212212212212222222222222222$	556991210932524240947748887200496075100122212221222122212221222122212221222	40723123859113314384610676046329900426771167839702709568895415 32483658605678477780984259133896932900426771167831699185162528519674 11032938464958509635344341393233890177513199918516228287022431 1098980123445454565566577778098820980017755131999185162288287022431	$\begin{array}{c} -0.042\\ 20.003382\\ -0.003382\\ -0.003391\\ -0.0003391\\ -0.0003391\\ -0.0003391\\ -0.0003391\\ -0.00003391\\ -0.000000\\ -0.000000\\ -0.0000000\\ -0.0000000\\ -0.0000000\\ -0.0000000\\ -0.0000000\\ -0.0000000\\ -0.0000000\\ -0.00000\\ -0.0000\\ -0.$	464426336610747729756666142107294802497086576534415647694568949 81142096010747729756666142107294802497086576534416210202021014412719 23323332333223333233323332333233333333	557577994937802878583297280003516468215111962735928188296222 8189633639698251482952929011912463907085239988400992265203803 81896336396982514829529290119124639070852399883400992265203803 80038804	7261501544549496969262300712756121788884456379485617148054129 6224645915445498706623300712754674053172987489160874379125502276 6.00000000000000000000000000000000000	40239041691295979471228608041873841863606209063005450919643752888 27796136726117538132336875288863080418738418636062090630054509196638 3222112217553813233647885379022511645551541136788926481628466417754988976498 3222112222222332333333333797777688899880678885785722223312245464121221	0.00036285042576614856288828982898289828982898289828982898289

FIGURE 3.3 Partial Output Printout of PSPEC

(subject SD1, 5th night, record #15322)





FIGURE 3.4 Flowchart Diagram

of ARA

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of spectral magnitudes. Plotting of a spectral point was suppressed if its magnitude plus a vertical scale increment was smaller than the magnitude of the corresponding point in the previously plotted spectrum. This algorithm was executed by subroutine GRAF. The size of the CSA plots is determined by the values assigned to the width in the horizontal direction (X), to the depth (Y), to the height (Z)and the viewing angle, which is the angle that an observer would be looking at the plots in a three dimensional space. In the present study we chose: the viewing angle = 45°, $X = 3^{"}, Z = 2^{"}, and Y = 10^{"}$. These plots were reduced by approximately 35% in order to fit four of them in a 8 1/2 by 11" page. This value of Y is for a CSA plot with 200 epochs, which means that the depth dimension of the plot is also a function of the number of epochs, the net effect of this is to cause an uniform separation between the epochs in the Y direction. The height of the peaks in the Z direction is a function of the largest spectral estimate for a given night, the value of Z and the viewing angle.

Once the CSA was obtained for a given night, it was examined very carefully for epochs with visible artifacts, that have been falsely selected by the artifact detection algorithm. Usually, we can expect their number to range from 1 to 3% of the total number of epochs for the night (see Table 2.5). After the epochs with artifacts were visually detected, their tag numbers were recorded. These numbers were entered into the computer before we proceeded to generate a new CSA with artifact-free epochs.

3.3 FEATURE EXTRACTION

3.3.1 Main Algorithm

In order to get more useful information from the all night power spectra, a computer program called FEATURE has been written. A flowchart diagram of FEATURE is given in Figure 3.5 and a detailed list of it appears in Appendix B. This program was run in the IBM-360 Computer and the input data was fed in by cards and by magnetic tape.

The first thing FEATURE did was to read in the total number of epochs with artifacts and their individual numbers. These numbers were the ones obtained from the CSA as described in Section 3.2. After that a value for the variable NSKIP is read in from a card. NSKIP can take the values of 0, 1, or 2 depending whether we wanted to process all epochs, only the odd or the even numbered ones, respectively

It was found very convenient to process only half of the epochs (either the odd or the even numbered ones), which means that the points in the output plots are spaced at five minute intervals in the horizontal axis. There are some advantages to this procedure. First the plots are not so crowded with points as in the case of considering all the epochs. Nevertheless, the patterns are essentially the same as in the case when all epochs were processed. Second, since some of the epochs are to be skipped due to artifacts,



we can reduce this number by only processing half of the epochs instead of all of them. An additional feature provided by the program allows for the reduction of this number even further. If an epoch is to be skipped, say epoch number 15, the program will then process the next one, epoch number 16, provided that its number is not included in the set of epochs with artifacts. In this way, we can keep track of the timming during the night, which will make it easier to compare our plots with the score sheet results obtained by manual scoring.

The input data to be processed by FEATURE was previously stored **on** a magnetic tape, as described in Section 3.1.6. Each time an epoch was to be processed, 90 data values are read in. The first 81 of these correspond to power spectral estimates from 0 to 40 Hz, and the remaining nine data values contained additional information about the epoch such as: maximum and minimum amplitude values, mean, variance, skewness, kurtosis, and chi-square coefficient from the sampled EEG, plus maximum and minimum values of the magnitude of the power spectrum estimates.

3.3.2 Plot of the Statistical Parameters

Three statistical parameters were processed by FEATURE: chi-square coefficient, kurtosis (plotted as excess of kurtosis), and skewness. The parameters for the entire night are stored in three different arrays for further processing. In order to get rid of the fast frequency variations between the points in each array, a simple moving average lowpass digital filter (Y(k) = 0.25X(k-1)+0.5X(k)+0.25X(k+1), where the X and Y are the input and output to the filter respectively) was used (Steiglitz, 1974). This operation is equivalent to a smoothing of the data. After this was performed, the arrays were put in the appropriate form to be plotted with respect to the time of the night by the subroutine PLOTR2. The subroutine SPARM was then called to compute maximum, minimum, mean and standard deviation values for each array, and these results were printed under each plot.

3.3.3 Separation of the Spectral Estimates in Frequency Bands

The power spectrum of each 30 seconds epoch was divided into seven bands, as follows:

Band	Frequency Range
Delta	0.0 - 3.5 Hz
Theta	4.0 - 7.5 Hz
Alpha	8.0 - 11.5 Hz
Sigma	12.0 - 15.5 Hz
Betal	16.0 - 20.5 Hz
Beta2	21.0 - 29.5 Hz
Fastf	30.0 - 40.0 Hz

After the 81 spectral estimates are read in from the magnetic tape, they are assigned to their respective bands. The power P_j in Band $j(B_j)$ was computed by adding the magnitude of the power spectrum estimates \hat{G}_{i} contained in B_i

$$P_{j} = \sum_{i=k}^{\ell} \hat{G}_{i}$$
 j = 1, 2, ..., 7

where k and l are indices computed from the lower and upper frequencies specified for B_j . The total power for the epoch, J TPOWR is given by:

$$TPOWR = \sum_{i=1}^{81} \hat{G}_i$$

The percentage of power for each B_{i} is then evaluated by:

$$\overline{P}_{j} = ((P_{j})/TPOWR) 100\%$$
 $j = 1, 2, ..., 7$

For each B_i the maximum value of \hat{G}_i was found and its corresponding frequency (F_i) stored. We then tested whether or not this value of \hat{G}_i corresponded to a frequency peak for the band. This test was performed by subroutine PIKCK. For a frequency peak to be classified as such, the intensity in the five adjacent spectral estimates, at least two in each side of the peak, had to be a monotonic decreasing function. In order for \hat{G}_i to be identified as a peak of B_{i} it was required that: $\hat{G}_{i-2} < \hat{G}_{i-1} < \hat{G}_{i} > \hat{G}_{i+1} > \hat{G}_{i+2} > \hat{G}_{i+3}$ or that $\hat{G}_{i-3} < \hat{G}_{i-2} < \hat{G}_i > \hat{G}_{i+1} > \hat{G}_{i+2}$. If \hat{G}_i was accepted as being a frequency peak, then its value and the corresponding frequency were saved into a two-dimensional array for further processing. If \hat{G}_i did not satisfy the above criterion, then PIKCK would search for a value of \hat{G}_i that was a new maximum in B_{i} and the same criterion would be applied again. The

entire procedure would be repeated until all the spectral estimates \hat{G}_i of B_j have been tested. If a frequency peak could not be encountered for B_j then a zero value is assigned to it. A flowchart diagram of PIKCK is given in Figure 3.6 and a detailed list of it is presented in Appendix B.

After all the P_j were computed, they were smoothed, as described in Section 3.3.2, and then plotted by subroutine PLOTR². Subroutine VOST was then called in order to compute maximum, minimum, mean and standard deviation values for the percentage of power in each B_j . These results were printed under each plot.

The frequency peaks and its spectral intensities for each B_j were plotted separately. The spectral intensity plots for all bands B_j were in the same scale. The spectral estimates were scaled by a square root function. The maximum and minimum values for the spectral intensities for all bands and for the entire night were computed by subroutine NERKA. Subroutine PLOTR2 used these values in all the spectral intensity plots for scaling purpose.

3.3.4 Mean Frequency Variation

In Section 3.3.3 it was described how the spectral estimates were separated in frequency bands and how the percentage of power in each of the bands B_j was computed. With this we can picture how the power was shifted among the bands during the night and from night to night. In



order to obtain the same kind of picture, for the mean frequency activity during the night, an algorithm was developed that would compute a mean frequency value for each epoch by plotting these values as a function of the time of the night,we can visualize how the mean frequency varies during a given night and from night to night.

The algorithm simply computes an average of the frequencies of the peak, weighted by their corresponding spectral intensity. Calling this average for each epoch Mean Frequency Coefficient (MFC_n), we have,

MFC_n =
$$(\sum_{j=1}^{7} F_{j}\hat{G}_{j})/(\sum_{j=1}^{7} \hat{G}_{j})$$
 for n = 1,2,...,N

where F_j is the frequency peak at band B_j , \hat{G}_j is its corresponding spectral intensity and N is the total number of epochs being processed. A FORTRAN subroutine called FMED has been written to carry out this algorithm.

CHAPTER IV

RESULTS

4.1 INTRODUCTION

In this chapter we present the results of the application of our data processing techniques to analyze the effect of toxic levels of secobarbital and methaqualone in the sleep EEG of two groups of six subjects each, Group 1 (drug addicts) and Group 2 (nondrug users), as described in Section 2.1. Each of these groups was divided into two subgroups of three subjects each. The subjects in each subgroup were given either secobarbital or methaqualone.

For each subject's night a compressed spectral array (CSA) was plotted (Section 3.2). This kind of plot emphasizes the dynamic shift in power between the frequency bands during a given night and from night to night throughout the withdrawal process. Since the EEG amplitudes for each subject were different, a calibration scale is presented with the CSA plots of each subject.

The algorithm FEATURE (Section 3.3) can generate three plots for each frequency band plus a plot for the MFC coefficients, and three for the statistical parameters (X^2 coefficient, excess of kurtosis and skewness), resulting in a total of 25 plots per night. In order to reduce the amount of output data we chose only four plots (percentage of power in the Delta and Sigma Bands, Mean Frequency Coefficients (MFC) and X^2 coefficient), which, in our opinion, best describe the effects of the drugs on the sleep EEG.

Due to the small size of our samples (three subjects), we could not make any statistical inference about the results. Therefore, we decided to treat each subject as its own control. After we describe the results for all subjects in a given subgroup, we proceed to compare these results among themselves and, if possible, with other subgroups.

4.2 GROUP 1 - SECOBARBITAL

4.2.1 Subject SD1

4.2.1.1 Subject's Protocol

Subject SD1, a 21 year-old male, was in treatment for 18 days. Of these, four nights were selected for automatic analysis. Table 4.1 shows the nights that were processed in a chronological order, the daily amount of drug administered, and the corresponding record number for each night. See Table 2.1 for the subject's complete withdrawal schedule.

TABLE 4.1

Analyzed Data for Subject SD1

Night	Record Number	Daily Dosage (mg)
5th	15322	1000
lOth	15340	200
13th	15358	0
16th*	15375	0
*Subjec	t complain	ed of intense
pain iņ	i nis arm c	luring the high

4.2.1.2 Compressed Spectral Array (CSA) Results

Figures 4.1a, b, c and d show the CSA plots for the nights listed in Table 4.1. The plot for the 5th night, Figure 4.1a, shows that the activity in the Delta Band was present most of the time as low intensity spectral peaks except at the beginning of the night, when there is a train of high intensity peaks. Appearing simultaneously with this train, there is a series of spectral peaks centered around 10-11 Hz, which probably correspond to slow spindle activity "riding" on the low frequency waves. At the very beginning of the night there are some peaks located at 7-7.5 Hz, which correspond to an alpha-like activity according to the raw EEG recorded on paper and to the subject's sleep stage score sheet for this night. The peaks in the Sigma Band are not well defined and their central frequencies are shifting between 12 and 14 Hz. Some low intensity activity in the Betal and Beta2 bands can also be seen in the plot.

In the 10th night, Figure 4.1b, when the subject was still taking the drug (200 mg daily), several changes have taken place mainly in the Alpha, Sigma, Betal and Beta2 Bands. At the onset of the night there are well-defined peaks in the Alpha Band. The Sigma Band shows peaks with lower intensity, centered around 13-13.5 Hz, which are now more well defined. No visible activity can be seen in the Betal and Beta2 Bands.



FIGURE 4.1 Subject SD1 - Compressed Spectral Array (CSA)
(a) 5th night, record #15322 (b) 10th night, record #15240
(c) 13th night, record #15358 (d) 16th night, record #15375

During the 13th and 16th nights, Figures 4.1c and 4.1d, when the subject was not taking the drug anymore, the main noticeable change is the appearance of a new train of high intensity peaks in the Delta Band before the first half of the night.

4.2.1.3 <u>Summary of the Output Results from the FEATURE</u> Algorithm

In Table 4.2, the results of the data reduction analysis performed by FEATURE are given. From this table it can be seen that there is a tendency for the mean percentage of power in the Delta and Theta Bands, the mean value and the percentage of rejections of the X^2 coefficient to increase as the drug was being withdrawn. On the other hand, the mean percentage of power in the remaining frequency bands and the mean value of the Mean Frequency Coefficients (MFC) have a tendency to decrease. Table 4.3 shows the percentage of change in the results of Table 4.2, between the last and the first nights.

It should be noticed that in Table 4.2, the values for the mean percentage of power and for the mean frequency peaks in the Theta and Alpha Bands are higher than what it would be expected. This was caused by the alpha-like activity (7.0-7.5 Hz) that was detected by the algorithm as being in the Theta Band and by the slow spindle activity (10.0-11.0 Hz), as described in Section 4.2.1.2, that was detected as being in the Alpha Band.

TABLE 4.2

Summary of the Output Results from FEATURE for Subject SD1

NIGHT			5th	lOth	13th	16th	
RECORD NUMBER			15322	15340	15358	15375	
DOSAGE (mg)			1000	200	0	0	
DELTA	PERC. OF	MEAN	27.3	38.0	44.3	46.3	
BAND	POWER	STD	8.2	9.9	12.9	12.4	
(0 0-3 547)	FREQ. PEAK	MEAN	1.5	2.0	1.7	1.8	
(0.0-3.3n2)	(Hz)	STD	0.5	0.8 0.5		0.5	
	PERC.	MEAN	16.2	24.9	25.1	22.5	
BAND	POWER	STD	4.2	4.2 6.9 5.6		4.9	
	FREQ. PEAK	MEAN	6.9	5.3	5.5	5.5	
(4.0-7.5Hz)	(Hz)	STD	0.8	1.4	1.1	1.3	
	PERC. OF	MEAN	23.3	15.2	16.7	14.5	
ALPHA	POWER	STD	5.0	5.2	11.1	10.6	
DAND	FREQ. PEAK	MEAN	10.3	9.3	9.3	9.4	
(8.0-11.5Hz)	(Hz)	STD	1.0	1.3	0.8	0.9	
C T CMA	PERC.	MEAN	17.4	10.3	6.4	5.6	
BAND	POWER	STD	5.0	6.9	2.9	2.5	
	FREQ. PEAK	MEAN	13.1	13.4	13.7	13.6	
(12.0-15.5Hz)	(Hz)	STD	1.0	1.0	0.6	0.8	

(continued)

NIGHT			5ht	10th	13th	16th
RECORD	NUMBER		15322	15340	15358	15375
DOSAGE (mg)			1000	200	0	0
D.5.00 3 1	PERC.	MEAN	8.7	3.5	1.9	2.5
BETAL	POWER	STD	3.1	1.7	1.1	1.2
(16.0-20.5Hz)	FREQ. PEAK	MEAN	18.0	18.5	18.5	18.4
(1010 2015h2)	(Hz)	STD	1.2	1.2	1.2	1.3
ጋር ምህን ጋ	PERC.	MEAN	3.9	3.8	1.8	2.3
BAND	POWER	STD	1.7	2.1	1.0	1.1
(21.0-29.5Hz)	FREQ. PEAK	MEAN	24.5	24.4	24.6	25.1
	(Hz)	STD	2.5	1.9	2.4	2.5
FASTF BAND	PERC. OF	MEAN	1.5	1.2	1.1	0.45
	POWER	STD	0.4	0.5	0.6	0.3
	FREQ. PEAK	MEAN	33.6	33.4	33.0	33.1
(30.0-40.0HZ)	(Hz)	STD	2.7	2.5	2.2	2.3
	MEAN		59.1	81.1	93.9	96.6
x ²	STD		8.0	13.2	17.2	15.6
	PERCENTAGE OF REJECTIONS		27.0	73.0	86.0	86.8
MFC	MEAN STD		7.9 1.7	6.2 1.6	5.0 1.6	5.0 1.4

	DELTA	THETA	ALPHA	SIGMA	BETAL	BETA2	FASTF	x ²	MFC
	BAND	BAND	BAND	BAND	BAND	BAND	BAND	COEF.	
MEAN PERC. OF POWER	+ 70%	+ 40% ¹	- 38% ¹	- 68%	- 71%	- 418	- 82%	-	-
MEAN FREQ. PEAK	+ 20%	- 20%l	- 9% ¹	+ 5%	+ 2%	+ 38	- 18	-	-
MEAN	-	-	-	-	-	_	-	+ 63%	- 37%
PERC. OF REJECTIONS	-	-	-	-	-	-	-	+210%	-

¹See text for comments

TABLE 4.3 Subject SD1 - Percentage Change in the Results of Table 4.2

Between the Last and the First Nights

In Figures 4.2a, b, c and d the plots of the percentage of power in the Delta Band vs time are shown. For the 5th night, Figure 4.2a shows a well-defined peak in the beginning of the night and the mean percentage of power in the band is 27.3%. In the 10th night, Figure 4.2b, the mean percentage of power has increased to 38% and the welldefined peak is still present at the beginning of the plot. Figure 4.2c, for the 13th night, illustrates the appearance of a second well-defined peak 110 minutes after the first The same pattern is repeated in the 16th night, Figure one. 4.2d, except that the two peaks are now separated by a 90 minutes interval. The mean percentage of power in the band during these two nights has increased to 44.3 and 46.3% respectively. From Table 4.3, we can see that the mean percentage of power in the Delta Band has increased 70% between the 5th and the 16th nights. In the same period the mean frequency in this band has increased 20%.

The plots for the percentage of power in the Sigma Band are shown in Figure 4.3a, b, c and d. In the 5th night, Figure 4.3a, the plot stays almost flat for the entire night, which reflects the "constancy" of the activity in the Sigma Band. From the 10th night on, well-defined peaks can be seen with a decreasing amplitude. The mean percentage of power has decreased 68% from the 5th to the 16th night, and the mean frequency peak has changed from 13.1 to 13.6 Hz in the same period.

The Mean Frequency Coefficients (MFC) for four nights are plotted in Figures 4.4a, b, c and d. In the 5th night, Figure 4.4a, most of the MFC points are above 7 Hz, except for a low frequency dip which is close to the 4 Hz frequency line, when the EEG was dominated by slow wave activity. The mean frequency for this night is 7.9 Hz. The plot for the 10th night, Figure 4.4b, shows a similar pattern to the one in the 5th night, except that the frequency dip in the beginning of the night is under the 3 Hz frequency line, which shows a slight increase in the slow wave activity. For the rest of the night the MFC are of lower frequency than in the preceeding nights. The mean frequency for the entire night decreased to 6.2 Hz. By the 13th night, Figure 4.4c, when the subject was out of the drug, the mean frequency for the night continued its downward trend reaching a value of 5 Hz. Now there are two low frequency dips that are under the 3Hz frequency line, and the rest of the MFC points are below the 6 Hz frequency line, except at the very beginning of the night when the activity in the Alpha Band was dominant as shown in Figure 4.1c. The plot for the 16th night, Figure 4.4d, shows a pattern similar to the one in the 13th night, except that the low frequency dips are of shorter duration and there is a high frequency peak around minute 400. According to the CSA plot, Figure 4.1a, this peak corresponds to activity in the Alpha Band (the subject was probably awaken). The mean frequency during the night has decreased 37% between the 5th and the 16th nights.


FIGURE 4.2 Subject SDl - Percentage of Power in the Delta Band (a) 5th night, record #15322 (b) 10th night, record #15240 (c) 13th night, record #15358 (d) 16th night, record #15375

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5th night, record #15322 (b) 10th night, record #15340 (a) 13th night, record #15358 (c) (d) 16th night, record #15375



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13th night, record #15358 (c) (d) 16th night, record #15375



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Figures 4.5a, b, c and d show the plots for the x^2 coefficients. In the 5th night, Figure 4.5a, the great majority of the points are below 72 and only 27% of the epochs were rejected, by the chi-square test of goodness-of-fit to a Gaussian distribution (CSQ). This means that the EEG amplitude in 73% of the epochs follows a Gaussian probability density function. As the drug is being withdrawn, the x^2 values start increasing and consequently more and more epochs are being rejected by the CSQ test as we can see in the remaining plots (Figures 4.5b, c and d). There is an increase of 210% in the percentage of rejections between the 5th and the 15th nights.

4.2.2 Subject SD2

4.2.2.1 Subject's Protocol

Subject SD2 is a 16 year-old male, who was in treatment for a period of 14 days. Four nights of sleep EEG, as shown in Table 4.4 were automatically processed. This table was compiled from Table 2.1. The titration night (daily dosage of 500 mg) was not processed because its record was not available on magnetic tape.

TABLE 4.4

Night	Record Number	Daily Dosage (mg)
2nd	15017	300
5th	15029	150
7th	15043	0
13th	15086	0

Analyzed Data for Subject SD2

4.2.2.2 Compressed Spectral Array

The CSA plots for the nights listed in Table 4.4 are shown in Figures 4.6a, b, c and d. In the 2nd night, Figure 4.6a, the activity in the Delta Band is dominated by low intensity peaks throughout the night, with the exception of two clusters of high intensity peaks that appear in the first third of the night. The Sigma Band shows well-defined peaks around 12-12.5 Hz and in some parts of the night it is the dominant frequency band. No peaks can be seen in the Beta frequency bands and very few peaks are present in the Theta Band. Several changes can be seen in the CSA plot for the 5th night, Figure 4.6b. The peaks in the Delta Band are of much higher intensity than before, and two new clusters of high peaks have appeared. The first one around the middle of the night and the other in the last third of The peaks in the Sigma Band are also of higher intensity. it. Several peaks can now be seen in the Theta Band. In the 7th night, Figure 4.6c, when the subject was not taking the



FIGURE 4.6 Subject SD2 - Compressed Spectral Array (CSA) (a) 2nd night, record #15017 (b) 5th night, record #15029 (c) 7th night, record #15043 (d) 13th night, record #15086

drug, the Delta Band continued to show large intensity peaks and the peaks in the Sigma Band are of smaller amplitude than before. No significant changes can be seen in the patterns of the 13th night as shown in Figure 4.6d.

4.2.2.3 Summary of the Output Results from the FEATURE Algorithm

Table 4.5 shows the output results from the data reduction analysis. From these results we can see that the drug affected the amount of SWS, which resulted in a reduction in the mean percentage of power in the Delta and Theta Bands. On the other hand, the drug also affected the high frequency activity, which resulted in an increase in the amount of power in the Alpha, Sigma, Betal, and Beta2 frequency bands. The above changes can be further summarized by the results from the mean value of the Mean Frequency Coefficient for each night.

Figures 4.7a, b, c and d show the plots vs time of the percentage of power in the Delta Band for the 2nd, 5th, 7th and 13th nights respectively. The 2nd and 5th nights, Figures 4.7a and b, show a similar pattern, with two large peaks in the first half of the night and another one at the last third of it. This last peak is of much higher amplitude in the 5th night. The mean percentage of power in the two nights was 40.4 and 45.3% respectively. In the 7th night, Figure 4.7c, there are three well-defined large peaks: two in the first third of the night and the other in the beginning of

TABLE 4.5

Summary of the Output Results from FEATURE for Subject SD2

NIGHT			2nd	5th	7th	13th
RECOPD	NUMBER		15017	15029	15043	15086
DOSAG	E (mg)		300	150	0	0
DELTA	PERC.	MEAN	40.0	45.3	51.8	57.7
BAND	POWER	STD	13.5	13.8	12.2	10.3
(0 0-3 5Hz)	FREQ. PEAK	MEAN	1.5	1.7	1.6	1.6
(0.0*3.5112)	(Hz)	STD	0.5	0.5	0.4	0.5
	PERC.	MEAN	18.3	19.9	23.2	21.7
THETA	POWER	STD	7.8	6.0	5.4	6.7
	FREQ.	MEAN	5.9	6.2	6.3	6.5
(4.0-7.5Hz)	PEAK (Hz)	STD	0.9	1.0	1.0	0.9
	PERC. OF	MEAN	14.2	13.1	11.9	9.0
ALPHA	POWER	STD	4.9	5.1	6.5	4.3
BAND	FREQ. PEAK	MEAN	9.7	9.1	9.2	9.4
(8.0-11.5Hz)	(Hz)	STD	1.2	1.1	1.0	1.0
GTCMA	PERC.	MEAN	14.8	11.8	6.6	4.9
BAND	POWER	STD	7.4	6.8	3.5	2.7
	FREQ. PEAK	MEAN	12.6	13.1	13.4	13.2
(12.U-15.5HZ)	(Hz)	STD	0.9	0.9	0.8	0.7

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NIGHT			2nd	5th	7th	13th
RECORD	NUMBER		15017	15029	15043	15086
DOSAG	E (mg)		300	150	0	0
BETAI	PERC. OF	MEAN	4.7	3.2	1.7	1.2
BAND	POWER	STD	2.1	1.8	1.0	0.6
(16 0-20 587)	FREQ. PEAK	MEAN	18.2	18.3	18.3	18.3
(10.0 20.512)	(Hz)	STD	1.3	1.4	1.4	1.3
ጋር ጥ እ ጋ	PERC.	MEAN	3.0	2.5	1.0	0.8
BAND	POWER	STD	1.3	1.4	0.7	0.4
	FREQ. PEAK	MEAN	24.0	24.2	24.2	24.4
(21.0-29.5Hz)	(Hz)	STD	2.3	2.0	2.4	2.3
	PERC.	MEAN	1.1	1.1	0.3	0.3
BAND	POWER	STD	0.5	0.6	0.2	0.2
	FREQ.	MEAN	33.4	32.8	33.2	32.5
(30.0-40.0Hz)	(Hz)	STD	2.4	2.0	2.5	2.3
	MEAN		75.0	76.3	85.6	86.6
v ²	STD		8.9	12.5	17.5	19.8
Δ	PERCENTAGE OF REJECTIONS		63.2	62.4	68.4	71.3
MFC	MEA STD	N	5.8 1.8	5.4 1.8	4.2 1.6	3.4 1.0

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETA1 BAND	BETA2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+ 43%	+ 19%	- 37%	- 67%	- 75%	- 73%	- 73%	-	-
MEAN FREQ. PEAK	+ 7%	+ 10%	- 3%	+ 5%	+0.5%	+ 2%	- 3%	-	-
MEAN	-	-	-	-	-	-	-	+ 16%	- 41%
PERC. OF REJECTIONS	_		-	-	-	-	-	+ 13%	-

TABLE 4.6 Subject SD2 - Percentage Change in the Results of Table 4.5

Between the Last and the First Nights

the second half of the night. It seems that a fourth one was appearing at the very end of the night. The mean percentage of power for the night has increased to 51.8%. In the 13th night there are four large peaks, two in the first third and two in the second third of the night. From these plots it seems that the drug has suppressed one of the peaks and somehow altered the timming of their occurance. Table 4.6 shows that the mean percentage of power in the Delta Band has increased 43% between the 2nd and the 13th nights. The mean frequency in the Delta Band has also increased 7% in the same period.

Figures 4.8a, b, c and d show the plots of the percentage of power in the Sigma Band. In the 2nd night, Figure 4.8a, there are peaks with large amplitude and long duration. From the 5th night on, Figures 4.8b, c and d, the amplitude and the latency of the peaks start decreasing. From Table 4.6 it can be seen that the mean percentage of power in the Sigma Band has decreased 67% between the 2nd and the 13th nights. In the same period, the mean frequency in the band increased from 12.6 to 13.2 Hz, (as shown in Table 4.5).

Figures 4.9a, b, c and d show the plots of the MFC's for the four nights of subject SD2. In the 2nd night, Figure 4.9a, the mean frequency was 5.8 Hz and for most of the night the MFC points were above 5 Hz, except for three low frequency dips that went below 3 Hz. Almost the same picture is still valid for the 5th night, Figure 4.9b, with



FIGURE 4.7 Subject SD2 - Percentage of Power in the Delta Band (a) 2nd night, record #15017 (b) 5th night, record #15029 (c) 7th night, record #15043 (d) 13th night, record #15086

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(c) 7th night, record #15043

(d) 13th night, record #15086



the exception of the latency of the low frequency dips, which is now longer. The mean frequency for this night was 5.4 Hz. Most of the MFC points in the 10th night, Figure 4.9c, were below 5 Hz and the mean frequency went further down to 3.4 Hz, with most of the MFC points staying below 4.5 Hz. For this subject the mean frequency in the MFC plots, has decreased 41% between the 2nd and the 13th nights.

The plots for the x^2 coefficient are shown in Figures 4.10a, b, c and d. As we can see from these plots, the mean and the percentage of rejections of the x^2 coefficients had their lowest values close to the subject's titration night. As the drug was being withdrawn, these values started increasing. From Table 4.6 it can be seen that the mean and the percentage of rejections increased 16 and 13%, respectively, between the 2nd and the 13th nights.

4.2.3 Subject SD3

4.2.3.1 Subject's Protocol

Subject SD3, a male 16 years of age, was a patient at the hospital for 13 days. A total of four nights were automatically processed. Table 4.7 shows these nights, their corresponding record number and the subject's daily intake of the drug. The data in Table 4.7 was derived from Table 2.1, where the complete subject's withdrawal schedule is shown.

TABLE 4.7

Daily Record Night Number Dosage (mg) lst 15030 350 3rd 15044 200 8th 15079 0 13th 15105 0

Analyzed Data for Subject SD3

4.2.3.2 Compressed Spectral Array Results

Figures 4.11a, b, c and d show the CSA plots corresponding to the four nights in Table 4.7. For the 1st and 3rd nights, Figures 4.11a and b, the activity in the Delta Band is characterized by low intensity peaks, except for two separate occurrence of sharp high intensity spectral peaks in the first half of the night. The delta peaks for the 3rd night have a higher intensity than the ones for the 1st night in the beginning of the night. The activity in the Sigma Band is about the same magnitude in both nights, however, it is better defined in the 3rd night. Furthermore, its peak frequency has shifted from 12.5 Hz in the beginning of the 1st night to 13.0 Hz in the beginning of the 3rd Little activity is also seen in the Betal and Beta2 night. frequency bands during the 1st night, but it is not visible anymore in the 3rd one.

In the 8th and 10th nights, Figures 4.11c and ϕ , when the subject was out of the drug, the large peaks in the



FIGURE 4.11 Subject SD3 - Compressed Spectral Array (CSA) (a) 1st night, record #15030 (b) 3rd night, record #15044 (c) 8th night, record #15079 (d) 13th night, record #15105

Delta Band at the beginning of each night are of higher intensity than the corresponding ones in the 1st and 3rd nights. In the 8th night, Figure 4.11c, there are also some large peaks after the first half of the night. Activity in the Alpha Band is clearly seen during the first 50 minutes of the 8th night. The frequency peaks in this band are first at 9 Hz and then shifted to 11.5 Hz during this period.

4.2.3.3 <u>Summary of the Output Results from the FEATURE</u> Algorithm

Table 4.8 shows the output results of the data reduction algorithm. As we can see from this table, there is an upward trend in the mean percentage of power in the Delta and Theta Bands, as well as in the mean value and in the percentage of rejection of the X^2 coefficient as the treatment progresses. However, a downward trend is shown in the mean percentage of power in the remaining frequency bands and by the MFC. Table 4.9 shows, in terms of percentages, the variations in the results of Table 4.8 between the last and the first nights of treatment.

Figures 4.12a, b, c and d show the plots vs time of the percentage of power in the Delta Band for the nights listed in Table 4.7. For the lst night, Figure 4.12a, there are two large peaks 100 minutes apart at the beginning of the night. The mean percentage of power in the band is 37%. These two peaks are still present in the 3rd night, except that now the first peak is of much higher amplitude, its

TABLE 4.8

Summary of the Output Results from FEATURE for Subject SD3

NIGHT			lst	3rd	8th	13th
RECOPD	NUMBER		15030	15044	15079	15105
DOSAG	E (mg)		300	200	0	0
DELTA	PERC. OF	MEAN	37.0	44.1	54.8	54.2
BAND	POWER	STD	9.2	12.8	13.5	13.5
	FREQ. PEAK	MEAN	1.5	1.6	1.6	1.7
(0.0-3.5HZ)	(Hz)	STD	0.3	0.5	0.4	0.3
mr rim a	PERC. OF	MEAN	13.4	18.7	18.1	19.9
BAND	POWER	STD	2.7	6.0	4.3	5.6
(4.0-7.5Hz)	FREQ. PEAK	MEAN	6.1	5.1	5.9	5.7
	(Hz)	STD	0.8	0.9	1.0	1.30
	PERC. OF	MEAN	10.4	10.2	9.8	9.6
ALPHA	POWER	STD	2.2	3.3	7.5	7.4
BAND	FREQ. PEAK	MEAN	9.6	9.8	10.2	9.7
(8.0-11.5Hz)	(Hz)	STD	1.1	1.1	1.0	1.0
GTCMA	PERC.	MEAN	13.5	13.4	7.4	8.3
BAND	POWER	STD	4.7	7.7	4.0	4.4
	FREQ. PEAK	MEAN	13.2	13.3	13.7	13.7
(12.0-15.5Hz)	PEAK (Hz)	STD	0.8	0.9	0.7	0.7

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NI	NIGHT			3rd	8th	13th
RECORD	NUMBER		15030	15044	15079	15105
DOSAG	E (mg)		300	200	0	0
BETAL	PERC. OF	MEAN	8.2	5.2	2.5	2.3
BAND	POWER	STD	2.6	2.8	1.8	1.3
(16.0-20.5Hz)	FREQ. PEAK	MEAN	18.0	18.5	18.6	18.5
(2000 2000)	(Hz)	STD	1.3	1.3	1.4	1.2
<u> ዝ</u> ድመል ጋ	PERC.	MEAN	9.6	4.5	2.4	1.8
BAND	POWER	STD	2.7	2.5	1.9	1.2
	FREQ. PEAK	MEAN	23.7	23.9	24.4	24.6
(21.0-29.5Hz)	(Hz)	STD	2.7	2.3	2.4	2.3
FACMF	PERC. OF	MEAN	4.5	1.0	1.3	0.7
BAND	POWER	STD	1.7	0.5	0.8	0.4
(20.0-40.00-	FREQ. PEAK	MEAN	33.1	33.4	33.0	33.2
(30.0-40.0H2)	(Hz)	STD	2.3	2.3	2.2	2.5
	MEA	N	75.7	76.2	83.0	84.7
x ²	STD		10.2	14.8	11.7	16.5
	PERCENTAGE OF REJECTIONS		61.1	60.2	78.4	70.7
MFC	MEA STD	N	7.3 1.9	5.9 1.8	4.4 2.0	4.4 1.7

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETAl BAND	BETA 2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+ 47%	+50%	- 48	- 39%	- 72%	- 81%	- 87%	-	-
MEAN FREQ. PEAK	+ 13%	- 78	+ 1%	+ 48	+ 3%	+ 48	0.3%	-	-
MEAN	-	-	-	-	-	-		+ 12%	- 40%
PERC. OF REJECTIONS	-	-	-	_	-	_	-	+ 16%	-

TABLE 4.9 Subject SD3 - Percentage Change in the Results of Table 4.8

Between the Last and the First Nights

duration is twice as longer and they are separated by 110 minutes interval (Figure 4.12b). The mean percentage of power in the 3rd night has increased to 44%. In the 8th night, Figure 4.12c, when the subject was not taking the drug anymore, there are three large peaks present in the plot. Two of these peaks are in the first half of the night and the other at the beginning of the second half of The first two peaks are separated by about 100 minutes it. and the third peak is 120 minutes apart from the second one. The mean percentage of power has now increased to 54.8%. In the last night the mean percentage of power in the Delta Band stayed at 54.2% and only two large peaks are shown at the beginning of the night (Figure 4.12d). These peaks are 50 minutes apart. The mean percentage of power in the Delta Band has increased 47% according to Table4.9. The mean frequency peak in this band has also increased from 1.4 to 1.7 Hz, between the 1st and the 13th nights.

Figures 4.13a, b, c and d show the plots of the percentage of power in the Sigma Band for the four nights. In the 1st night, Figure 4.13a, the percentage of power stays relatively high for long periods of time with a mean of 13.5%, and no well-defined peaks can be seen. The plot for the 3rd night, Figure 4.13b, shows some high amplitude peaks. The mean percentage of power for the night remained at 13.4%. In the 8th and 13th nights the mean percentage of power has decreased to 7.4 and 8.3% respectively. Their plots, Figure 4.13c and d, show that the amplitude of the peaks has also decreased. From Table 4.9 we can see that the mean percentage of power in the Sigma Band has decreased 39% between the 1st and the 13th nights. In the same period the mean frequency peak in the band has increased from 13.2 to 13.7 Hz.

In Figures 4.14a, b, c and d we can see the plots for the MFCs. In the 1st night, Figure 4.14a, the mean frequency is high for most of the night, except for two well-defined, short duration low frequency dips in the first third of the night. The mean frequency for the entire night is 7.3 Hz. The plot for the 3rd night, Figure 4.14b, shows a large low frequency dip of long duration in the beginning of the night. A second low frequency dip is present at the beginning of the second third of the night, however, its duration is much shorter than that of the first one. The mean frequency for the night was 5.9 Hz, which is lower than the previous value for the 1st night. By the 8th night, Figure 4.14c, when the subject's daily drug intake was reduced to zero, the two low frequency dips are still present and a third one appeared towards the end of the second third of the night. The mean frequency during the night went down even further to reach a value of 4.4 Hz. In the last night, Figure 4.14d, the low frequency activity dominates most of the first third of the night, which results in two low frequency dips in this interval. The mean frequency for the night stayed at 4.4 Hz. There is a 40% reduction in the mean frequency from the 1st to the 13th night as shown by Table 4.9.







FIGURE 4.12 Subject SD3 - Percentage of Power in the Delta Band (a) 1st night, record #15030 (b) 3rd night, record #15044 (c) 8th night, record #15079 (d) 13th night, record #15105



(c) 8th night, record #15079 (d) 13th night, record #15105



(a) 1st night, record #15030 (c) 8th night, record #15079

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(d) 13th night, record #15105



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The plots for the X^2 coefficients are shown in Figures 4.15a, b, c and d. The mean and the percentage of rejection for the X^2 coefficient followed the same upward trend observed in the previous subjects. The mean and the percentage of rejections increased 12 and 16%, respectively, between the 1st and the 13th nights.

4.2.4 Discussion of Results of the Subjects in Group 1 Withdrawn from Secobarbital

Table 4.10 contains comparisons of the changes in the output results of FEATURE between the last and the first nights that were processed for each of the subjects in the Group 1, that were withdrawn from secobarbital. This table summarizes the results already presented in Tables 4.3, 4.6, and 4.9.

From Table 4.10 we can see that the mean percentage of power in the Delta Band increased with the withdrawal of the drug in all subjects. Subject SDl, who started with the highest titration dosage (1000mg of secobarbital) in the Group, had the largest increase (70%) and the other two subjects showed increases above 40%. In addition, two of the subjects, SDl and SD3, showed a marked increase in the mean frequency peak in this band as the drug was withdrawn. This is equivalent to saying that the drug caused not only a reduction in the percentage of power in the Delta Band, but that it also slowed down the "frequency" of the "delta waves."

TABLE 4.10

SUBJ	ECTS	SD1	SD2	SD3	
TITRATION	DOSAGE (mg)	1000	500	350
DELTA BAND	PERC. OF POWER	MEAN	+70%	+43%	+47%
(0.0-3.5Hz)	FREQ. PEAK (Hz)	MEAN	+20%	+ 78	+13%
THETA	PERC. OF POWER	MEAN	+40%	+19%	+50%
BAND (4.0-7.5Hz)	FREQ. PEAK (Hz)	MEAN	-20%	+10%	- 7%
ALPHA	PERC. OF POWER	MEAN	-38%	-37%	- 48
BAND (8.0-11.5Hz)	FREQ. PEAK (Hz)	MEAN	- 98	- 38	+ 1%
_ SIGMA	PERC. OF POWER	MEAN	-68%	-67%	-39%
BAND (12.0-15.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 5%	+ 5%	+ 4%

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Summary of the Results of Tables 4.3, 4.6 and 4.9 for in Subjects in Group 1 - Secobarbital

TABLE 4.10

(continued)

SUBJ	ECTS		SD1	SD2	SD3
TITRATION	DOSAGE (mg)	1000	500	350
BETAL	PERC. OF POWER	MEAN	_71%	-75%	-72%
BAND (16.0-20.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 2%	+0.5%	+ 38
BETA2	PERC. OF POWER	MEAN	-41%	-73%	-81%
BAND (21.0-29.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 38	+ 2%	+ 4%
FASTF	PERC. OF POWER	MEAN	-70%	-73%	-87%
BAND (30.0-40.0Hz)	FREQ. PEAK (Hz)	MEAN	- 1%	- 38	0.3%
x ²	MEAN	1	+63%	+16%	+12%
	PERCENTAGE OF REJECTIONS		+210%	+13%	+16%
MFC	MEAN		-378	-418	-40%

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an increase in the power in the higher frequency bands, as we can clearly see from the CSA plots. The amount of power in a frequency band is a function of the amplitude as well as the amount of activity present in that particular frequency. In this way, using only spectral analysis results, we cannot say for sure that the reduction of power in the Delta Band is caused only by a reduction in the amplitude of the delta waves, because it can also be caused by a decrease in the number of these waves, or by both. Spectral analysis is, by its own nature, a good descriptor of the "background" EEG activity in the different sleep stages, especially regarding the "frequency content" in the different bands. Α good example of this occurs in the 5th night of subject SD1, in which the results of the manual scoring show no sleep stages 3 or 4 present during the night. However, the CSA plot for this night, Figure 4.1a, shows the presence of relatively high intensity peaks in the Delta Band at the beginning of the night. This could suggest that the patient even under the effects of a large dosage of the drug, may generate sufficient activity in the delta frequency band to qualify as stage 3, or, perhaps, stage 4 sleep, although the EEG voltage did not reach the 75 μ V threshold, required by the visual scorer, for acceptance as delta activity. (Williams et al, 1964).

There is disagreement among investigators on to the effects of barbiturates on sleep. Most studies report no significant change in delta sleep after barbiturates (Kay, 1973). Lester et al (1968) reported changes in the distribution of the delta sleep throughout the night, after the administration of 200 mg of secobarbital to a group of 14 college students. However, the drug did not cause a significant change in the total amount of delta sleep, computed by using period analysis techniques. In the present study we do observe changes in the 90-100 minutes delta rhythm that is present in normal subjects (Lubin et al, 1973) as shown in the plots vs time of the percentage of power in the delta band (Figures 4.2, 4.7, and 4.12). The drug also caused the suppression of some of the delta peaks in these plots. This can be further illustrated by the plots for subject SDL., Figures 4.2a, b, c and d, in which the second delta peak was completely suppressed in the first two nights

The average percentage of power in the Sigma Band decreased with the withdrawal of the drug in all three subjects. Two of them showed changes above 60%. The mean frequency peak in this band increased 4-5% for all the subjects, which is equivalent to a slowdown of 0.5-0.6 Hz in the frequency of the spindle activity due to the effects of the drug. These results agree with several studies in the literature (Kay et al, 1972 and Kay, 1973).

From Table 4.10 we can also see that there is a reduction in the average percentage of power in the Alpha, Betal and Beta2, and FASTF frequency bands with the withdrawal of the drug. An opposite trend is observed in the mean percentage of power in the Theta Band. The average frequencies computed for each night showed a downward trend with the withdrawal of the drug. Table 4.10 shows that the subjects in this group had a reduction in the mean frequency between 37 and 41% during the treatment. For two of the subjects, SD1 and SD2, we can see from Tables 4.2 and 4.8 that although the mean frequency for each night was decreasing as the drug was being withdrawn, the standard deviation for each subject stayed practically constant from night to night. This could suggest that the basic mechanism responsible for the EEG's frequency shifts during the night was not seriously affected by the drug and that the changes in the mean frequency from night to night were probably due to the changes in the amount of slow and fast wave activities.

The mean value and the percentage of rejections (percentage of epochs in which the computed X^2 fell outside the region of acceptance for the chi-square test for goodness-of-fit to a Gaussian distribution) of the X^2 coefficients increased in all subjects with the withdrawal of the drug. The highest changes were shown by subject SD1. From Table 4.2 we can see that during the night following the subject's titration procedure (5th night, record #15322) the EEG amplitude followed a Gaussian probability density function in 73% of the epochs during the night. By the l6th night this number was reduced to 13.2%. The other two subjects followed the same trend, but with much less variability. This suggest that the EEG's amplitude distribution had become more Gaussian due to the effects of the drug. Furthermore, it seems that the degree of Gaussianity is related to the initial amount of drug given to the subjects. It has been suggested that, on the basis of the Central Limit theorem, increased Gaussianity in observed EEG activity may reflect an increased degree of independence among individual cortical neural generators (Elul, 1969). A similar increase was also observed by McEwen et al (1975) in subjects under halothane anesthesia.

4.3 GROUP 1 - METHAQUALONE

4.3.1 Subject SD4

4.3.1.1 Subject's Protocol

Subject SD4, a 25 year-old female, was intreatment at the Poly Drug Treatment Unit for 14 days. After the discharge the subject reported to the hospital for two additional nights. The subject's complete withdrawal schedule is shown in Table 2.2. Table 4.11 shows the four nights that were selected for analysis, the daily amount of drug administered and the corresponding record number for each night. It was not possible to process any of the nights close to the subject's discharge from the hospital, due to the lack of good analog tape recordings for these nights.
TABLE 4.11

Daily Record Night Number Dosage (mg) lst 14497 2100 3rd 15006 1500 7th 15024 600 llth^l 15050 0 ¹Night following the first day of zero drug intake.

Analyzed Data for Subject SD4

4.3.1.2 Compressed Spectral Array Results

Figures 4.16a, b, c and d show the CSA plots for the nights listed in Table 4.11. For the 1st and 3rd nights. Figures 4.16a and b, the activity in the Delta Band was present most of the time as low intensity peaks, except at the beginning of the nights when there are large peaks. These delta peaks are, however, of less intensity in the 3rd night. For both nights the subject's alpha activity produced spectral peaks centered around 7.0-7.5 Hz. For the 3rd night this is only true at the beginning of the night, since some spectral peaks are seen in the Alpha Band at 10.0-10.5 Hz in the second half of the night. The Sigma Band shows welldefined high intensity peaks around 13.0-13.5 Hz. These peaks, however, are of lower intensity in the 3rd night. Some low intensity activity can also be seen in the Betal and Beta2 Bands during the 1st night.

In the 7th night, Figure 4.16c, when the subject's daily intake of drug has been reduced to 600 mg, several



FIGURE 4.16 Subject SD4 - Compressed Spectral Array (CSA) (a) lst night, record #14497 (b) 3rd night, record #15006 (c) 7th night, record #15024 (d) llth night, record #15050

changes have taken place mainly in the Delta and Alpha Bands. The frequency peaks corresponding to the subject's alpha activity has moved from 7.0-7.5 Hz to 10.0-11.0 Hz and its intensity is much lower. In the Delta Band a new cluster of high intensity peaks are present in the beginning of the second half of the night. The high intensity peaks in the Sigma Band are still present and its center frequency is now shifted between 13.5 and 14 Hz.

The first 90 minutes of the 11th night, Figure 4.16d, are dominated by alpha activity which resulted in spectral peaks centered around 10.5-11.5 Hz. In the Delta Band there are two clusters of high intensity spectral peaks The central frequency of the spectral peaks in the Sigma Band is now between 14 and 14.5 Hz.

4.3.1.3 Summary of the Output Results from the FEATURE Algorithm

In Table 4.12 the results of the data reduction analysis performed by FEATURE are given. From these results we can see that due to the effects of the drug there is a tendency for the mean percentage of power in the low frebands (Delta and Theta) to decrease in the first night when compared to the results in the llth night. The mean value and the percentage of rejection of the X^2 coefficient also follow the same trend. An upward trend is, however, shown by the mean percentage of power in the Sigma, Betal, Beta2 and FASTF frequency bands and by the mean of the MFCs.

TABLE 4.12

Summary of the Output Results from FEATURE for Subject SD4

NI	GHT		lst	3rd	7th	llth
RECORD	NUMBER		14497	15006	15024	15050
DOSAG	E (mg)		2100	1500	600	0
DELTA	PERC.	MEAN	31.8	28.7	36.3	38.8
BAND	POWER	STD	11.8	10.4	8.6	10.5
(0, 0, 2, ETT-)	FREQ. PEAK	MEAN	1.7	1.8	1.9	2.0
(0.0-3.5HZ)	(Hz)	STD	0.5	0.6	0.7	0.6
	PERC.	MEAN	15.5	16.9	23.2	24.4
THETA	POWER	STD	3.9	3.0	5.0	3.5
DARD	FREQ.	MEAN	6.4	5.9	6.0	6.0
(4.0-7.5Hz)	(Hz)	STD	1.1	1.2	1.1	0.8
	PERC. OF	MEAN	12.1	15.5	13.5	15.9
ALPHA	POWER	STD	3.4	4.4	3.3	3.7
BAND	FREQ.	MEAN	9.6	9.3	9.5	10.2
(8.0-11.5Hz)	(Hz)	STD	1.1	1.1	1.2	1.3
CTCMD	PERC.	MEAN	18.5	15.7	14.2	10.6
BAND	POWER	STD	9.4	6.7	6.6	2.4
	FREQ. PEAK	MEAN	13.2	13.4	13.6	14.1
(12.0-15.5Hz)	(Hz)	STD	0.7	0.7	0.5	0.6

TABLE 4.12

(continued)

NI	GHT		lst	3rd	7th	llth
RECORD	NUMBER		14497	15006	15024	15050
DOSAG	E (mg)		2100	1500	600	0
BETAI	PERC. OF	MEAN	8.1	7.2	4.0	3.8
BAND	POWER	STD	4.0	2.8	1.9	1.9
(16.0-20.587)	FREQ. PEAK	MEAN	18.4	18.4	18.7	18.4
(10.0 20.512)	(Hz)	STD	1.2	1.3	1.3	1.3
<u> ጋር መን</u> ሳ	PERC.	MEAN	8.6	8.9	4.5	4.3
BAND	POWER	STD	4.6	3.9	3.4	3.5
	FREQ. PEAK	MEAN	24.7	24.5	24.9	24.5
(21.0-29.5Hz)	(Hz)	STD	2.2	2.4	2.6	2.5
EACHE	PERC. OF	MEAN	2.8	4.8	1.7	1.9
BAND	POWER	STD	1.5	2.0	1.0	2.0
	FREQ.	MEAN	33.6	33.4	33.3	33.6
(30.0-40.0Hz)	(Hz)	STD	2.3	2.3	2.4	2.3
	MEA	N	72.9	76.2	83.6	87.6
x ²	STD		13.4	14.9	12.4	19.2
25	PERCEN OF REJECT	TAGE IONS	57.0	59.6	77.8	75.8
MFC	MEA STD	N	8.2 2.1	8.9 2.1	6.9 1.7	6.5 1.7

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	DELTA	THETA	ALPHA	SIGMA	BETA1	BETA2	FASTF	x ²	MFC
	BAND	COEF.							
MEAN PERC. OF POWER	+ 22%	+ 56%	+ 31%	- 43%	- 53%	- 49%	- 32%	-	
MEAN FREQ. PEAK	+ 18%	- 6%	+ 7%	+ 78	08	- 1%	08	-	-
MEAN	_	-	-	_	-	-	-	+ 20%	- 21%
PERC. OF REJECTIONS	_		_	_	_	-		+ 33%	-

TABLE 4.13 Subject SD4 - Percentage Change in the Results of Table 4.12

Between the Last and the First Nights

Table 4.13 shows, in terms of percentage, the changes in the results of Table 4.12 between the last and the first nights in consideration. However, these results should be compared with caution with the previous ones, mainly due to the shorter interval of time between the llth night and the last day the subject took the drug.

Figures 4.17a, b, c and d show the plots of the percentage of power in the Delta Band for the 1st, 3rd, 7th and llth nights respectively. The 1st and 3rd nights, Figures 4.17a and b, show a similar kind of pattern, with two large peaks in the first third of the night. The mean percentage of power for the two nights was 31.8 and 28.7%, respectively. In the 7th night, Figure 4.17c, two large peaks can be seen, located around minutes 115 and 280 respectively. In between them there are two other peaks with smaller amplitudes. The mean percentage of power for the night has increased to 36.3%. The plot for the 11th night, Figure 4.17d, shows two high amplitude peaks in the first half of the night, and the mean percentage of power for the night was 38.4%. Table 4.13 shows that the mean percentage of power in the Delta Band has increased 21% between the 1st and the llth nights. The mean frequency in this band has also increased 18% in the same period.

In Figures 4.18a, b, c and d the plots of the percentage of power in the Sigma Band for the four nights are presented. The first three nights show well-defined high amplitude peaks. The means of the percentage of power for

these three nights were 18.5, 15.7 and 14.2% respectively. By the llth night, Figure 4.18d, the high amplitude peaks have disappeared and the mean percentage of power is reduced to 10.6%. From Table 4.13 we can see that the average percentage of power in the Sigma Band has decreased 43% between the 1st and the llth nights. In the same period the mean frequency peak in the band has changed from 13.2 to 14.1 Hz (Table 4.12).

The Mean Frequency Coefficients (MFC) for the nights listed in Table 4.12, are plotted in Figures 4.19a, b, c and d. For the 1st night the mean frequency in most of the epochs was above 7.5 Hz, except for a low frequency dip in the beginning of the night which reached the frequency of The mean frequency for the entire night was 8.2 Hz. 3Hz. In the 3rd night the low frequency dip was still present at the beginning of the night as shown in Figure 4.19b, and the activity for the rest of the night was dominated by fast frequency waves which resulted in the MFC points being above 8 Hz. The mean frequency during this night was 8.9 Hz. By the 7th night, Figure 4.19c, the mean frequency has decreased to 6.9 Hz and there is only one frequency dip of short duration in the first third of the night. In the first 90 minutes of the 11th night, Figure 4.19d, the MFC's in the epochs stayed above 7.5 Hz, reflecting the activity in the Alpha Band, as shown by the CSA plot in Figure 4.16d. There are two low frequency dips before the first half of the night. In the last 60 minutes the alpha activity was again dominant.





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(d) 11th night, record #15050







(c) 7th night, record #15024

(d) 11th night, record #15050

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The mean frequency for this night was 6.5 Hz. According to the results of Table 4.13, the mean frequency decreased 21% between the 11th and the 1st nights.

In Figures 4.20a, b, c and d the plots of the X^2 coefficient for the four nights are shown. In the first two nights, Figures 4.20a and b, more than 40% of the epoch had an EEG with a Gaussian amplitude distribution and the mean value of the X^2 coefficients were 72.9 and 74.4 respectively. These values were up to 83.6 and 83.5 in the last two nights (Figures 4.20c and d), and the percentage of epochs with a Gaussian distribution has decreased to 22.2 and 34.7 respectively. The average value and the percentage of rejections for the X^2 coefficient have increased 20 and 33% between the lst and the llth nights respectively (Table 4.13).

4.3.2 Subject SD5

4.3.2.1 Subject's Protocol

Subject SD5, a 35 year-old male, was patient at the Poly Drug Unit for 15 days. After the discharge, the subject slept at the hospital on three consecutive nights (16th, 17th and 18th). It was reported that he had been drinking heavily since after leaving the hospital. Table 4.14 shows the nights that were automatically processed, the daily amount of drug given to the subject and the corresponding record number for each night. This table was derived from Table 2.2, in which the subject's withdrawal schedule is shown. Due to technical problem with the tape recorder, the first 40 minutes of the 13th night (record #15063) were not recorded on magnetic tape. According to the score sheet for this night, the subject spent the first 61 minutes of the night in stage 0 sleep. The only reason to process this record is that no other record was available before the subject's discharge from the hospital, and due to his drinking problem the data collected after this time would be questionable.

TABLE 4.14

Analyzed Data for Subject SD5

	Record	Daily
Night	Number	Dosage (mg)
<u></u>		
lst	14498	3000
3rd	15007	2100
JIU	20007	2200
5+b	15016	1975
Jui	1010	10/2
12421	15062	0
<u>13tn-</u>	12002	<u> </u>
¹ This 1	night corre	spond to the
first d	lav the sub	piect was out
or the	arua.	

4.3.2.2 Compressed Spectral Array

The CSA plots for the nights listed in Table 4.14 are shown in Figures 4.21a, b, c and d. In the last night, Figure 4.21a, the activity in the Delta Band is characterized by low intensity spectral peaks throughout the night, except for some high intensity peaks towards the end of the night. Most of the energy, however, is concentrated in the



FIGURE 4.21 Subject SD5 - Compressed Spectral Array (CSA) (a) 1st night, record #14498 (b) 3rd night, record #15007 (c) 5th night, record #15016 (d) 13th night, record #15063

Sigma Band, where high intensity peaks are present around 12.0-12.5 Hz. The peaks in the Alpha Band are centered at 8.5 Hz. Activity in the Betal and Beta2 frequency bands can also be seen. In the 3rd night, Figure 4.21b, the CSA plot shows that there is a little increase in the intensity of the peaks in the Delta Band at the beginning of the night. However, the main observable change in this plot is the decrease in the intensity of the peaks in the Sigma Band. Activity in the Beta Bands can still be seen. The decrease in activity in the Sigma Band continued in the 5th night, Figure 4.21c, and a cluster of high intensity peaks in the Delta Band started forming at the beginning of the night. As we had already explained in Section 4.3.2.1, 40 minutes of EEG are missing in the beginning of the 13th night. This was indicated in the CSA plot, Figure 4.21d, by starting the time axis at minute 40. As we can see from this plot, after the first hour of sleep some high intensity peaks are present in the Delta Band. The activity in the Sigma Band is much lower than in the previous nights. The peaks in the Alpha Band are now centered around 10 Hz.

4.3.2.3 Summary of the Output Results from the FEATURE

Algorithm

The results of the data reduction analysis, performed by FEATURE are shown in Table 4.15. These results show the same kind of trends already described for subject SD4. In Table 4.16, the changes in the results of Table 4.15, between the first and last nights that were processed are

TABLE 4.15

Summary of the Output Results from FEATURE for Subject SD5

· NIC	GHT		lst	3rd	5th	13th
RECOPD	NUMBER		14498	15007	15016	15063
DOSAG	E (mg)		3000	1875	1650	0
DELTA	PERC. OF	MEAN	34.0	34.0	40.1	45.1
BAND	POWER	STD	8.5	10.0	10.0	9.8
	FREQ. PEAK	MEAN	1.6	1.7	1.6	1.8
(0.0-3.5HZ)	(Hz)	STD	0.4	0.7	0.6	0.5
	PERC.	MEAN	17.6	22.4	22.0	25.1
THETA	POWER	STD	2.5	4.6	2.4	3.2
BAND	FREQ.	MEAN	6.2	6.5	6.2	6.2
(4.0-7.5Hz)	(Hz)	STD	0.9	0.8	1.0	1.1
	PERC. OF	MEAN	15.5	18.1	14.9	13.4
ALPHA	POWER	STD	3.4	4.3	4.3	5.6
BAND	FREQ.	MEAN	9.0	9.6	9.5	9.3
(8.0-11.5Hz)	(Hz)	STD	1.2	1.3	1.3	1.1
C T C M A	PERC.	MEAN	15.8	11.6	10.5	6.6
BAND	POWER	STD	4.8	3.6	4.2	2.6
	FREQ.	MEAN	12.5	12.7	12.7	13.0
(12.0-15.5Hz)	(Hz)	STD	0.7	1.1	1.0	0.9

TABLE 4.15

(continued)

NI	GHT		lst	3rd	5th	13th
RECORD	NUMBER		14498	15007	15016	15063
DOSAG	E (mg)		3000	1875	1650	0
ይፑጥኔነ	PERC.	MEAN	7.5	5.9	4.9	3.4
BAND	POWER	STD	2.9	3.0	3.1	2.3
(16.0-20.5Hz)	FREQ. PEAK	MEAN	18.3	18.2	18.0	18.3
(10.0 20.512)	(Hz)	STD	1.21	1.2	1.4	1.2
BFTA 2	PERC.	MEAN	5.7	4.4	3.5	2.6
BAND	POWER	STD	1.7	2.4	2.3	1.8
	FREQ. PEAK	MEAN	23.7	23.8	23.9	23.8
(21.0-29.5Hz)	(Hz)	STD	2.0	2.4	1.9	2.1
EXCHE	PERC. OF	MEAN	0.9	0.9	0.7	0.6
BAND	POWER	STD	0.4	0.8	0.4	0.4
	FREQ.	MEAN	33.3	33.5	33.1	32.6
(30.0-40.0Hz)	(Hz)	STD	2.3	2.4	2.3	2.2
	MEA	N	68.5	66.0	85.6	88.6
x ²	STD		12.1	7.7	16.4	17.2
	PERCEN OF REJECT	TAGE IONS	46.6	47.5	80.4	81.5
MFC	MEA STD	N	7.3 1.2	6.5 1.7	6.0 1.9	4.9 1.7

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETA1 BAND	BETA2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+ 33%	+ 43%	- 14%	- 58%	- 55%	- 54%	- 30%	-	-
MEAN FREQ. PEAK	+ 13%	08	+ 3%	+ 48	08	+0.4%	- 28	-	•
MEAN	-	_	_	-	-	_		+ 29%	-33%
PERC. OF REJECTIONS	_	_	_	-	-	-	-	+ 75%	-

TABLE 4.16 Subject SD5 - Percentage Change in the Results of Table 4.15

Between the Last and First Nights

shown. The results in this table should be looked with caution due to the following: (1) technical problems with the analog magnetic tape in the 13th night (record #15063), as we have already mentioned in Section 4.3.2.1 and (2) the date of this night is not far enough from the date of the last ingestion of the drug, as it was in the other patients (with the exception of SD4). In fact, as we can see from Table 2.3, this was the first night the subject was not taking the drug.

The plots for the percentage of power in the Delta Band are shown in Figures 4.22a, b, c and d. In the 1st night, Figure 4.22a, only two peaks can be seen, one at the beginning of the night and the other around minute 380. The plot for the 3rd night, Figure 4.22b, shows three welldefined peaks. Of those the first one is of higher amplitude than the peaks in the 1st night. The average percentage of power stayed at 34% for both nights. By the 5th night, the mean percentage of power in the band was up to 40%, four peaks are present in the plot (Figure 4.22c) and the amplitude of the first peak has risen even further. In the 13th night, Figure 4.22d, three peaks are present and of those the second one has the largest amplitude. The average percentage of power for the night has increased to 45%. Table 4.16 shows that the average percentage of power and the mean frequency in the Delta Band have increased 33 and 13% respectively between the first and the last nights.

Figures 4.23a, b, c and d show the plots for the percentage of power in the Sigma Band. In the first three nights the amplitude of the peaks stayed relatively high, declining sharply in the last night. From Table 4.16 we can see that the average percentage of power in the band has decreased 58% between the lst and l3th nights. In the same period the mean frequency in the band has increased 4%.

The plots for the Mean Frequency Coefficients (MFC) are shown in Figures 4.24a, b, c and d. In the 1st night, Figure 4.24a, most of the MFC points are above the 6.5 Hz frequency line, with a mean frequency for the night of In the 2nd night, Figure 4.24b, a low frequency dip 7.3 Hz. can be clearly seen in the beginning of the plot. This dip went below the 2.5 Hz line. The mean frequency for the night is down to 6.5 Hz, reflecting mainly a decrease in the activity in the Sigma Band as shown by the CSA plot in Figure 4.21b. Figure 4.24c shows the plot for the 5th night, in which the low frequency dip is still present in the beginning of the night, except that now it went close to 2 Hz. The mean frequency for the night continued its downward trend reaching now 6 Hz. In the 13th night, Figure 4.24d, there are two low frequency dips that went below 3 Hz, and the mean frequency for the night is 4.9 Hz. This last result would probably be higher if the first 40 minutes of the night were not deleted, since, according to the manual scoring, the subject was in stage 0 sleep during that time. From Table 4.16, we can see that the mean frequency during the night



(a) lst night, record #14498 (b) 3rd night, record #15007 (c) 5th night, record #15016 (d) 13th night, record #15063



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FIGURE 4.23 Subject SD5 - Percentage of Power in the Sigma Band (a) 1st night, record #14498 (b) 3rd night, record #15007 (c) 5th night, record #15016 (d) 13th night, record #15063





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has decreased 33% between the 1st and the 13th nights.

In Figures 4.25a, b, c and d the plots for the X^2 coefficients are shown. In the first two nights, Figures 4.25a and b, the EEG's amplitude follows a Gaussian probability function in more than 50% of the epochs. In the last two nights this number has decreased to less than 20%. Table 4.16 shows that the mean and the percentage of rejections for the X^2 coefficient has increased 29 and 75% respectively between the lst and the l3th nights.

4.3.3 Subject SD6

4.3.3.1 Subject's Protocol

Subject SD6, a 16 year-old male, was a patient at the hospital during a period of 10 days. A total of four nights were automatically perocessed. Table 4.17 shows these nights, their corresponding record number and the subject's daily intake of the drug. The data in Table 4.17 was derived from Table 2.2, in which the subject's complete withdrawal schedule is shown.

TABLE 4.17

Analyzed Data for Subject SD6

	Record	Daily
Night	Number	Dosage (mg)
2nd	15341	600
3rd	15351	300
4th	15359	0
7th	15376	0

4.3.3.2 Compressed Spectral Array

Figures 4.26a, b, c and d show the CSA plots corresponding to the four nights listed in Table 4.17. For the 2nd and the 3rd nights, Figures 4.26a and b, the activity in the Delta Band is characterized by low intensity spectral peaks for most of the night, except for some peaks with high intensities at the beginning of each night. The peak activity in the Sigma Band is about the same magnitude in both nights and its center frequency is around 12.0-12.5 Hz. In the 4th and 7th nights, Figures 4.26 c and d, two clusters of high intensity peaks can be clearly seen in the Delta Band. The intensity of the spectral peaks in the Sigma Band is now lower than in the previous two nights.

4.3.3.3 <u>Summary of the Output Results from the FEATURE</u> Algorithm

Table 4.18 shows the output results from the data reduction algorithm. From these results we can see that the drug affected the amount of SWS, which resulted in a reduction in the mean percentage of power in the Delta Frequency Band. On the other hand the drug also affected the high frequency activity which resulted in an increase in the percentage of power in the Sigma, Betal and Beta2 frequency bands. The above changes can be further summarized by the results from the mean value of the MFC's for each of the nights.

Figures 4.27a, b, c and d show the plots of the percentage of power in the Delta Band, for the 2nd, 3rd, 4th



FIGURE 4.26 Subject SD6 - Compressed Spectral Array (CSA) (a) 2nd night, record #15341 (b) 3rd night, record #15351 (b) 4th night, record #15359 (d) 7th night, record #15376

TABLE 4.18

Summary of the Output Results from FEATURE for Subject SD6

NI	GHT		2nd	3rd	4th	7th
RECOPD	NUMBER		15341	15351	15359	15376
DOSAG	E (mg)		600	300	0	0
DELTA	PERC.	MEAN	42.7	39.3	43.8	52.4
BAND	POWER	STD	14.4	14.4	16.5	15.4
(0 0-2 5Hz)	FREQ. PEAK	MEAN	1.5	1.6	1.6	1.6
(0.0-5.5h2)	(Hz)	STD	0.4	0.5	0.4	0.3
ጠጠን	PERC. OF	MEAN	18.5	19.9	21.8	21.7
BAND	POWER	STD	6.5	5.6	7.2	6.5
	FREQ.	MEAN	6.7	6.3	6.3	6.8
(4.0-7.5Hz)	PEAK (Hz)	STD	0.8	1.1	1.2	0.8
	PERC. OF	MEAN	12.0	15.9	17.1	12.6
ALPHA	POWER	STD	4.9	9.0	13.5	13.7
BAND	FREQ. PEAK	MEAN	8.9	8.7	9.0	8.9
(8.0-11.5Hz)	(Hz)	STD	1.0	0.9	1.2	0.9
STGMA	PERC.	MEAN	12.9	15.4	9.0	7.5
BAND	POWER	STD	8.3	9.3	5.9	5.6
(12 0.15 57-)	FREQ. PEAK	MEAN	12.6	12.6	13.0	13.0
(12.0-13.3HZ)	(Hz)	STD	0.6	0.8	1.1	0.8

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(continued)

NI	GHT		2nd	3rd	4th	7th
RECORD	NUMBER		15341	15351	15359	15376
DOSAG	E (mg)		600	300	0	0
BETAI	PERC. OF	MEAN	5.0	3.5	2.8	1.4
BAND	POWER	STD	2.8	2.5	2.0	1.0
(16 0-20 5Hg)	FREQ. PEAK	MEAN	18.2	18.6	18.8	18.5
(10.0-20.5h2)	(Hz)	STD	1.3	1.4	1.4	1.2
	PERC.	MEAN	4.0	2.5	2.3	0.8
BAND	POWER	STD	2.3	1.7	1.9	0.5
	FREQ. PEAK	MEAN	24.0	24.2	24.4	24.4
(21.0-29.5Hz)	(Hz)	STD	2.1	1.9	2.5	2.0
EI A CIMEI	PERC. OF	MEAN	1.2	0.8	0.6	0.2
BAND	POWER	STD	0.7	0.5	0.6	0.1
	FREQ.	MEAN	32.7	32.9	32.9	33.0
(30.0-40.0Hz)	(Hz)	STD	2.1	2.1	2.0	2.3
	MEA	N	74.4	73.9	78.7	85.8
x ²	STD		10.8	10.4	16.8	16.3
4	PERCEN OF REJECT	PERCENTAGE OF REJECTIONS		61.2	62.8	78.4
MFC	MEA STD	N	5.9 2.0	6.4 2.0	5.4 2.1	4.3 1.8

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETAl BAND	BETA2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+ 23%	+ 17%	+ 5%	- 42%	- 72%	- 80%	- 83%	-	-
MEAN FREQ. PEAK	+ 7%	+1.4%	0%	+ 3%	+ 2%	+ 2%	+ 1%	-	-
MEAN	-	-	_	-	-		-	+ 15%	- 27%
PERC. OF REJECTIONS	_	-	_	-	-	-	-	+ 30%	-

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TABLE 4.19 Subject SD6 - Percentage Change in the Results of Table 4.18

Between the Last and the First Nights

and 7th nights respectively. The 2nd and 3rd nights, Figures 4.27a and b, show a similar kind of pattern, with a high amplitude and long duration peak in the beginning of both nights. The mean values of the percentage of power in the two nights are 42.7 and 39.3% respectively. In the 4th and 7th nights, Figures 4.27c and d, there are two high amplitude peaks in the first half of the night. However, the peaks in the 7th night are of higher amplitude and the time interval between them is shorter when compared to the peaks in the 3rd night. The mean percentage of power in the 7th night is 52.4% for the 4th night. Table 4.19 shows that the mean percentage of power in the Delta Band has increased 23% from the 1st to the 7th night.

Figures 4.28a, b, c and d show the plots of the percentage of power in the Sigma Band, for the four nights. In the 1st and 2nd nights, Figures 4.28a and b, there are peaks with large amplitude and long duration. The values of the mean percentage of power for these two nights are 12.9 and 15.4% respectively. The peaks in the 4th and 7th nights, Figures 4.28c and d, are of lower amplitude and short latency than the ones in the previous nights. The values of the mean percentage of power in these two nights have decreased to 9.0 and 7.5% respectively. Table 4.19 shows that the mean percentage of power in the Sigma Band has decreased 42% between the 2nd and the 7th nights.

The plots of the Mean Frequency Coefficients for the four nights of subject SD6 are shown in Figures 4.29a, b, c

and d. In the 2nd night, Figure 4.29a, the mean frequency is 5.9 Hz and there is a low frequency dip in the first third of the night, which reached the frequency of 1.5 Hz. The MFC points stayed above 5 Hz for the remaining of the In the 3rd night, Figure 4.29b, the pattern is alnight. most the same, except that the MFC points were above the 6.5 Hz line for the rest of the night. The mean frequency for the entire night has increased to 6.4 Hz. In the 4th and 7th nights, Figures 4.29c and d, there is the appearance of a new low frequency dip in the first third of the night, in addition to the one already present in the previous plots. The mean frequency for the nights has now decreased to 5.4 Hz and 4.3 Hz respectively. From Table 4.18, we can see that the mean frequency has declined from 5.8 Hz in the 2nd night to 4.3 Hz in the 7th night which represents a 27% change (Table 4.19).

Figures 4.30a, b, c and d show the plots of the X^2 coefficients for the four nights listed on Table 4.17. In the first three nights the EEG's amplitude followed a normal distribution in almost 40% of the epochs. In the last night, however, this value has decreased to 21.6%. From Table 4.19, we can see that the mean and the percentage of rejections of the X^2 coefficient had increased 15 and 30% respectively from the 2nd to the 7th night.



FIGURE 4.27 Subject SD6 - Percentage of Power in the Delta Band (a) 2nd night, record #15341 (b) 3rd night, record #15351 (c) 4th night, record #15359 (d) 7th night, record #15376



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FIGURE 4.28 Subject SD6 - Percentage of Power in the Sigma Band (a) 2nd night, record #15341 (b) 3rd night, record #15351 (c) 4th night, record #15359 (d) 7th night, record #15376






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4.3.4 Discussion of the Results for the Subjects in Group 1 Withdrawn from Methaqualone

In order to compare the results between the subjects in Group 1 that were titrated and withdrawn from methaqualone, we grouped the results presented in Tables 4.13, 4.16 and 4.19 in Table 4.20. As we pointed out before, the results in this table for subjects SD4 and SD5 should be looked with caution due to the fact that the data used as baseline was obtained from nights in which the subject were still under the effects of the drug.

From Table 4.20 we can see that the mean percentage of power in the Delta Band increased in all subjects with the withdrawal of the drug. The largest change (+33%) is shown by subject SD5, who was titrated with the highest dosage (3000 mg of methaqualone). In two of the subjects, SD4 and SD5, a marked increased in the mean frequency in the Delta Band accompanied the withdrawal of the drug (18 and 13% respectively). The delta rhythm was altered by the drug in a manner similar to the one described in Section 4.2.4, for the subjects under secobarbital. The three subjects in this group showed changes in the mean percentage of power in the Delta Band that were smaller when compared to the ones presented by the subjects that were withdrawn from secobarbital. On the other hand, the changes in the mean frequency in the band were essentially the same in both groups. The above results must be looked at with caution due to the differences not only in the amount of drug given to the subjects, but also in the withdrawal schedule for the two groups.

Most of the studies in the literature report either no significant change (Williams and Agnew, 1969; Kales et al, 1970; Risberg et al, 1975), or a little decrease in delta sleep (Goldstein et al, 1970; Itil et al, 1974) after the administration of methaqualone. In all these studies the subjects received no more than 300 mg of methaqualone.

In all three subjects, the average percentage of power in the Sigma Band decreased with the withdrawal of the drug. Subject SD5 showed the largest change (58%) and the others had changes around 40%. These changes, however, are smaller than the ones observed in two of the subjects under secobarbital (Section 4.2.4). The mean frequency in this band also increased between 3 and 7%, which is equivalent to a slowdown of 0.4-0.9 Hz in the frequency of the spindle activity due to the chronic administration of the drug.

From Table 4.20 we can also see that there is an increase in the average percentage of power in Theta Band with the withdrawal of the drug. However, an opposite trend is observed in the mean percentage of power in the Betal, Beta2 and FASTF Bands. Two of the subjects, SD4 and SD5, showed smaller changes in these last results when compared to the ones in the subjects withdrawn from secobarbital. This is probably due to the different time interval between the last and first nights that were compared in these two groups. As we can see from the last columns in Tables 4.12 and 4.15, the values for the mean percentage of power in the Beta1, Beta2 and FASTF Bands are the highest among all the subjects in the present study, with the exception of subject SC5.

This probably reflects the fact that although the subjects were not taking the drug anymore, their organisms did not have enough time to wash out the drug (Kay et al, 1976).

The mean frequency in each night decreases with the withdrawal of the drug. From Table 4.20 it can be seen that the subjects in this group showed reduction in this parameter between 21 and 33%. Again the highest change is observed in subject SD5. On the other hand, subject SD6, had a larger change than subject SD4, despite the fact that this subject was titrated with an amount of drug at least three times as large as that of subject SD6. A partial explanation for this discrepancy could be in the choice of the baseline night for subject SD4, as we already mentioned. We should also keep in mind that these subjects had a long history of polydrug abuse and that these discrepancies could also be the results of intersubject sensitivity to the drug.

The mean and the percentage of rejections of the x^2 coefficient decreased in all subjects with the administration of the drug. Table 4.20 shows these results as a percentage increase in the mean and in the percentage of rejections of the x^2 coefficient between the titration night and the night used as baseline (the last night that was processed for each subject). Subject SD5 had increases of 29 and 75% for the mean and for the percentage of rejection respectively. The other subjects had changes of 30% in the percentage of rejections and 15-20% in the mean value of x^2 coefficient with the withdrawal of the drug. As we can see

SUBJ	SD4 ¹	SD5 ²	SD 6		
TITRATION	DOSAGE (mg)	2100	3000	600
DELTA	PERC. OF POWER	MEAN	+22 %	+33%	+23%
(0.0-3.5Hz)	FREQ. PEAK (Hz)	MEAN	+18%	+13%	+ 7%
THETA	PERC. OF POWER	MEAN	+56%	+43%	+17%
BAND (4.0-7.5Hz)	FREQ. PEAK (Hz)	MEAN	- 6%	08	+1.4%
ALPHA	PERC. OF POWER	MEAN	+31%	-14%	+ 5%
(8.0-11.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 7%	+ 3%	08
. SIGMA	PERC. OF POWER	MEAN	-438	-58%	-428
(12.0-15.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 7%	+ 4%	+ 38

Summary of the Results of Tables 4.13, 4.16 and 4.19 for Subjects in Group 1 - Secobarbital

¹The results of the titration night were compared with the ones in the night following the first day of zero drug intake.

 2 The results of the titration night were compared with the ones in the night of the first day of zero drug intake.

(continued)

SUBJ	ECTS		SD4	SD5	SD6	
TITRATION	DOSAGE (mg)	2100	2100 3000		
BETAL	PERC. OF POWER	MEAN	<u>-</u> 53%	-55%	-72%	
BAND (16.0-20.5Hz)	FREQ. PEAK (Hz)	MEAN	08	08	+ 2%	
BETA2	PERC. OF POWER	MEAN	-49%	-54%	-80%	
BAND (21.0-29.5Hz)	FREQ. PEAK (Hz)	MEAN	- 1%	+0.4%	+ 2%	
FASTF	PERC. OF MEAN POWER		-32%	-30%	-83%	
BAND (30.0-40.0Hz)	FREQ. PEAK (Hz)	MEAN	08	- 2%	+ 1%	
2	MEAN	1	+20%	+29%	+15%	
X	PERCENTAGE OF REJECTIONS		+33%	+75%	+30%	
MFC	MEAN	I	-21%	-33%	-27%	

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from Table 4.15, during the nights close to the titration night of subject SD5, the EEG amplitude followed a Gaussian probability density function in more than 50% of the epochs. However, this was reduced to less than 20% when the subject stopped taking the drug. The other subjects in this group show the same trend but with less variability. As we suggested in Section 4.2.2, it seems that the EEG's amplitude distribution become more Gaussian with the administration of the drug and that this is also dosage dependent.

4.4 GROUP 2 - SECOBARBITAL

4.4.1 Subject SCl

4.4.1.1 Subject's Protocol

Subject SCl is a 22 year-old male. A total of four sleep sessions were automatically processed. Table 4.21 shows these sessions in a chronological order, their corresponding record number and the amount of drug given to the subject. This table was derived from Table 2.3, in which the complete procedure for the subject is shown. The baseline night was recorded one month after the first period.

TABLE 4.21

Analyzed Data for Subject SCl

Sleep	Record	Titration			
Sessions	Number	Dosage (mg)			
Nap ¹	15983	600			
lst Night ¹	15989				
1 Both sessions day.	were run	in the same			

(continued)

Sleep	Record	Titration
Sessions	Number	Dosage (mg)
2nd Night	15994	none
Baseline ² Night	16093	none
$2_{\text{Recorded on}}$ ministration	e month aft of the dru	er the ad- q.

4.4.1.2 Compressed Spectral Array

Figures 4.31a, b, c, and d show the CSA plots corresponding to the four sleep records listed in Table 4.21. For the nap, Figure 4.31a, the activity in the Delta Band is characterized by some high intensity spectral peaks at the beginning of the record. The spectral peaks in the Sigma Band are not well-defined in the beginning of the nap. In the 1st night, Figure 4.31b, the activity in the Delta Band is characterized by low intensity spectral peaks throughout the night. Well-defined peaks in the Sigma Band are the dominant activity for most of the night. The CSA plot for the 2nd night, Figure 4.31c, shows the return of the high intensity peaks in the Delta Band and that the magnitudes of the peaks in the Sigma Band have declined from their previous values. A similar pattern is also shown by the baseline night (Figure 4.31d).

4.4.1.3 <u>Summary of the Output Results from the FEATURE</u> <u>Algorithm</u>

Table 4.22 shows the output results from the data



FIGURE 4.31 Subject SC1 - Compressed Spectral Array (CSA) (a) nap, record #15983 (b) lst night, record #15989 (c) 2nd night, record #15994 (d) baseline night, record #16093

Summary of the Output Results from FEATURE for Subject SC1

SLEEP	SESSION	· · · · · · · · · · · · · · · · · · ·	Nap	lst	2nd	Baseline
RECOPD	NUMBER		15983	15989	15994	16093
DOSAG	E (mg)		600	-	0	0
DELTA	PERC. OF	MEAN	42.4	32.6	43.4	43.5
BAND	POWER	STD	16.9	10.8	13.7	10.8
(0 0-3 5Hz)	FREQ. PEAK	MEAN	1.6	1.8	1.7	1.7
	(Hz)	STD	0.5	0.5	0.5	0.6
milion a	PERC. OF	MEAN	22.9	27.1	33.4	34.7
BAND	POWER	STD	9.2	7.6	8.2	7.4
	FREQ. PEAK	MEAN	6.3	6.4	6.2	6.2
(4.0-7.5Hz)	(Hz)	STD	1.0	0.9	0.8	1.0
	PERC. OF	MEAN	13.7	13.2	11.4	11.0
ALPHA	POWER	STD	5.0	3.9	4.5	4.4
BAND	FREQ. PEAK	MEAN	10.2	9.1	9.5	8.9
(8.0-11.5Hz)	(Hz)	STD	1.2	1.3	1.2	0.7
CTCMA	PERC.	MEAN	9.9	17.1	5.9	4.9
BAND	POWER	STD	6.1	10.3	3.8	2.8
	FREQ. PEAK	MEAN	13.1	13.3	13.6	13.4
(12.0-15.5HZ)	(Hz)	STD	1.0	0.8	0.8	0.7

(continued)

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SLEEP	SESSION		Nap	lst	2nd	Baseline
RECORD NUMBER			15983	15989	15994	16093
DOSAG	E (mg)		600	_	0	0
BETAL	PERC. OF	MEAN	4.3	4.1	1.6	1.1
BAND	POWER	STD	2.4	1.9	0.7	0.4
(16.0-20.5Hz)	FREQ. PEAK	MEAN	18.2	18.5	18.7	18.4
((Hz)	STD	1.3	1.3	1.2	1.2
BETA 2	PERC. OF	MEAN	2.7	3.1	1.2	0.8
BAND	POWER	STD	1.7	1.8	0.6	0.3
	FREQ. PEAK	MEAN	23.7	23.7	23.8	24.1
(21.0-29.5Hz)	(Hz)	STD	1.9	2.0	2.2	2.3
EXCUE	PERC.	MEAN	0.5	0.8	0.4	0.3
BAND	POWER	STD	0.4	0.6	0.2	0.1
	FREQ.	MEAN	33.3	34.0	33.0	33.9
(30.0-40.0Hz)	(Hz)	STD	2.3	2.6	2.3	2.3
	MEA	N	79.5	79.4	86.8	87.0
x ²	STD		10.9	12.7	11.4	14.4
Α-	PERCEN OF REJECT	TAGE IONS	68.4	66.3	75.9	82.1
MFC	MEA	N	5.2	6.9	4.5	4.4
	STD		2.1	1.7	1.5	1.4

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETA1 BAND	BETA2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+ 33%	+ 28%	- 17%	- 71%	- 73%	- 748	- 63%	-	_
MEAN FREQ. PEAK	- 5%	- 3%	- 2%	+ 1%	-0.5%	- 2%	-0.3%	-	-
MEAN	-	-	-			-	-	+ 10%	- 36%
PERC. OF REJECTIONS	-	-	-	-	-	-	-	+ 24%	-

TABLE 4.23 Subject SCl - Percentage Change in the Results of Table 4.22

Between the Baseline and the First Nights

reduction algorithm. As we can see from this table, there is a tendency for the mean percentage of power in the Delta Band to decrease in the first night from their previous values during the nap. This tendency is however reversed in the 2nd and in the baseline nights. An upward trend is shown by the mean percentage of power in the Sigma, Betal and Beta2 Bands, and by the mean of MFC's between the nap and the 1st night. This trend is also reversed in the 2nd and in the baseline nights. The mean percentage of power in the Theta Band has it lowest value during the nap, increasing afterwards.

Figures 4.32a, b, c and d show the plots of the percentage of power in the Delta Band, for the sleep records shown in Table 4.21. For the nap, Figure 4.32a, there are two large peaks: one with a long duration at the beginning of the plot and the other towards the end of the nap. The mean percentage of power in the band is 42.4%. In the 1st night, Figure 4.32b, there is only one high amplitude peak with a short duration at the beginning of the night, and the mean percentage of power in the band has declined to 32.6%. By the 2nd night this value has increased to 43% which is close to the previous value computed for the nap. Figure 4.32c shows the corresponding plot for the 2nd night, in which two large peaks can be clearly seen in the first half of the night. In the baseline night, Figure 4.32d, there are three high amplitude peaks in the first half of the night and the mean percentage of power for the band

stayed at 43.5%. There is an increase of 33% in the mean percentage of power in the Delta Band, between the 1st and the baseline nights, as shown by Table 4.23. In the same period the mean frequency in the band has decreased 5%.

The plots for the percentage of power in the Sigma Band are shown in Figures 4.33a, b, c and d. The plot corresponding to the nap, Figure 4.33a, shows that the amplitude of the peaks is increasing as the nap progresses. The mean percentage of power in the band during the nap is 9.9%. In the 1st night the mean percentage of power has increased to 17.1% and its percentage of power plot, Figure 4.33b, shows high amplitude peaks most of the time. By the 2nd night, the mean percentage of power was reduced to 5.9%, and the number of peaks and their amplitudes have also diminished as illustrated in Figure 4.36c. In the baseline night the mean percentage of power in the band was 4.9%, and Figure 4.36d shows only few low amplitude peaks. From Table 4.23 we can see that there is a reduction of 71% in the percentage of power in the Sigma Band between the 1st and the baseline nights. In the same period the mean frequency has increased 1%.

Figures 4.34a, b, c and d show the plots for the Mean Frequency Coefficients. During the nap, Figure 4.34a, the mean value of the MFCs stayed above the 5 Hz line for most of the time, except for two low frequency dips that reached the 2.0 Hz frequency line, one of long duration at the beginning of the nap and the other towards its end. Figure 4.34b



(a) nap, record #15983
(c) 2nd night, record #15994

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(d) baseline night, record #16093

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shows that for the 1st night there is one low frequency dip reaching the 3 Hz frequency line and that the MFC's points stayed above 6 Hz for the remaining of the night. The mean frequency for the entire night was up to 6.9 Hz in contrast to the one from the previous night of 5.2 Hz. In the 2nd and in the baseline nights the mean frequency values were 4.5 Hz and 4.4 Hz respectively. Figures 4.34c and d show low frequency dips in the first half of the night. The mean frequency has decreased 36% between the 1st and the baseline nights, as shown in Table 4.23.

The plots for the X^2 coefficient are shown in Figures 4.35a, b, c and d. During the nap, Figure 4.35a, we can see that the hypothesis that the EEG's amplitude follows a Gaussian distribution was rejected in 68.4% of the epochs. This value has decreased to 66.3% in the 1st night, Figure 4.35b. In the 2nd and in the baseline nights the percentages of rejections have increased to 75.9 and 82% respectively. From Table 4.23 we can see that the percentage of rejections for the X^2 coefficient, in the baseline night, has increased 24% over the computed value for the 1st night.

4.4.2 Subject SC2

4.4.2.1 Subject's Protocol

Subject SC2 is a 23 year-old male. The sleep sessions that were automatically processed are listed in Table 4.24, which also shows their corresponding record number and the amount of drug taken by the subject. This table was derived from Table 2.3.

Sleep Sessions	Record Number	Titration Dosage (mg)
Nap ¹	15945	600
lst Night ¹	15952	_
2nd Night	15956	none
Baseline ² Night	16328	none
¹ Both sessions same day.	were run	in the same

Analyzed Data for Subject SC2

²Recorded one month after the administration of the drug.

4.4.2.2 Compressed Spectral Array

The CSA plots for subject SC2 are shown in Figures 4.36a, b, c and d. The CSA for the nap, Figure 4.36a, shows high intensity spectral peaks in the Delta and Sigma Bands. The later ones persisted throughout the 1st night, Figure 4.36b, however the peaks in the Delta Band are of less intensity than the ones in the nap plot. In the 2nd night, Figure 4.36c, there exist two trains of high intensity spectral peaks in the Delta Band, and the intensity of the peaks in the Sigma Band has been reduced a little. The CSA of the baseline night, Figure 4.36d, shows that the two trains of high intensity peaks in the Delta Band are still present and that the peaks in the Sigma Band have reached their lowest level for the four plots. The activity in the Alpha Band is located at 8.5 Hz in the 1st night, Figure 4.36b,



FIGURE 4.36 Subject SC2 - Compressed Spectral Array (CSA) (a) nap, record #15945 (b) 1st night, record #15952 (c) 2nd night, record #15956 (d) baseline night, record #16328

but it is shifted to 10 Hz in the 2nd and in the baseline nights (Figures 4.36c and d).

4.4.2.3 <u>Summary of the Output Results from the FEATURE</u> Algorithm

The output results from the data reduction algorithm are listed in Table 4.25. From there we can see that in the 1st night there is a decrease in the average percentage of power in the Delta Band and in the percentage of rejection of the X^2 coefficient in relation to the values computed during the nap. This trend is, however, reversed in the 2nd and in the baseline nights. The mean percentage of power in the Sigma, Betal, Beta2 and FASTF Bands and the average value of the MFCs increased from the nap to the 1st night and then started decreasing to their baseline values. The mean percentage of power in the Theta Band has its lowest value during the nap but it starts increasing after the 1st night.

The plots for the percentage of power in the Delta Band are shown in Figures 4.37a, b, c and d. From Figure 4.37a, we can see the presence of two high amplitude peaks in the first two thirds of the nap. The average percentage of power for this record is 43.5%. Only one large peak is present in the plot for the lst night, Figure 4.37b, and the mean percentage of power has decreased to 33.5%. In the 2nd night, Figure 4.37c, there are two high amplitude peaks in the beginning of the first and second halves of the

Summary of the Output Results from FEATURE for Subject SC2

SLEEP S	SESSION		Nap	lst	2nd	Baseline
RECORD	15945	15952	15956	16328		
DOSAG	E (mg)		600	-	0	0
DELTA	PERC. OF	MEAN	43.5	33.5	40.3	50.7
BAND	POWER	STD	8.4	8.5	11.0	9.4
(0, 0-3, 547)	FREQ. PEAK	MEAN	1.5	1.8	1.8	1.7
(0.0-3.3hz)	(Hz)	STD	0.5	0.6	0.7	0.7
MITTONA	PERC. OF	MEAN	15.6	16.3	21.4	22.4
BAND	POWER	STD	1.4	1.7	3.3	4.3
	FREQ. PEAK	MEAN	6.4	6.2	5.4	5.1
(4.0-7.5Hz)	(Hz)	STD	1.0	1.2	1.2	1.1
	PERC. OF	MEAN	9.5	10.0	10.0	7.5
ALPHA	POWER	STD	1.4	2.5	2.7	2.8
BAND	FREQ. PEAK	MEAN	9.8	9.5	9.7	9.8
(8.0-11.5Hz)	(Hz)	STD	1.2	1.1	1.2	1.1
STGMA	PERC.	MEAN	16.0	24.5	18.6	6.8
BAND	POWER	STD	6.5	9.4	10.0	3.4
	FREQ. PEAK	MEAN	13.3	13.7	13.6	13.9
(12.U-15.5HZ)	(Hz)	STD	0.6	0.7	0.6	0.7

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SLEEP	SESSION		Nap	lst	2nd	Baseline
RECORD	NUMBER		15945	15952	15956	16328
DOSAG	E (mg)		600	_	0	0
ר גייינו	PERC.	MEAN	5.7	6.3	3.5	2.5
BAND	POWER	STD	4.0	4.1	2.3	1.6
(16.0-20.5Hz)	FREQ. PEAK	MEAN	18.6	18.9	18.9	19.2
(10.0 20.512)	(Hz)	STD	1.1	1.1	1.3	0.9
BETLA 2	PERC.	MEAN	3.8	5.0	2.7	2.0
BAND	POWER	STD	2.0	3.5	2.0	1.8
(2) 0 20 57-	FREQ. PEAK	MEAN	23.9	25.1	24.9	24.9
(21.0-29.5HZ)	(Hz)	STD	2.0	2.3	2.5	3.1
FASTF	PERC. OF	MEAN	1.4	1.6	0.9	0.6
BAND	POWER	STD	0.9	1.3	0.6	0.5
	FREQ. PEAK	MEAN	33.3	33.2	33.5	34.6
(30.0-40.0H2)	(Hz)	STD	2.4	2.1	2.2	2.9
	MEA	N	93.9	85.4	93.5	91.7
x ²	STD		12.6	10.8	14.3	10.1
	PERCEN OF REJECT	PERCENTAGE OF REJECTIONS		80.7	88.2	88.7
MFC	MEA STD	N	6.3 1.9	8.5 2.0	6.7 2.2	4.9 2.0

	DELTA	THETA	ALPHA	SIGMA	BETA1	BETA2	FASTF	X ²	MFC
	DAND	BAND	DAND	BAND	DAND	DAND	DAND	COEF.	
MEAN PERC. OF POWER	+ 51%	+ 37%	- 25%	- 72%	- 60%	- 60%	- 56%	_	-
MEAN FREQ. PEAK	- 5%	- 18%	+ 3%	+1.5%	+1.6%	- 1%	+ 48	-	-
MEAN	-	-	-	-	-	-	-	+ 88	- 42%
PERC. OF REJECTIONS	_		-	_	-	-	_	+ 10%	-

TABLE 4.26 Subject SC2 - Percentage Change in the Results of Table 4.25

Between the Baseline and the 1st Nights

night respectively. The average percentage of power in this night has returned to the value of 40.3%. During the baseline night, Figure 4.37d, the two high amplitude peaks are still present with the difference being that they now have moved close to each other. The mean percentage of power in the baseline night is 50.7%, which is the highest value for all nights. From Table 4.26 we can see that the mean percentage of power in the Delta Band for the baseline night is 51% above the same quantity computed during the 1st night. The mean frequency in this band has decreased 5% between the first and the baseline nights.

The percentage of power in the Sigma Band is plotted, as a function of time, in Figures 4.38a, b, c and d. During the nap, Figure 4.38a, we can see that the amplitude of the peaks is becoming larger as the nap evolved. In the 1st night, Figure 4.38b, the amplitude and the number of peaks have increased in relation to the nap. The average percentage of power has also augmented from 16.0% during the nap, to 24.5%. Some high amplitude peaks still remain during the 2nd night, Figure 4.38c, but the mean percentage of power for this night has decreased to 18.6%. During the baseline night, Figure 4.38d, the high activity that was predominant in the first two nights has disappeared, which in turn resulted in a decrease in the average percentage of power to 6.8%. The total change in the mean percentage of power between the 1st and the baseline nights was -72%, as shown in Table 4.26. The mean frequency in this band has

increased by 1.5% in the same period.

Figures 4.39a, b, c and d show the Mean Frequency Coefficient (MFC) plotted as a function of time for the records listed in Table 4.24. During the nap, Figure 4.39a, there are two low frequency dips that are close to the 3 Hz frequency line in the first two third of the plot. The plot for the 1st night, Figure 4.39b, shows one low frequency dip above 3 Hz in the beginning of the night. The mean frequency for this night is up to 8.5 Hz from the value of 6.3 Hz during the nap. In the 2nd night, Figure 4.39c, there are two low frequency dips in the beginning of the first and second halves of the night respectively. Of these, the first one went below the 3 Hz frequency line. The mean frequency for this night is 6.7 Hz which is below the computed value for the previous night. The plot for the baseline night, Figure 4.39d, shows two low frequency dips in the first half of the night which are below the 3 Hz frequency line. The mean frequency for the night is down to 4.9 Hz. From table 4.26 it can be seen that the mean frequency in the baseline night has decreased 42% from its corresponding value during the 1st night. As we pointed out before the low frequency dips in these plots coincide with the delta peaks in the corresponding plots for the percentage of power in the Delta Band.

The plots for the X^2 coefficients are shown in Figures 4.40a, b, c and d. During the nap, Figure 4.40a, the EEG's amplitude did not follow **a** Gaussian probability dis-





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(d) baseline night, record #16328



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(c) 2nd night, record #15956

(d) baseline night, record #16328



- (c) 2nd night, record #15956
- (d) baseline night, record #16328







tribution function in 87% of the epochs. This value decreased to 80.7% in the 1st night and returned to its previous level in the 2nd and in the baseline nights (88.2 and 86.7 respectively). Table 4.26 shows that the mean value and the percentage of rejections for the x^2 coefficients have increased in the baseline night by 8 and 10% over their previous values in the 1st night.

4.4.3 Subject SC3

4.4.3.1 Subject's Protocol

Subject SC3 is a 24 year-old male. A total of three sleep sessions were automatically processed. Since this subject slept straight from the beginning of the nap until the next morning, the nap and the 1st night records are merged into a single long record. In order to facilitate the comparison of the results between the subjects, we divided the entire record into two parts of equal time duration and called them "nap" and "1st" night, respectively. Table 4.27 shows the sleep sessions, their corresponding record number and the amount of drug given to the subject.

TABLE 4.27

Analyzed Data for Subject SC3

Sleep	Record	Titra	Titration			
Session	Number	Dosage	e (mg)			
"Nap"l	15971	100	00			
"lst" Night ^l	15971	-				
¹ Both session record.	s are in	a single	e long			

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Sleep	Record	l Titration				
Sessions	Number	Dosage (mg)				
2nd Night	15979	none				
Baseline ² Night	16326	none				
² Recorded one month after the ad- ministration of the drug.						

4.4.3.2 Compressed Spectral Array

Figures 4.41a, b, c and d show the CSA plots of the four sleep sessions listed in Table 4.27. The CSA plot for the "nap", Figure 4.41a, shows some high intensity spectral peaks in the delta range at the beginning and towards the end of the nap. The spectral peaks in the Sigma Band are not well-defined in this plot and their central frequency is shifting between 13 and 14 Hz. Appearing with the first train of peaks in the Delta Band, there is a series of spectral peaks centered around 11.0 to 11.5 Hz which probably correspond to slow spindle activity "riding" on the delta waves. Activity in the Theta and Betas Bands can be also Figure 4.41b shows that in the "1st" Night, the high seen. intensity peaks in the Delta Band have disappeared. The peaks in the Sigma Band are better defined and most of them are centered around 13.5 Hz. Activity in the Theta and Betas Bands can still be seen.



FIGURE 4.41 Subject SC3 - Compressed Spectral Array (CSA) (a) "nap", record #15971 (b) "1st" night, record #15971 (c) 2nd night, record #15979 (d) baseline night, record #16326

The CSA plot for the 2nd night, Figure 4.41c, shows trains of high intensity delta-activity spectral peaks in the first half of the night and at the very end of it. The number of peaks in the Sigma Band has lessened in comparison to the ones in the first two plots and their peak frequencies are now between 14.0 and 14.5 Hz. Very little activity can be seen in the Beta's frequency range. For the baseline night, Figure 4.41d, the CSA plot shows that the peaks in the Delta Band are of higher intensity than the corresponding ones in the 2nd night, and the magnitude of the peaks in the Sigma Band remained essentially unchanged.

4.4.3.3 Summary of the Output Results from the FEATURE Algorithm

Table 4.28 shows the output results computed by the data reduction algorithm (FEATURE). As we can see from these results, the mean percentage of power in the Delta and Theta Bands decreased between the "nap" and the "lst" nights. On the other hand the average percentage of power in the Sigma, Betal, Beta2 and the mean value and the percentage of rejection of the X^2 coefficient increased in the same period. However an opposite trend is followed by all of the above results in the 2nd and in the baseline nights. Table 4.29 shows the percentage changes in all the results of Table 4.28 between the "lst" and the baseline nights.

Figures 4.42a, b, c and d show the plots of the percentage of power in the Delta Band for the sleep records

Summary of the Output Results from FEATURE for Subject SC3

SLEEP SESSION			Nap	lst	2nd	Baseline
RECORD	15971	15971	15979	16326		
DOSAGE (mg)			1000			
DELTA	PERC. OF	MEAN	36.1	25.0	37.5	58.3
BAND	POWER	STD	15.1	9.8	12.2	8.4
(0, 0, 2, 5H-)	FREQ. PEAK (Hz)	MEAN	1.4	1.5	1.5	1.3
(0.0-3.5HZ)		STD	0.4	0.4	0.4	0.3
mitema	PERC. OF	MEAN	13.2	13.0	17.5	16.3
BAND	POWER	STD	3.5	3.3	4.2	2.8
	FREQ. PEAK	MEAN	6.7	6.6	6.6	7.0
(4.0-7.5Hz)	(Hz)	STD	0.6	0.8	0.9	0.8
	PERC. OF	MEAN	14.4	14.4	12.5	9.6
ALPHA	POWER	STD	3.0	2.0	3.3	4.1
BAND	FREQ.	MEAN	10.6	9.9	10.0	9.7
(8.0-11.5Hz)	(Hz)	STD	1.1	1.1	1.3	1.0
CTCMA	PERC. OF POWER	MEAN	18.9	21.8	13.5	5.7
BAND		STD	9.0	8.4	6.2	2.7
	FREQ. PEAK (Hz)	MEAN	13.5	13.7	14.2	14.2
(12.0-15.5Hz)		STD	0.8	0.7	0.8	0.6
(continued)

SLEEP	SESSION		Nap	lst	2nd	Baseline
RECORD	NUMBER	· · · · · · · · · · · · · · · · · · ·	15971	15971	15979	16326
DOSAG	E (mg)		1000			
ר השבום	PERC.	MEAN	8.8	12.8	6.9	1.83
BAND	POWER	STD	5.8	3.9	3.5	0.9
	FREQ.	MEAN	18.6	18.4	18.8	18.8
(16.0-20.5HZ)	(Hz)	STD	1.4	1.5	1.3	1.0
	PERC.	MEAN	9.2	10.1	6.5	1.5
BAND	POWER	STD	0.6	3.5	4.1	1.1
	FREQ. PEAK	MEAN	23.4	23.6	24.5	24.7
(21.0-29.5Hz) PEA (Hz	(Hz)	STD	2.1	2.0	2.7	2.5
	PERC.	MEAN	0.4	0.9	0.7	0.4
BAND	POWER	STD	0.2	0.4	0.5	0.3
	FREQ.	MEAN	34.4	33.9	34.6	34.7
(30.0-40.0Hz)	(Hz)	STD	2.3	2.2	2.5	2.8
	MEA	N	65.9	61.2	83.9	105.5
x ²	STD		9.4	7.1	15.7	23.0
A	PERCEN OF REJECT	TAGE IONS	38.5	35.6	71.0	94.3
MFC	MEA STD	N	6.5 2.5	9.0 2.0	6.7 2.1	3.5 1.2

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETA1 BAND	BETA 2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+133%	+ 18%	- 33%	- 73%	- 85%	- 84%	- 56%	-	-
MEAN FREQ. PEAK	- 13%	+ 6%	- 2%	- 48	+ 2%	+ 5%	+ 2%	-	-
MEAN	-	-	-	-	-	_		+ 72%	- 61%
PERC. OF REJECTIONS	_		-	_	-	_	-	+165%	-

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TABLE 4.29 Subject SC3 - Percentage Change in the Results of Table 4.28

Between the Baseline and the 1st Nights

listed on Table 4.27. During the "nap", Figure 4.42a, there are three large peaks, of which two are located at the first half of the plot and the other towards its end. The first peak is of higher amplitude and longer duration compared to the other two. The mean percentage of power in the band for the "nap" is 36.1%. The plot for the "lst" night, Figure 4.42b, shows only one high amplitude peak located close to the center of the plot, and the mean percentage of power in the band is down to 25%. By the 2nd night this value is up to 37.5%, which is close to the computed value during the "nap". The plot for the 2nd night, Figure 4.42c, shows four high amplitude and short duration peaks, two in the first third and two in the last third of the night. In the baseline night, Figure 4.42d, the mean percentage of power is up to 58.3% and only two large amplitude peaks can be clearly distinguished agains the high activity background. From Table 4.28 it can be seen that the mean percentage of power in the Delta Band during the baseline night has increased 133% over the corresponding value computed for the "1st" night. The mean frequency in this band has decreased 13% between these two nights.

The plots for the percentage of power in the Sigma Band are shown in Figure 4.43a, b, c and d. Figure 4.43a, shows that during the "nap" the amplitude of the peaks in the band is increasing as the "nap" progresses. The mean percentage of power in the band is 18.9%. This value has increased to 21.8% during the 1st night, Figure 4.43b. However the peaks in the 2nd night, Figure 4.43c, are of less amplitude than the ones in the previous plots, and the mean percentage of power for the night is down to 13.5%. In the baseline night, Figure 4.43d, the average percentage of power is reduced to 5.7% and the peaks are of low amplitude. There is a reduction of 73% between the mean percentage of power in the "lst" night and the one in the baseline night. The mean frequency in this band has also increased 4% between the two nights.

Figures 4.44a, b, c and d show the plots for the Mean Frequency Coefficients (MFC) for the records as listed in Table 4.27. The plot for the "nap," Figure 4.44a, shows two low frequency dips. The first which is located at the beginning of the "nap" is of longer duration and it is closer to the 2 Hz frequency line. The second one is in the last third of the "nap", and its frequency is above the 3 Hz frequency line. After the first dip the MFC points are of increasing frequencies as the nap progresses. During the "1st" night, Figure 4.44b, most of the MFCs are located at the higher frequency part of the plot, except for a low frequency dip around the middle of the night that is below the 3 Hz line. The mean frequency now has increased to 9 Hz. In the 2nd night this is down to 6.7Hz, which is close to the previous value for the "nap". The plot for this night, Figure 4.44c, shows three low frequency dips that are either close or below the 3 Hz frequency line. Two of them are located in the first third of the night and the other at the



(a) "nap", record #15971 (b) "1st" night, record #15971 (c) 2nd night, record #15979

(d) baseline night, record #16326



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(a) "nap", record #15971 (b) "1st" night, record #15971 (c) 2nd night, record #15979 (d) baseline night, record #16326



- (a) "nap", record #15971 (b) "1st" night, record #15971
- (c) 2nd night, record #15979

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(d) baseline night, record #16326

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very end of it. The baseline night, Figure 4.44d, is dominated by low frequency activity and the low frequency dips are not well-defined. The mean frequency for this night is down to 3.5 Hz. According to Table 4.29, this result is 61% below the value computed for the "1st" night.

Figures 4.45a, b, c and d show the plots for the x^2 coefficients. In the first two plots, Figures 4.45a and b, most of the points are below the 72 line and the EEG's amplitude distribution follows a Gaussian probability density function in more than 60% of the epochs. However, this percentage is reduced to less than 30% in the 2nd night and to 5% in the baseline night. The last two plots, Figure 4.45c and d, show that the X^2 points are more scattered than in the previous plots (Figures 4.45a and b). This can also be seen by an increase in the standard deviation in these last two records. From Table 4.29 we can see that the mean value and the percentage of rejections for the x^2 coefficient have increased 72 and 165% between the "lst" and the baseline nights.

4.4.4 Discussion of the Results of Subjects in Group 2 Receiving Secobarbital

Table 4.30 shows the comparisons of the percentage changes in the output results of the data reduction algorithm between the baseline night and the first night for each of the subjects in Group 2, that were given secobarbital.

The mean percentage of power in the Delta Band decreased in all subjects during the first night when they were under the effects of the drug. Table 4.30 shows this as an increase in the mean percentage of power in the baseline night compared to the corresponding value during the first night. Karacan et al (1970) has found that afternoon naps, during which there was significantly more stage 4 sleep, were followed by nights during which stage 4 was reduced below normal levels. Since we cannot correlate especifically stage 4 with the percentage of power in the Delta Band, we can only speculate that probably part of the reduction in the percentage of power in the Delta Band during the first night could be caused by the fact that the subjects took a nap following the administration of the drug. Subject SC3, to whom was given the highest dosage (1000 mg of secobarbital) had the largest increase (133%). The other two subjects who were given 600 mg of secobarbital showed increases in the mean of 33 and 15%. In addition, one of the subjects, SC3, showed a marked decrease in the mean frequency peak in this band. This later results is in contradiction with the previous results obtained for the subjects in Group 1 that were withdrawn from the same drug (Section 4.2.4).

The average percentage of power in the Sigma Band decreased below 70% in all the subjects during the baseline night, when compared to the results obtained in the first night. Two of the subjects, SCl and SC2, showed a decrease in the mean frequency peak in this band during the first night. However, only subject SC3 had a frequency change (4%) compared to the ones observed in the drug abusers (Table 4.10).

From Table 4.30 we also can see that there is a reduction in the average percentage of power in the Alpha, Betal, Beta2 and FASTF frequency bands in the baseline night when compared to the same results during the first night. However an opposite trend is observed in the mean percentage of power in the Theta Band.

The mean frequency during the baseline night decreased in all subjects from the previous high values computed for the first night. The highest reduction (61%) was shown by subject SC3. The other two subjects showed changes between -36 and -42%.

The mean value and the percentage of reduction of the X^2 coefficient decreased in all subjects with the administration of the drug. Table 4.30 shows these results as a percentage increase in the mean value and in the percentage of rejections in the X^2 coefficient between the first and the baseline nights. Again subject SC3, showed the highest increases of 72% for the mean and 165% for the percentage of rejections. Furthermore, as we can see from Table 4.28, during the first night in almost 65% of the epochs the EEG's amplitude was distributed according to a Gaussian probability density function. This value was reduced

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SUBJ	ECTS	SC1	SC2	SC3	
TITRATION	DOSAGE (mg)	600	600	1000
DELTA	PERC. OF POWER	MEAN	33%	+51%	+133%
(0.0-3.5Hz)	FREQ. PEAK (Hz)	MEAN	- 5%	- 5%	- 13%
THETA	PERC. OF POWER	MEAN	+25%	+37%	+ 18%
BAND (4.0-7.5Hz)	FREQ. PEAK (Hz)	MEAN	- 38	+ 3%	+ 6%
ALPHA	PERC. OF POWER	MEAN	-17%	-25%	- 33%
(8.0-11.5Hz)	FREQ. PEAK (Hz)	MEAN	- 28	+ 3%	- 2%
SIGMA	PERC. OF POWER	MEAN	-71%	-72%	- 73%
(12.0-15.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 1%	-1.5%	+ 4%

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Summary of the Results of Tables 4.23, 4.26 and 4.29 for Subjects in Group 1 - Secobarbital

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SUBJ	ECTS		SC1	SC2	SC3
TITRATION	DOSAGE (1	mg)	600	600	1000
BETAl	PERC. OF POWER	MEAN	-73%	-60%	-85%
BAND (16.0-20.5Hz)	FREQ. PEAK (Hz)	MEAN	-0.5%	+1.6%	+ 2%
BETA2	PERC. OF POWER	MEAN	-748	-60%	-84%
BAND (21.0-29.5Hz)	FREQ. PEAK (Hz)	MEAN	- 28	- 1%	+ 5%
FASTF	PERC. OF POWER	MEAN	-63%	-56%	-56%
BAND (30.0-40.0Hz)	FREQ. PEAK (Hz)	MEAN	-0.3%	+ 48	+ 2%
2	MEAN	1	+10%	+ 88	+72%
X	PERCENT OF REJECT I	'AGE IONS	+24%	+10%	+165%
MFC	MEAN	I	- 36%	-42%	-61%

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to less than 5% in the baseline night. The other subjects followed the same trend but with much less variability.

4.5 GROUP 2 - METHAQUALONE

4.5.1 Subject SC4

4.5.1.1 Subject's Protocol

Subject SC4 is a 25 year-old male. Table 4.31 lists the sleep sections that were automatically processed. This table was derived from Table 2.4. Record #15912, for the lst night, showed a very low frequency artifact that was present for most of the night. It was possibly caused by the subject's sweating or by the fact that blood samples were drawn during the night. This artifact resulted in large spectral peaks around 0 Hz in the CSA plot. In order to minimize its effects, the raw EEG data was passed through a 9th order high-pass Chebyshev digital filter (Papp, 1975) with a cutoff frequency at 1 Hz, before the algorithm REDUCE was applied to it (See Section 2.4.2).

TABLE 4.31

Analyzed Data for Subject SC4

Sleep	Record	Titration			
Sessions	Number	Dosage (mg)			
Nap ¹	15909	4200			
lst Night ^l	15912	-			
l Both sessions day.	were run	in the same			

TABLE 4.	21	
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Sleep	Record	Titration
Sessions	Number	Dosage (mg)
2nd Night	15917	none
Baseline ² Night	16070	none
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Recorded one month after the administration of the drug.

4.5.1.2 Compressed Spectral Array

Figures 4.46a, b, c and d show the CSA plots for the sleep sessions listed on Table 4.31. The CSA plot for the nap, Figure 4.46a, shows some high intensity spectral peaks in the Delta, Alpha and Sigma Bands. The peaks in the Alpha Band are centered around 8.5-9.0 Hz and the ones in the Sigma Band are between 12.5 and 13.0 Hz. In the 1st night, Figure 4.46b, the peaks in the Delta Band are of lower intensities than the ones during the nap and the activity in the Sigma Band is present most of the night, centered at 13.0 Hz. The peaks in the Alpha Band are still located at 8.5-9.0 Hz. Figure 4.46c, shows the CSA plots corresponding to the 2nd night. From there we can see that the main change from the previous plots is an increase in the amplitude of the peaks in the Delta, Sigma, and Theta Bands. Furthermore, this increase is much larger in the Delta Band. During the baseline night, Figure 4.46d, there exist very high amplitude peaks in the Delta Band and the number of peaks in the Sigma





Band has diminished in comparison to the 2nd night. The peaks in the Alpha Band are now located at 10 Hz, and some activity can also be seen in the Theta Band.

4.5.1.3 <u>Summary of the Output Results from the FEATURE</u> Algorithm

In Table 4.32, the results of the data reduction analysis performed by FEATURE are given. From this table it can be seen that there is a decrease in the percentage of rejections for the x^2 coefficient in the 1st night in relation to the values previously computed for the nap. This trend is, however, reversed in the 2nd and in the baseline nights. The mean percentage of power in the Sigma, Betal, Beta2 and Fastf Bands, and the mean value of the increase from the nap to the 1st night, and then MFCs start decreasing to their baseline values. The mean percentage of power in the Theta Band reached its lowest value during the nap, but it starts increasing in the subsequent The mean percentage of power in the Delta Band stays nights. almost the same during the nap, and in the 1st night, but it increases in the baseline night. Table 4.33 shows the percentage change in the results of Table 4.32 between the 1st and the baseline nights.

In Figures 4.47a, b, c and d the plots of the percentage of power in the Delta Band vs time are shown. Only one peak is present at the beginning of the nap, Figure 4.47a. The plot for the 1st night, Figure 4.47b, shows three high

Summary of the Output Results from FEATURE for Subject SC4

SLEEP	SESSION		Nap	lst	2nd	Baseline
RECORD	NUMBER		15909	15912	15917	16090
DOSAG	E (mg)	.	4200	_	0	0
DELTA	PERC. OF	MEAN	42.0	41.5	43.7	51.9
BAND	POWER	STD	15.1	9.6	15.2	11.4
(0, 0=3, 547)	FREQ. PEAK	MEAN	1.5	1.7	1.4	1.5
	(Hz)	STD	0.6	0.4	0.5	0.5
	PERC. OF	MEAN	15.5	18.3	19.1	20.9
BAND	POWER	STD	3.7	5.0	6.3	5.8
	FREQ. PEAK	MEAN	6.5	6.7	6.3	6.6
(4.0-7.5Hz)	(Hz)	STD	1.0	0.8	1.2	1.0
	PERC. OF	MEAN	14.9	12.9	14.0	12.4
ALPHA	POWER	STD	9.8	4.5	6.9	7.3
DAND	FREQ. PEAK	MEAN	9.1	9.0	8.6	8.4
(8.0-11.5Hz)	(Hz)	STD	0.7	1.0	0.6	0.6
ST(M)	PERC.	MEAN	12.0	15.6	12.4	5.7
BAND	POWER	STD	4.6	5.6	5.9	2.4
	FREQ. PEAK	MEAN	12.9	13.1	13.1	13.6
(12.U-15.5HZ)	(Hz)	STD	0.6	0.6	0.6	0.6

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SLEEP	SESSION		Nap	lst	2nd	Baseline
RECORD	NUMBER		15909	15912	15917	16090
DOSAG	E (mg)		4200	-	0	0
BETAL	PERC. OF	MEAN	4.5	4.6	2.8	2.1
BAND	POWER	STD	2.5	1.8	1.8	1.1
(16 0-20 547)	FREQ. PEAK	MEAN	18.4	18.7	18.7	18.5
(10.0-20.5h2)	(Hz)	STD	0.8	1.2	1.3	1.3
ר געשים	PERC.	MEAN	4.4	4.8	2.6	1.5
BAND	POWER	STD	2.4	2.0	1.9	0.9
	FREQ. PEAK	MEAN	24.3	22.9	24.2	24.7
(21.0-29.5Hz)	(Hz)	STD	1.9	2.1	1.9	2.4
53.000	PERC.	MEAN	1.0	1.1	0.7	0.5
BAND	POWER	STD	0.4	0.4	0.5	0.3
	FREQ.	MEAN	34.2	34.0	33.3	34.1
(30.0-40.0Hz)	(Hz)	STD	2.4	2.4	2.4	2.4
	MEA	N	71.0	67.1	88.6	88.5
x ²	STD		10.8	9.8	15.5	12.8
	PERCEN OF REJECT	TAGE IONS	52.0	46.7	4.6 2.8 1.8 1.8 1.8 1.8 8.7 18.7 1.2 1.3 4.8 2.6 2.0 1.9 2.9 24.2 2.1 1.9 1.1 0.7 0.4 0.5 4.0 33.3 2.4 2.4 7.1 88.6 9.8 15.5 6.7 79.6 6.2 5.3 1.2 1.9	83.9
MFC	MEA STD	N	5.9 2.2	6.2 1.2	5.3 1.9	3.8 1.4

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETAl BAND	BETA 2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+ 26%	+ 14%	- 48	- 63%	- 54%	- 69%	- 55%	_	-
MEAN FREQ. PEAK	- 12%	-1.5%	- 78	+ 48	+ 1%	+ 3%	+0.3%	-	-
MEAN	-	-	-	-	-	-	-	+ 32%	- 39%
PERC. OF REJECTIONS	-		-	-	-	-	-	+ 80%	-

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TABLE 4.33 Subject SC4 - Percentage Change in the Results of Table 4.32

Between the Baseline and the 1st Nights

amplitude peaks, two in the first half and the other in the last third of the night. The average percentage of power in this night has remained essentially the same computed for the nap (41.5 and 42% respectively). In the 2nd night, Figure 4.47c, there are two high amplitude peaks in the first and in the 2nd half of the night and the average percentage of power in the band has slightly increased to 43.7%. In the plot for the baseline night, Figure 4.47d, three high amplitude peaks can be seen, two in the first half and the other in the last third of the night. The mean percentage of power is now 51.9%, which represents an increase of 26% over the value computed in the 1st night, as shown in Table 4.33. The mean frequency in the band has decreased 12% in the same interval. The change in the mean percentage of power in the band, for the amount of drug administered, is small compared to the other subjects in the group. This could be a result of the high background activity in the Delta Band caused by the artifact mentioned in Section 4.5.1.1 which was not totally eliminated by the filter.

The plots for the percentage of power in the Sigma Band are shown in Figures 4.48a, b, c and d. During the nap, Figure 4.48a, it can be seen that the amplitude of the peaks is becoming larger as the nap evolved. In the 1st night, Figure 4.48b, the plot stays almost flat, for a long period of time in the first half of the night, reflecting the "constancy" of the activity in the Sigma Band. After the second half of the night, some well-defined peaks can be seen. The average percentage of power has increased to 15.5% from the previous value of 12% obtained during the nap. By the 2nd night this value has been reduced to 12.4% and only two high amplitude peaks can be seen in the plot (Figure 4.48c). In the baseline night, Figure 4.48d, the amplitude of the peaks has reached its lowest value and the average percentage of power in the band is down to 5.7%. From Table 4.33 we can see that the mean percentage of power in the Sigma Band has decreased 63% from the 1st to the baseline night. In the same period the mean frequency in the band has increased from 13.1 to 13.6 Hz (Table 4.32).

Figures 4.49a, b, c and d show the plots for the Mean Frequency Coefficients as a function of time. During the nap, Figure 4.49a, there is only one low frequency dip that went close to the 1.5 Hz frequency line. In the 1st night, Figure 4.49b, there are three low frequency dips located between the 4.5 and the 3.0 Hz frequency lines, coinciding in time with the delta peaks in Figure 4.47b. As we can see from this night's CSA plot (Figure 4.46b), there is a high background activity in the Delta Band and relatively low amplitude peaks in the Sigma Band. These two facts together result in a relatively small mean frequency for this night (6.2 Hz), when compared to the same quantity computed for other subjects, who took smaller dosages of the drug. Figure 4.49c, shows the plot for the 2nd night, in which two low frequency dips can be seen. The first is below the 3 Hz and the second reached the 1.5 Hz frequency line, which is mainly the result of an increase in the activity in the Delta Band. The mean frequency for this





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night decreased to 5.3 Hz. In the baseline night, Figure 4.49d, there exist three low frequency dips. Two of them are close to the 1.5 Hz frequency line. The mean frequency for the night is down to 3.8 Hz, reflecting the dominance of the low frequency activity. The mean frequency was reduced 39% between the 1st and the baseline nights as we can see from Table 4.33.

The plots for the X^2 coefficient are shown in Figures 4.50a, b, c and d. During the nap, Figure 4.50a, 52% of the epochs were rejected by the CSQ test. In the 1st night, Figure 4.50b, the great majority of the points are below 72 and only 46.7% of the epochs were rejected by the CSQ test. This means that the EEG amplitude in 53% of the epochs follows a Gaussian probability density function. As the subject is recovering from the drug, the x^2 values start increasing as we can see in Fgiures 4.50c and d. There is an increase of 80% in the percentage of rejections for the x^2 coefficients between the 1st and the baseline nights.

4.5.2 Subject SC5

4.5.2.1 Subject's Protocol

Subject SC5 is a 23 year-old male. This subject was titrated with 3000 mg of methaqualone according to the procedure described in Section 2.2.2. The subject slept straight from the afternoon of the titration day until the next morning and, therefore, the EEG data was stored in a single long record. In order to facilitate the comparisons of this subject's results with the others, we divided that record into two equal duration records, which we called "nap" and "lst" night respectively. The subject's 2nd night was not processed because its analog record was not available on magnetic tape. Table 4.34 shows the sleep sessions and their corresponding record numbers that were automatically processed.

TABLE 4.34

Analyzed Data for Subject SC5

Sleep	Record	Titration
Sessions	Number	Do sa ge (mg)
"Nap" ¹	15928	3000
"lst" Night ^l	15928	-
Baseline ² Night	16070	none
l Both session	s are in a	single long
record.		
² Recorded one	month aft	er the ad-
ministration	or the aru	1g.

4.5.2.2 Compressed Spectral Array Results

Figures 4.51a, b, and c show the CSA plots for the sleep sessions listed in Table 4.34. During the "nap", Figure 4.51a, the activity in the Delta Band is characterized by low amplitude spectral peaks for most of the time, except at the beginning and towards the end of the nap, when some high amplitude peaks are present. The peaks in Alpha Band are centered around 8.5-9.0 Hz. The Sigma Band



FIGURE 4.51 Subject SC5 - Compressed Spectral Array (CSA) (a) "nap", record #15928 (b) "lst" night, record #15928 (c) baseline night, record #16090

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shows well-defined high amplitude peaks located at 13.5-14.0 Hz and for most parts of the "nap" it is the dominant frequency band. Some activity can also be observed in the Beta Bands. The main difference in the CSA plot for the "1st" night, Figure 4.51b, compared to the one for the "nap", is the absence of high amplitude peaks in the Delta Band. In the baseline night, Figure 4.51c, we can see two trains of high amplitude peaks in the Delta Band in the first half of the night. The activity in the Sigma Band is greatly reduced in comparison to the previous plots and no visible activity can be seen in the Beta Bands.

4.5.2.3 <u>Summary of the Output Results from the FEATURE</u> Algorithm

Table 4.35 shows the output results from the data reduction analysis. From these results we can see that the drug affected the amount of the SWS, which resulted in a reduction in the mean percentage of power in the Delta Band. On the other hand, the drug also affected the high frequency activity by increasing the percentage of power in the Sigma, Betal, Beta2 and Fastf frequency bands. The above changes can be further summarized by the results of the average value of the MFCs for each sleep record.

Figures 4.52a, b and c show the plots of the percentage of power in the Delta Band vs time, for the sleep sessions listed on Table 4.34. During the "nap", Figure 4.52a, two high amplitude peaks can be seen, one in the first

Summary of the Output Results from FEATURE for Subject SC5

SLEEP	SESSION		Nap	lst	2nd	Baseline
RECOPD	NUMBER		15928	15928		16090
DOSAG	E (mg)		3000	-		0
DELTA	PERC. OF	MEAN	30.0	27.7		42.3
BAND	POWER	STD	9.3	6.8		11.7
(0 0-3 5Hz)	FREQ. PEAK	MEAN	1.6	1.9		1.7
(0.0-3.3nz)	(Hz)	STD	0.5	0.6		0.6
	PERC.	MEAN	14.4	19.0		22.8
THETA	POWER	STD	1.6	4.5		6.3
DIMD	FREQ. PEAK	MEAN	6.6	6.0		5.9
(4.0-7.5Hz)	(Hz)	STD	1.0	1.2		1.3
	PERC. OF	MEAN	11.5	11.7		12.0
ALPHA	POWER	STD	2.4	1.4		3.4
BAND	FREQ. PFAK	MEAN	9.5	9.3		9.2
(8.0-11.5Hz)	(Hz)	STD	1.0	1.0		0.9
CTCMA	PERC.	MEAN	24.0	21.8		11.7
BAND	POWER	STD	7.6	9.8		7.2
	FREQ. PEAK	MEAN	14.0	14.2		14.2
(12.0-15.5Hz)	(Hz)	STD	0.6	0.7		0.7

(continued)

SLEEP	Nap	lst	2nd	Baseline		
RECORD NUMBER			15928	15928		16090
DOSAGE (mg)			3000	-		0
BETA1 BAND	PERC.	MEAN	8.4	8.7		3.9
	POWER	STD	5.0	4.7		1.9
(16.0-20.5Hz)	FREQ. PEAK	MEAN	18.9	18.6		18.6
	(Hz)	STD	1.1	1.4		1.3
BETA2 BAND	PERC.	MEAN	6.8	7.0		2.8
	POWER	STD	3.3	4.2		1.7
(21.0-29.5Hz)	FREQ. PEAK (Hz)	MEAN	24.3	24.5		24.6
		STD	2.4	2.7		2.7
	PERC. OF	MEAN	1.9	2.1		0.9
BAND	POWER	STD	1.1	1.1		0.6
(30.0-40.0Hz)	FREQ. PEAK (Hz)	MEAN	33.3	33.7		34.7
		STD	2.0	2.4		2.8
x ²	MEAN STD		92.2	88.8		91.1
			17.0	13.7		12.5
	PERCENTAGE OF REJECTIONS		75.0	73.9		87.6
MFC	MEA STD	N	9.1 2.0	8.9 1.5		5.9 1.8

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETAl BAND	BETA2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+ 53%	+ 20%	+ 3%	- 46%	- 55%	- 59%	- 57%	-	-
MEAN FREQ. PEAK	- 11%	-1.7%	+ 3%	08	08	+0.4%	+ 38	-	_
MEAN	-	-	-	_	-	-	-	+ 3%	- 34%
PERC. OF REJECTIONS	-	-	-	-	-	-	-	+ 19%	-

TABLE 4.36 Subject SC5 - Percentage Change in the Results of Table 4.35

Between the "1st" and the Baseline Nights

third and the other in the last third of the plots, and the average percentage of power is 30%. These peaks are no more present during the "lst" night, Figure 4.52b, and the mean percentage of power decreases slightly to 27.7%. The plot for the baseline night, Figure 4.52c, shows three high amplitude peaks in the first half of the night. The mean percentage of power is now 42.3% which represents an increase of 53% over the previous value computed for the "lst' night (Table 4.36). From Table 4.35 we can see that the mean frequency in the band decreased from 1.9 Hz during the "lst" night to 1.7 Hz in the baseline night.

The plots for the percentage of power in the Sigma Band vs time are shown in Figures 4.53a, b and c. The first two plots, Figures 4.53a and b, are dominated by high amplitude peaks most of the time, and the mean percentage of power in the band is above 21% in both sleep sessions. In the baseline night, Figure 4.53c, this value is reduced to 11.7% and the majority of the peaks are of smaller amplitudes. From Table 4.36 we can see that the mean percentage of power in the Sigma/Band has decreased 46% between the lst and the baseline nights. However, the mean frequency in the band has not changed in the same period.

Figures 4.54a, b and c show the plots for the MFCs. The plot for the "nap", Figure 4.54a, shows two low frequency dips above the 4.5 Hz frequency line. However, most of the points in the plot are above 8 Hz. The computed mean frequency for the "nap" was 9.1 Hz. In the "lst" night the



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FIGURE 4.52 Subject SC5 - Percentage of Power in the Delta Band (a) "nap", record #15928 (b) "1st" night, record #15928 (c) baseline night, record #16090



FIGURE 4.53 Subject SC5 - Percentage of Power in the Sigma Band (a) "nap", record #15928 (b) "1st" night, record #15928 (c) baseline night, record #16090

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mean frequency remained high at 8.9 Hz and no low frequency dips can be seen in this plot. During the baseline night, the mean frequency went down to 5.9 Hz and three low frequency dips are present in the plot (Figure 4.54c). For this subject, the mean frequency in the baseline night has decreased 34% from its previous value during the "lst" night (Table 4.36).

The plots for the X^2 coefficients are shown in Figures 4.55a, b and c. As we can see from these plots, the percentage of rejections for the X^2 coefficients has its lowest values on the subject's titration day. This value increases on the baseline night. From Table 4.36 it can be seen that the percentage of rejections increased 19% between the "lst" and the baseline nights.

4.5.3 Subject SC6

4.5.3.1 Subject's Protocol

Subject SC6 is a 22 year-old male. The sleep sessions that were automatically processed and their record numbers are shown in Table 4.37. This table was derived from Table 2.4.

TABLE 4.37

Analyzed Data for Subject SC6

Sleep	Record	Titration
Sessions	Number	Dosage (mg)
Nap^1	15984	1800
lst Night ¹	15990	

¹The two sessions were run in the same day.

TABLE 4.37

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Sleep	Record	Titration
Sessions	Number	Dosage (mg)
2nd Night	15995	none
Baseline ² Night	16339	none
² Recorded one ministration	e month a of the d	after the ad- lrug.

4.5.3.2 Compressed Spectral Array

Figures 4.56a, b, c and d show the CSA plots corresponding to the sleep sessions listed on Table 4.37. During the nap, Figure 4.56a, the activity in the Delta Band is characterized by the presence of high amplitude peaks most of the time. The peaks in the Sigma Band have also high amplitude and are centered around 13.5-14.0 Hz. The high activity in the Sigma Band persisted throughout the 1st night, Figure 4.56b, but the activity in the Delta Band is greatly reduced. In the 2nd night, Figure 4.56c, there are two trains of high intensity peaks in the Delta Band and the peaks in the Sigma Band are of reduced amplitudes. In the baseline, Figure 4.56d, the peaks in the Delta Band are of even higher amplitudes and the activity in the Sigma Band remained low.

4.5.3.3 <u>Summary of the Output Results from the FEATURE</u> <u>Algorithm</u>

Table 4.38 shows the output results of the data re-





TABLE 4.38

Summary of the Output Results from FEATURE for Subject SC6

SLEEP S	SESSION		Nap	lst	2nd	Baseline
RECORD NUMBER			15984	15990	15995	16339
DOSAG	E (mg)		1800	-	0	0
DELTA	PERC. OF	MEAN	41.6	36.5	46.3	47.7
BAND	POWER	STD	8.4	8.0	13.9	13.8
(0 0-2 5ug)	FREQ. PEAK	MEAN	1.6	1.8	1.8	1.9
(0.0-5.5HZ)	(Hz)	STD	0.3	0.6	0.6	
	PERC.	MEAN	15.2	19.5	21.0	22.3
THETA	POWER	STD	2.2	4.5	5.8	5.8
BAND	FREQ. PEAK	MEAN	6.7	5.9	5.9	6.2
(4.0-7.5Hz)	(Hz)	STD	0.9	1.1	1.3	1.0
	PERC. OF	MEAN	11.0	13.3	12.4	13.4
ALPHA	POWER	STD	1.7	3.6	3.9	4.5
BAND	FREQ. PFAK	MEAN	10.0	9.4	9.0	9.0
(8.0-11.5Hz)	(Hz)	STD	1.1	0.9	0.9	0.9
CTCVD	PERC.	MEAN	18.7	19.0	13.7	9.3
BAND	POWER	STD	6.0	6.9	7.3	5.4
	FREQ.	MEAN	13.6	13.6	13.5	13.6
(12.0-15.5Hz)	(Hz)	STD	0.7	0.7	0.5	0.7
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SLEEP SESSION			Nap	lst	2nd	Baseline
RECORD NUMBER			15984	15990	15995	16339
DOSAGE (mg)			1800	-	0	0
BETAI	PERC. OF	MEAN	7.2	4.9	2.8	2.7
BAND	POWER	STD	3.6	2.5	1.6	2.0
(16 0-20 547)	FREQ.	MEAN	18.2	18.4	18.3	18.4
(10.0-20.5h2)	(Hz)	STD	1.5	1.2	1.5	1.2
DE(0) 2	PERC.	MEAN	3.9	2.5	1.7	1.7
BAND	POWER	STD	1.6	1.3	1.0	1.5
	FREQ. PEAK	MEAN	24.4	24.5	24.5	24.1
(21.0-29.5Hz)	(Hz)	STD	2.3	2.4	2.3	2.2
	PERC. OF	MEAN	1.0	0.6	0.5	0.4
BAND	POWER	STD	0.4	0.3	0.3	0.3
	FREQ.	MEAN	32.9	33.3	33.3	33.2
(30.0-40.0Hz)	(Hz)	STD	2.2	2.3	2.6	3.4
	MEA	N	80.1	79.9	82.6	88.2
x ²	STD		9.7	12.7	17.8	23.6
4	PERCENTAGE OF REJECTIONS		73.6	70.2	77.8	85.0
MFC	MEA STD	N	6.7 1.6	6.8 1.3	5.2 1.9	5.0 1.9

	DELTA BAND	THETA BAND	ALPHA BAND	SIGMA BAND	BETAL BAND	BETA 2 BAND	FASTF BAND	x ² COEF.	MFC
MEAN PERC. OF POWER	+ 27%	+ 14%	+ 1%	- 51%	- 45%	- 48%	- 33%	-	-
MEAN FREQ. PEAK	+ 6%	+ 5%	- 5%	08	08	- 2%	-0.3%	_	-
MEAN	-	-	-	_	-	_	-	+ 10%	- 27%
PERC. OF REJECTIONS	_	-	-	-	-	-	-	+ 21%	-

TABLE 4.39 Subject SC6 - Percentage Change in the Results of Table 4.38

Between the 1st and the Baseline Nights

duction algorithm. As we can see from this table, the results show essentially the same trends that were observed in the two previous subjects.

Figures 4.57a, b, c and d show the plots of the percentage of power in the Delta Band vs time. The plot for the nap, Figure 4.57a, shows three high amplitude peaks, two located in the first half and the other in the last half of the nap. The mean percentage of power during the nap is 41.6%. This quantity is down to 36.5% in the 1st night. The plot for this night, Figure 4.57b, shows two high amplitude peaks in the first half of the night. Of these, the first one is of longer duration. The mean percentage of power has now increased to 46.3%. During the baseline night, Figure 4.57d, we can see a very high amplitude long duration peak at the beginning of the night and the mean percentage of power stayed at 47.7%. Table 4.39 shows that the mean percentage of power in the baseline night is 27% above the same quantity during the 1st night.

The plots for the percentage of power in the Sigma Band are shown in Figures 4.58a, b, c and d. As we can see from these plots, the mean percentage of power is high during the nap and in the 1st night and then it starts decreasing in the 2nd night to reach its lowest value in the baseline night. From Table 4.39 we can see that the mean percentage of power in the Sigma Band has decreased 51% in the baseline night in comparison to the value computed in the 1st night. -1

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(a) nap, record #15984
(b) 1st night, record #15990
(c) 2nd night, record #15995
(d) baseline night, record #16339











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(c) 2nd night, record #15995

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(d) baseline night, record #16339



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The plots for the Mean Frequency Coefficient vs time are shown in Figures 4.59a, b, c and d. The plot for the nap, Figure 4.59a, shows three low frequency dips between 4.5 and 3 Hz, and that the mean frequency for the night is 6.7 Hz. This quantity is at 6.8 Hz in the 1st night, and the plot for this night, Figure 4.59b shows a low frequency dip between4.5 and 3 Hz. In the 2nd night, Figure 4.59c, there are two low frequency dips between 3 and 1.5 Hz. The mean frequency for this night is now 5.2 Hz. This value is down to 5 Hz during the baseline night, Figure 4.59d, in which there is a low frequency dip that went down to the 1.5 Hz frequency line. There is a decrease of 27% in the mean frequency between the 1st and the baseline nights (Table 4.39).

Figures 4.60a, b, c and d show the plots for the x^2 coefficient. During the nap, Figure 4.60a, we can see that the hypothesis that the EEG's amplitude follows a Gaussian distribution was rejected in 73.6% of the epochs. This value has decreased to 70.2% in the 1st night, Figure 4.60b. In the 2nd and in the baseline nights the percentage of rejections have increased to 77.8 and 85% respectively. From Table 4.39 we can see that the percentage of rejections for the x^2 coefficient, in the baseline night, has increased 21% over the same quantity in the 1st night.

4.5.4 Discussion of the Results of the Subjects in Group 2 Receiving Methaqualone

Table 4.40 shows the comparison of the percentage

changes in the output results of the data reduction algorithm between the baseline night and the first night for each of the subjects in Group 2, that were given methaqualone.

The mean percentage of power in the Delta Band increases between the first and the baseline night, in all three subjects. This is equivalent to saying that the mean percentage of power decreases in the first night when the subjects were under the effects of the drug. As we pointed out in section 4.4.4 this could be partially caused by the fact that the subjects had an afternoon nap preceeding the first night. Subject SC3 who received 3000 mg of methagualone had the largest increase (53%). However, subject SC4 who was given the highest dosage (4200 mg) showed an increase of 26%. This could be the result of the low frequency artifact that was present during the first night (see Section 4.5.1). In addition, two of the subjects, SC4 and SC5, showed a marked increase in the mean frequency peak in this band during the first night. This is shown in Table 4.40 as a percentage decrease in the mean frequency peak in the baseline night when compared to the same quantity during the first night. These results are in contradiction with the previous ones obtained for the subjects in Group 1 that were withdrawn from the same drug (section 4.3.4).

In all three subjects the average percentage of power in the Sigma Band decrease in the baseline night from the values computed during the 1st night. The highest change is shown by subject SC4 (-63%) and the other subjects have changes of 46 and 51%. These results are close enough to the ones obtained from the abuser group that were withdrawn from the same drug. Only subject SC4 showed a decrease in the mean frequency peak during the first night. This result is in agreement with the ones observed in the abusers group.

From Table 4.40 we also can see that there is a reduction in the average percentage of power in the Betal, Beta2 and Fastf frequency bands in the baseline night in comparison to the first night. However, an opposite trend is observed in the mean percentage of power in the Theta Band.

The mean frequency during the baseline night (computed as the mean of the MFCs) decreased in all subjects from the previous high values calculated in the first night. The highest reduction (39%) was shown by subject SC4, the other two subjects had changes between -27 and -34%. These results compare well with the ones computed for the abusers group (Section 4.3.4), but in absolute values these changes are smaller than the ones for the subjects in Group 2, that were given secobarbital.

The mean value and the percentage of rejection of the X^2 coefficient decreased in all subjects with the administration of the drug. Table 4.40 shows these results as a percentage increase in the mean value and in the percentage of reduction in the X^2 coefficient between the first and the baseline nights. Subject SC4 shows the highest increase in both the mean and in the percentage of rejection (32 and 80% respectively). The other two subjects show the same trend but with much smaller changes.

TABLE 4.40

SUBJ	SC4	SC5	SC6		
TITRATION	DOSAGE (mg)	4200	3000	1800
DELTA BAND (0.0-3.5Hz)	PERC. OF POWER	MEAN	+26%	+53%	+27%
	FREQ. PEAK (Hz)	MEAN	-12%	-11%	+ 6%
THETA	PERC. OF POWER	MEAN	+14%	+20%	+14%
BAND (4.0-7.5Hz)	FREQ. PEAK (Hz)	MEAN	-1.5%	-1.7%	+ 5%
ALPHA	PERC. OF POWER	MEAN	- 48	+ 3%	+ 1%
(8.0-11.5Hz)	FREQ. PEAK (Hz)	MEAN	- 7%	+ 3%	- 5%
SIGMA	PERC. OF POWER	MEAN	-63%	-46%	-51%
(12.0-15.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 4%	0%	0%

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Summary of the Results of Tables 4.33, 4.36 and 4.39 for Subjects in Group 1 - Secobarbital

TABLE 4.40

(continued)

SUBJ	SC4	SC5	SC6		
TITRATION	DOSAGE (4200	3000	1800	
BETAl	PERC. OF POWER	MEAN	-54%	-55%	-45%
BAND (16.0-20.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 1%	08	0%
BETA2	PERC. OF POWER	MEAN	-69%	-59%	-48%
BAND (21.0-29.5Hz)	FREQ. PEAK (Hz)	MEAN	+ 3%	+0.4%	- 2%
FASTF	PERC. OF POWER	MEAN	-55%	-578	-338
BAND (30.0-40.0Hz)	FREQ. PEAK (Hz)	MEAN	+0.3%	+ 3%	-0.3%
2	MEAN		+32%	+ 38	+10%
X	PERCENTAGE OF REJECTIONS		+80%	+19%	+21%
MFC	MEAN	I	-39%	-34%	-27%

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CHAPTER V

CONCLUSIONS AND RECOMMENDATIONS

5.1 CONCLUSIONS

Before we start this section we want to make it clear that whenever we refer to the activity or to the frequency in one of chosen frequency bands (Delta, Theta, Alpha, Sigma, Betal, Beta2 and Fastf) we are talking from the point of view of spectral analysis. These parameters are not necessarily the same ones for Electroencephalographers, since they look at the EEG waves in terms of periods and amplitudes. For example, a spectral peak in the Delta Band for us can be the result of: (1) delta activity as the visual scorer sees it; (2) activity with the same period as the delta, but whose amplitude does not reach the $75\mu V$ peak-to-peak threshold required to be called delta (Rechtschaffen and Kales, 1968); (3) some low frequency artifact; (4) amplitude modulation in the EEG waveform; (5) any combination of the above.

From the results of Chapter IV we can see that the administration of either drug reduced the mean percentage of power in the Delta Band in all subjects. This value, however, increased with the withdrawal of the drug as in the case of the subjects in Group 1, or returned to its normal value during the baseline night for subjects in Group 2. The highest reductions in this parameter were observed in subjects SDl and SC3, who received the largest dosages of secobarbital (1000 mg). The subjects with the highest titration dosages of methaqualone did not show this kind of dosage related changes, except for subject SC5. Since with the other subjects that were also titrated with large doses of methaqualone, we had problem with either choosing their baseline nights (SD4 and SD5), or with a low frequency artifact present in the first night (SC4), we cannot say, based on the available data, that the mean percentage of power in the Delta Band is less affected by methaqualone than it is by secobarbital.

All the subjects in Group 1 showed a decrease in the mean frequency in the Delta Band with the administration of the drug. Furthermore, four of the subjects (SD1, SD3, SD4, and SD5) had a marked decrease in this parameter. This is equivalent to saying that the drugs caused not only a reduction in the percentage of power in the Delta Band, but that they also slowed down the "frequency" of the "delta waves". On the other hand, the subjects in Group 2, with the exception of subject SC6, had an increase in the mean frequency of the Delta Band with the acute administration of the drugs. This discrepancy could be explained by the fact that the subjects in Group 1 were all chronic drug abusers who have been using these drugs for quite some time. To date we have not found any references in the literature related to these frequency shifts.

As we already pointed out in section 4.2.4, the energy in a given frequency band is a function of the EEG amplitude, as well as of the amount of activity present in that particular frequency. In this manner, spectral analysis

is a blunt tool in the sense that it cannot tell us if the reduction of power in a given band is due to the reduction in the amplitude of the EEG waves in that frequency, or to the decrease in their total number or both. A good illustration of this was shown in section 2.4.3, example 1, when a low frequency high amplitude artifact was present in an epoch of relatively low voltage background EEG. The power spectrum of the epoch resulted in a high intensity peak in the Delta Band. However, this peak was larger than the one corresponding to the dominant alpha activity, despite the fact that the time duration of the artifact was much shorter than the total time occupied by the alpha activity.

Furthermore, it must be clear by now that the peaks we see in the Delta Band (either in the CSA plots or in the plots of the percentage of power in this band vs time) are not necessarily due to the presence of delta activity as the visual scorer sees it.

The above comments on the "weakness" of spectral analysis as compared to human EEG scoring should not imply that spectral analysis as an EEG quantification tool should be abandoned. On the contrary, we believe it provides a sensitive and reliable tool for a gross examination of noise-like nocturnal EEG activity in artifact-free data. A good example of this are the plots for the percentage of power in the Delta Band vs time, which shows a time profile of the low frequency EEG activity during a night's sleep. These profiles illustrate clearly the cyclic nature of the "delta activity" which occurs throughout the night. Due to the effects of the drugs this "delta rhythm" usually was not detected by visual scoring, since some of the delta activity did not reach the required 75_µV amplitude threshold to be scored as EEG activity belonging to stages 3 or 4, and the Electroencephalographer scored the EEG as stage 2. An illustration of this was already presented in section 2.2.4. It referred to the 5th night of subject SD1, in which the results of the manual scoring showed no sleep stages 3 or 4 present. However, the plot for the percentage of power in the Delta Band vs time showed a large peak at the beginning of the night. A further illustration of the usefullness of these low frequency profiles can be seen in the four nights of subject SD5 (1st, 3rd, 5th and 13th nights). In none of these nights sleep stages 3 or 4 were ever scored. However, the plots for the percentage of power in the Delta Band, Figures 4.22a, b, c and d show cyclic variations in the Delta Band throughout the night. Two subjects in Group 2, SC4 and SC5, have also no sleep stages 3 or 4 scored during their first nights, but the plots for the percentage of power in the Delta Band show peaks in The above examples suggest that even under the this band. effects of large doses of the drugs, the subjects may generate sufficient activity in the delta range to qualify for sleep stages 3 or 4. We observed in some of the corresponding EEG records that this activity indeed had the same period of the delta waves, but its amplitude did not reach the 75 µV threshold required by the visual scoring system for acceptance as

delta activity. Similar observations were also found in some of the score sheets that we looked at, in which the scorer stated that, although there was "delta activity" present, its amplitude was not high enough to be called delta, and so the EEG epochs were scored as stage 2 sleep. The times of the night when these observations were made correspond, in our case, to the times of occurrence of the peaks in the plots of the percentage of power in the Delta Band vs time.

By looking at the plots of the percentage of power in the Delta Band, it can be seen that there is a tendency for a shortening of the period of the "delta rhythm" during the nights as the drug was being withdrawn, for most subjects in Group 1 (SD1, SD2, SD3, SD5 and SD6), or in the 2nd and in the baseline nights, when compared to the first night, for subjects in Group 2 (SC1, SC2, SC3 and SC4). Furthermore, it seems that these peaks tend to move towards the early parts of the night as the drug was withdrawn. Lubin et al (1973) have tried to model this "delta rhythm" (also obtained by spectral analysis) as a damped sinusoid function. However, we did not find any references in the literature with respect to changes in this "rhythm" caused by barbiturates or methaqualone.

The mean percentage of power in the Sigma Band increased in all subjects with the administration of the drugs. It seems that the amount of changes in this parameter is not much related to the dosage the subjects received, and that the two drugs produced similar effects. It has been reported

that barbiturates cause an increase in sleep spindle activity (Allnutt and O'Connor, 1971; Kay et al (1972); Feinberg et al The drugs also caused a decrease in the mean fre-1974). quency in the Sigma Band for all subjects in Group 1. Only two subjects in Group 2, SC3 and SC4, had changes in this parameter that were comparable to the ones exibited by the subjects in Group 1. This slowing of the sigma frequency may be related to the observation by Kayet al (1972), in which they reported that the incidence of sleep spindle bursts increased after pentobarbital and that the frequency within the bursts tended to decrease. However, no reference was made as to the amount of this reduction. The lack of information with respect to changes in frequency of the various EEG waveforms can be explained by the fact that the Electroencephalographers do not look for these kind of changes when they score sleep EEGs into various stages.

The mean percentage of power in the Betal, Beta2 and Fastf Bands increased in all subjects with the administration of either drug, and it seems that there is no apparent drugrelated type of changes. No appreciable changes can be observed in the mean frequencies in these bands. Lester et al (1968) also observed an increase in beta activity after the administration of 200 mg of secobarbital, however no quantification of this increase was presented.

The mean percentage of power in the Theta Band decreased with the administration of the drugs in all subjects. However, these results, in some subjects, could have been artifactual due to the slowing down of the frequency of their alpha activity. This system artifact could be also responsible for the high variability among the subjects in the results for the mean percentage of power in the Alpha Band.

An efficient summary of the effects of the drugs on the activity in the various frequency bands was shown in the plots for the mean frequency coefficients (MFC) for each These plots illustrate in a quantitative fashion night. how the dominant EEG frequencies were shifted during the In all subjects the mean frequency during the night nights. (computed as the mean of the MFCs for an entire night) increases with the administration of the drugs, and then it starts decreasing to its baseline values as the drug is withdrawn. Furthermore, in most of the subjects, this value also seems to be related to the amount of drug administered. In six of the subjects, SD1, SD3, SD4, SD5, SC1 and SC2 we observed that although the mean frequency for each night was decreasing with withdrawal from the drug,, the standard deviation stayed pratically constant from night to night. This suggests that the basic mechanism responsible for the EEG frequency shifts during the night was not seriously affected by the drugs, and that the changes in the mean frequency from night to night were probably due to changes in the amount of slow (Delta, Theta) and fast (Sigma, Betas) wave activities.

The mean value and the percentage of rejections for the x^2 coefficient decreased in all subjects with the administration of either drug. What this suggests is that the EEG amplitude distribution had become more Gaussian due to the effects of the drugs. Furthermore, it seems that the degree of Gaussianity is related to the titration dosage given to the subjects. However, this relation is more clearly seen in the chronic users (Group 1). Furthermore, this drug-related effect is more pronounced in the subjects of both groups that were titrated with the highest doses of either drug. Elul (1969) suggested that, on the basis of the central limit theorem, increased Gaussianity in observed EEG activity may reflect an incresed degree of independence among individual cortical neural generators. A similar increase was also observed by McEwen et al (1975) in subjects under halothane anesthesia. To date, we have not found any other studies that report the effects of abusive or therapeutic dosage of sedative-hypnotic agents on the Gaussianity of the EEG amplitude distribution.

From the above discussion it can be seen that the proposed system seems to be quite efficient in showing drug related patterns in the sleep EEG that otherwise cannot be seen by using the classical approach of sleep staging. Karacan et al (1975) reported that conventional sleep staging parameters obtained from the manual scoring of the sleep EEG of subjects in Group 1, did not exhibit clearly dose-related effects. However, these effects could be better seen in the

EEG waveform characteristics, mainly with respect to spindle activity, which was virtually continuous throughout stage 2 in the titration night and in some nights following it.

The development of an automatic artifact detection algorithm made possible, not only the computation of more reliable power spectra, but also the processing of an entire night of EEG sleep automatically. With the implemented system, about 45 minutes are required to process an entire night of EEG sleep (based on an average of eight hours of EEG data), which is much faster than the two or three hours required by an experienced Electroencephalographer to score the same night. This time can be further reduced if in future applications the sampling rate is reduced to 64 samples/sec instead of the 128 samples/sec used in this study. This reduction in the sampling rate will result in a 50% reduction in the number of data points to be processed. The upper frequency in the analog bandpass filter (Section 2.3.1) can also be reduced to 30 Hz, since no appreciable activity was observed between 30 and 40 Hz.

5.2 RECOMMENDATIONS FOR FURTHER STUDY

In order to better understand the application of the Mean Frequency Coefficients and that of the X^2 coefficients for drug action evaluation, results from normative data are necessary. We suggest the application of the present system to process the sleep EEGs of normal individuals to obtain values for these coefficients that will permit further comparisons with the ones computed in the present work.

Further work is also necessary in order to correlate the values of the X² coefficients with the different EEG frequencies, and to investigate how the Gaussianity of the sleep EEGs is associated with slow wave sleep (SWS) and/or non-SWS. Weiss (1973) found in a normal subject that low frequency activity was correlated in time with non-Gaussian properties of the EEG amplitude distribution.

The quantification of the changes in the "delta rhythm" as a result of drug withdrawal with respect to changes in the amplitude and in the periodicity of the peaks is necessary.

Application of other automated techniques, such as the one proposed by Smith and Karacan (1973), are important to investigate the low frequency activity in order to see if the decrease we observed in the intensity of the spectral peaks, due to the effects of the drug, were the result of a reduction in the amplitude or in the amount of delta waves, or both.

More subjects and more nights per subject are needed if any statistical inference is to be made from the results.

BIBLIOGRAPHY

- Aird, T. J. and Kainer, D. G., "Fourier transform of a real vector of even length-FFTR", IMSL Library, 5th ed., 1975.
- Allnutt, M. F. and O'Connor, P. J., "Comparison of the electroencephalographic behavioral and subjective correlates of natural and drug-induced sleep at atypical hours," Aerosp Med, 42:1006-10, Sept. 1971.
- Aserinsky, E. and Kleitman, N., "Regularly occuring periods of eye motility, and concomitant phenomena, during sleep," Science, 118:273-274, 1953.
- Aserinsky, E. and Kleitman, N., "Two types of ocular motility occuring in sleep," J. Appl. Physiol., 8:1-10, 1955.
- Bendat, J. S. and Piersol, A. G., <u>Measurement and analysis of</u> random data, Wiley, New York, 1966.
- Bendat, J. S. and Piersol, A. G., <u>Random Data: Analysis and</u> <u>Measurement Procedures</u>, New York: Wiley, 1971.
- Beauchamp, K. G., Signal Processing Using Analog and Digital Techniques, New York: Wiley, 1973.
- Bergland, G. D., "A guided tour of the fast Fourier transform," IEEE Spectrum, Vol. 6, pp. 41-52, July 1969.
- Bickford, R. G., Billinger, T. W., Fleming, N. I. and Stewart, L., "The compressed spectral array (CSA) - a pictorial EEG," Froc. 1972 San Diego Biomed. Symp., 365-370.
- Bingham, C., Godfrey, M. D. and Tukey, J. W., "Modern Techniques of Power Spectrum Estimation," Trans. IEEE Audio and Electroacoustics, AU-15, June 1967.
- Blackman, R. B. and Tukey, J. W., The measurement of power spectra. Dover Publications, New York, 1958.
- Brenner, N., "Cooley-Tukey fast Fourier transform FOURL," IBM-Contributed Program Library, 1968.
- Brigham, E. O., <u>The fast Fourier transform</u>, New Jersey, Prentice-Hall, 1974.
- Brigham, E. O. and Morrow, R. E., "The fast Fourier transform," IEEE Spectrum, Dec. 1967, 64-70.
- Bronzino, J. D., Brusseau, J. N., Soern, J. C. and Morgane, P. J., "Power density spectra of cortical EEG of the cat in sleep and waking." Electroenph. Clin. Neurophysiol., 35:187-191, 1973.

- Burch, N. R. and Childers, H., "Physiological data acquisition," In: Psychophysiological Aspect of Space Flight, 1:;95-211. Columbia Univ. Press, New York, 1961.
- Burch, N. R., Nettleton, W. J., Sweeney, J., and Edwards, R. J., "Period analysis of the electroencephalogram on a generalpurpose digital computer," Ann. N. Y. Acad. Sci., 115:827-843, 1964.

Caille, E. J., "Psychotropic drug-induced EEG changes based on power spectrum analysis," In: Turan Itil (ed.), Psychotropic durgs and the human LEG mod. probl. pharmacopsychiat., Vol. 8, pp. 99-116, New York, (Karger, Basel), 1974.

- Cooley, J. W. and Tukey, J. W., "An Algorithm for the machine calculation of Fourier series," Math. Comput., Vol. 19, pp. 297-301, 1965.
- Cox, J. R., Nolle, F. M. and Arthur, R. M., "Digital analysis of the electroencephalogram, the blood pressure wave, and the electrocardiogram," Proc. IEEE, Vol. 60, pp. 1137-1200, Oct. 1972.
- Dean, J. L., "Optimization Techniques for FORTRAN IV (G and H) Programs Written for the IBM 360 under OS," Goddard Space Flight Center, Greenbelt, Maryland, Report NASA-TM-X-70477, 1971.
- Dummermuth, G., Huber, P. J., Kleiner, B., and Gasser, T., "Numerical Analysis of electroencephalographic data," IEEE Trans. Audio Electroacoustics, AU-18:404-411, 1970.
- Dummermuth, G., Walz, W., Scollo-Lavizzari, G., Kleiner, B., "Spectral analysis of EEG activity in different sleep stages in normal adults," Europ. Neurol. 7:265-296, 1972.
- Elul, R., "Gaussian behavior of the electroencephalogram changes during performance of mental task," Science, 164, pp. 328-331, 1969.
- Feinberg, I., Hibi, S., Cavners, C. and March, J., "Absence of REM rebound after barbiturate withdrawal," Science, 185:534-535, 1974.
- Fink, M., Itil, T. M., and Shapiro, D., "Digital Computer Analysis of the human EEG in psychiatric research," Comphrehens. Psychiat., 8:521-538, 1967.
- Fraser, H. F., Wikler, A., Essig, C. F. and Isbell, H., "Degree of physical dependence induced by secobarbital or pentobarbital," J. Am., Med. Ass., pp. 126-129, Jan. 1958.

- Gevins, A. S., Yeager, C. L., Diamond, S. L., Spire, J. P., Zeitlin, G. M. and Gevins, A. H., "Automated analysis of the electrical activity of the human brain (EEG): a progress report," Proc. IEEE, Vol. 63, pp. 1382-1399, Oct., 1975.
- Goldstein, L., Graedon, J., Willard, D., Goldstein, F., Smith, R. R., "A comparative study of the effects of methaqualone and glutethimide on sleep in male chronic insomniacs," J. Clin. Pharmacol. 110:258-268, 1970.
- Grass, A. M. and Gibbs, F. A., "A Fourier transform of the electroencephalogram," J. Neurophysiol, 1, 521-526, 1938.
- Hagne, I., Persson, J., Magnusson, R., Petersen, J., "Spectral analysis via Fast Fourier Transform of waking EEG in normal infants," In: P. Kellaway and I. Petersen (eds.) <u>Auto-</u> <u>matation of Clinical Electroencephalography</u>, New York, Raven Press, 1973.
- Hord, D. J., Johnson, L. C., Lubin, A. and Austin, T. M., "Resolution and stability in the autospectra of EEG," Electroenceph. Clin. Neurophysiol., 1965, 19: 305-308.
- Itil, T. M., Shapiro, D. and Fink, M., "Differentiation of psychotropic drugs by quantitative EEG analysis," Agressologie, Vol. 9, pp. 267-280, 1968.
- Itil, T. M., Saletu, B. and Akpinar, "Classification of psychotropic drugs based on digital computer sleep prints," In: Turan M. Itil (ed.), Psychotropic drugs and the human EEG mod. probl. phamacopsychiat. Vol. 8, pp. 117-130, New York, Karger, Easel), 1974.
- Jenkins, G. M. and Watts, D. G., <u>Spectral analysis and its</u> applications, San Francisco, Calif.: Holden-Day, 1968.
- Johnson, L. C., Lubin, A., Nute, C., and Austin, M. "Spectral analysis of the EEG of dominant and nondominant alphasubjects during waking and sleeping," Electroenceph. Clin. Neurophysiol., 26:361-370, 1969.
- Jones, R. H., "A reappraisal of the periodogram in spectral analysis," Technometrics 7:531-542, 1965.
- Joy, R. M., Hance, A. J., Killam, K. F., "Spectral Analysis of long EEG samples for comparative purposes," Neurophamacology, 1971, 10:471-481.
- Joy, R. M., Hance, A. J., and Killam, K. F., "A quantitative electroencephalographic comparison of some benzodiazepines in the primate," Neuropharmacology, 10:483-497, 1971.

- Kales, A., Kales, J. D., Scharf, M. B., Tan, T-L,I. "Hypnotics and altered sleep-dream patterns, II. All-night EEG studies of chloral hydrate flurazepam and methaqualone," Ach. Gen. Psychiatry, 23:219-225, 1970.
- Kales, A. and Bixler, E., "Sleep profiles of insomnia and hypnotic drug effectiveness," In: N. Burch and H. Altshuler (eds.), <u>Behavior and Brain Electrical Activity</u>, New York, Plenum Press, 1975.
- Karacan, I. Williams, R. L., Finley, W. W., Hurch, C. J., "The effects of naps on nocturnal sleep: Influence on the need for stage-1 REM and stage-4 sleep," Biol. Psychiat., 2:391-399 (1970).
- Karacan, I., O'Brien, G. S., Williams, R. L., Salis, P. J., and Thornby, J. I., "Methodology for Electroencephalographyc sleep evaluation of drugs," In: W. P. Koella and P. Levin (eds.), Sleep, New York, S Krager, p. 463, 1973.
- Karacan, I., Okawa, M., Salis, P. J., Williams, R. L., Comstock, E. G. and Comstock, B. S., "Hypnotic drug abusers: sleep during titration and withdrawal," Sleep Research, 4/103, 1975.
- Kay, D. C. Jasinski, D. R., Eiesenstein, R. B., Kelly, O. A., "Quantified human sleep after pentobarbital," Clin. Pharmacol. Ther., 13:221-231, 1972.
- Kay, D. C., "Sleep and some psychoactive drugs," Psychosomatics, 14:108-118, 1973.
- Kay, D. C., "Human sleep and EEG through a cycle of methalone dependence," Electroenceph., Clin. Neurophysiol., 38: 35-43, 1975.
- Kay, D. C., "Human pharmacology of sleep," In: R. L. Williams and I. Karacan (eds.) <u>Pharmacology of Sleep</u>, New York, Wiley, 1976 (in press).
- Killam, K. F., and Killam. E. K., "Electroencephalographic studies of withdrawal from addition in the papio papio," In: N. Burch and H. Altscnuler, (eds.) <u>Behavior and Brain</u> <u>Electrical Activity</u>, New York, Plenum Press, 1975.
- Knott, J. R., Gibbs, F. A. and Henry C. E., "Fourier transforms of the electroencephalogram during sleep," J. Exp. Psychol. 31:465-477, 1942.
- Larsen, L. E. and Walter, D. O., "On automatic methods of sleep staging by spectra of electroencephalograms," Agressologie, vol. 10, pp.1-13, 1969.

- Larsen, L. E. and Walter, D. O., "On automatic methods of sleep staging by EEG spectra," Electroenceph. Clin. Neurophysiol., 28:459-467, 1970.
- Larsen, L. E., "Sequential methods for parametric discriminant analysis of EEG during sleep," In: N. Burch and H. Altshuler (eds.), <u>Behavior and Brain Electrical</u> Activity, New York, Plenum Press, 1975.
- Lester B. K., Coulter, J. D., Cowden, L. C., Williams, H. L., "Secobarbital and nocturnal physiological patterns," Psychopharmacologia (Berlin), 13:275-286, 1968.
- Lester, L., "A new method for the determination of the effectiveness of sleep-inducing agents in humans," Compr. Psychiatry, 1:301-307, 1960.
- Lubin, A., Johnson, L. C. and Austin, M. T., "Discrimination among states of consciousness using EEG spectra," Psychophysiology, 1969, 6:122-132.
- Lubin, A., Nute, C. and Naitoh, P., "EEG delta activity during human sleep as a damped ultradian rhythm," Psychophysiology, Vol, 10. pp. 27-35, Jan. 1973.
- Mann, H. B., Wald, A., "On the choice of the number of intervals in the application of the chi-square test," Annals of Mathematical Statistics, Sept. 1942.
- Martin, W. B., Johnson, L. C., Viglione, S. S., Naitoh, P., Joseph, R. D., and Moses, J. D., "Pattern recognition of EEG-EOG as a technique for all-night sleep state scoring Electroenceph." Clin. Neurophysio., 32:417-427, 1972.
- Maynert, E. W., "Sedatives and hypnotics II. Bartiturates," In: J. R. DiPalma (ed.), Drill's Fharmacology in Medicine 4th edition, New York, McGraw-Hill, 1971, pp. 250-274.
- McEwen, J. A., Anderson, G. B., "Modeling the stationarity and Gaussianity of spontaneous electroencephalographic activity," IEEE Trans. Bio-Med. Eng., vol.BME-22, pp.361-369, Sept. 1975.
- Meyer, S. L., <u>Data Analysis for Scientists and Engineers</u>, New York; Wiley, 1975.
- Otnes, K. R., and Enochson, L., <u>Digital Time Series</u>, New York, Wiley, 1972.
- Papp, N., "Studio dei filtri numerics d'onda e attuazione di alcune strutture," Tesi di Laurea in Ingegneria Elettronica, Universita di Firenze, 1975.

Parzen, E., "An approach to empirical time series analysis." Radio Sci., 680:937-951, 1964.

- Rechtschaffen, A. and Kales, A., (Eds.), <u>A Manual of Standardized Terminology</u>, Techniques, and Scoring System for <u>Sleep Stages of Human Subjects</u>, National Institutes of Health, Fublication No. 204, U.S. Government Printing Office, Wasnington, 1968.
- Rechtschaffen, A., Robison, T. M., Winsor, M. Z., "The Effect of Methaqualone on nocturnal sleep," Fsychophysiology, 7:346, 1970.
- Richards, F. I., "Computing reliable power spectra," IEEE Spectrum, Jan. 1967.
- Risberg, A. M., Risberg, J., Elmquist, D., Ingvar, D. H., "Effects of dixyrazine and methaqualone on the sleep pattern in normal man," Eur J. Chin, Pharmacol., 8:227-231, 1975.
- Saito, M., "Effects of Psychotropic Drugs on the Human EEG based on Analog Frequency Analysis," In: Turan Itil (Ed.), Psychotropic drugs and the human EEG Mod. probl. pharmaco. psychiat., Vol. 8, pp.117-130, New York, (Karger, Basel), 1074.
- Saltzberg, B. and Burch, N. R., "A rapidly convergent orthogonal representation for EEG time series and related methods of automatic analysis," IRE Wescon Conv., Rec., 8:35-43, 1959.
- Schallek, W., Lewinson, T. and Thomas, J., "Power Spectrum analysis as a tool for statistical evaluation of drug effects on electrical activity of brain," Int. J. Neuropharmacol., 7:35-46, 1968.
- Schneider, S. Y. K., Correlation of the conventional Electroencephalogram with the compressed spectral array electroencephalogram during halothane anestesia in the dog. (M. S. Thesis, University of Houston, 1974).
- Sharpless, S. K., "Hypnotic and sedatives: I. Barbiturates II. Miscellaneous agents,' In: L. S. Goodman and A. Gilman (eds.) The Pharmacological Basis of Therapeutics, 4th edition, New York, AcMillam Company, 1970, pp. 98-134.
- Simpson, R. S., Houts, R. C., <u>Fundamentals of Analog and</u> <u>Digital Communication Systems</u>, Boston: Allyn and Bacon, 1971.
- Singleton, R. C. and Poulter, T. C., "Spectral analysis of the call of the male killer wale," IEEE Trans. Audio and Electroacoustics, AU-15, June 1967.

- Singleton, R. C., "On computing the Fast Fourier transform," Comm. ACM, 10(10), pp.647-654, 1967.
- Smith, J. R. and Karacan, I., "EEG sleep stage scoring by an automatic hybrid system," Electroenceph. Clin. Neurophysiol., 31:231-237, 1971.
- Smith, J. R. and Karacan, I., "Quantification of the effects of a hypnotic-like drug on slow-wave sleep," In: W. P. Koella and P. Levin (eds.) <u>Sleep</u>, New York: S. Kager, p. 504, 1973.

Smith, J. R., Personal Communication, 1976.

- Steiglitz, K., "An introduction to discrete systems," New York: Wiley, 1974.
- Swinyard, E. A., "Sedatives and Hypnotics," In: A Hoover (ed.), <u>Remington's Pharmaceutical Sciences</u>, 15th edition, Pennsylvania, Mac Publishing Co., 1975, p. 1006.
- Tukey, J. W., "An introduction to the calculation of numerical spectrum analysis," In: Spectral Analysis of Time Series Bernard Harris, ed., New York: Wiley, 1967, pp. 25-46.
- Walter, W. G., "An automatic low frequency analyser," Electron Eng., 16:9-13, 1943.
- Walter, D. O., "Spectral analysis for electroencephalogram: mathematical determination of neurophysiological relationships from records of limited duration," Exptl. Neurol., 8:151-181, 1963.
- Walter, D. O. and Adey, W. R., "Analysis of brainwave generators as multiple statistical time series," IEEE Trans. biomed. Eng., 12:8-13, 1965.
- Weiss, M. S., "Non Gaussian properties of the EEG during sleep," Electroencepha. Clin. Neurophysiol., 34:200-202, 1973.
- Welch, P., "The use of fast Fourier transform for the estimation of power spectra: a method based on time averaging over short, modified periodograms," IEEE Trans. Audio and Electroacoustics, Vol. AU-15, No. 2, June 1967.
- Williams, C. A., Jr., "On the choice of the number and width of classes for chi-square test of goodness-of-fit," J. Am. Statistical Assoc., 45, March, 1950, pp. 77-86.
- Williams, R. L., Agnew, H. W. and Webb, W. B., "Sleep patterns in young adults: an EEG study," Electroenceph. Clin. Neurophysiol., 1964, 17:376-381.
- Williams, R. L., Agnew, H. W., "The effects of drugs on the EEG sleep patterns of normal humans," Exp. Med. Surg., 27:53-64, 1969.
- Williams, R. L., Karacan, I., Hursch, C., <u>Electroencephalo-</u> graphy (EEG) of Human Sleep: Clinical Applications, New York: Wiley, 1974.

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APPENDIX A

CHI-SQUARE GOODNESS-OF-FIT TEST

A Chi-square goodness-of-fit test is commonly employed to test whether or not a sample was drawn from a population having a specified distribution. In order to carry out this test, we first set up a null hypothesis stating that the sample was drawn from a set with a known distribution, in our case the normal distribution. Next we compute the chi-square coefficient according to:

$$x^{2} = \sum_{i=1}^{k} (f_{i} - Np_{i}) / Np_{i} = \sum_{i=1}^{k} (f_{i} - F_{i})^{2} / F_{i}$$
(A.1)

where f_i is the actual or observed number of frequencies in the ith class, p_i the probability under the null hypothesis that an observation will fall into the ith class, $F_i = Np_i$ is the expected frequency in the ith class, N the number of observation in the sample, and k is the number of classes. It can be shown that as the size of the sample approaches infinity the distribution of this statistic approaches the chi-square distribution with n=k-l-s degrees of freedom where s is the number of parameters estimated from the sample. In our case s=2 since two parameters (mean and variance) must be estimated to fit a normal distribution function. Hence the number of degrees of freedom for X^2 in Equation (A.1) is n=k-3.

In practice the chi-square distribution is assumed to hold for finite values of N, and the region of acceptance for the test is

$$x^2 \leq \chi^2_{n;\alpha}$$

where the value of χ^2 is available from a table of the chisquare distribution and α is the level of significance. If the sample value χ^2 is greater than $\chi^2_{n;\alpha}$, the hypothesis that the sample distribution is Gaussian is rejected at the α level of significance, otherwise the null hypothesis is accepted at the α level of significance.

The number of classes was computed according to the formula:

$$K = 4 \frac{5}{2(N-1)^2/C^2}$$
(A.2)

where K is the number of classes, N the number of observations in the sample, and c is obtained from a table of areas under the normal (Gaussian) curve such that (Mann and Wald, 1942)

$$(1/\sqrt{2\pi}) \int_{c}^{\infty} e^{-y^{2}/2} dy = \alpha$$
 (A.3)

For $\alpha = 0.05$ (5% level of significance) it follows from Eq. (A.3) that c = 1.64.

In our case N = 3840 and the test is to be conducted at a 5% level of significance. The value of k calculated from Eq. (A.2) is 102.3. Williams (1950) has shown that for large values of N (N \geq 1000) the number of classes can be cut in half with a relatively small effect on the power of the test. In the present study we use k = 52 and it follows that the number of degrees of freedom will be n=49. The acceptance region will be given by:

$$x^2 \leq \chi^2_{49;0.05} = 66.2$$

A FORTRAN subroutine called GOODFT was written to perform the chi-square goodness-to-fit test.

This subroutine was extensively tested both with normally distributed random numbers generated by the digital computer and with digitized Gaussian white noise with a bandwidth of 15 Hz from a noise generator (Hewlett Packard 3722A with infinite sequence length). To generate the normal distributed random numbers we use two IBM Library subroutines: GAUSS and RANDU. Subroutine GAUSS uses 12 uniform distributed random numbers to compute normal random numbers by the central limit theorem. The subroutine RANDU produces the uniform distributed random numbers to be used by GAUSS.

Table A.1 below shows the result of the X^2 test for 10 sets of 3840 normally distributed random number generated by GAUSS. As we can see the null hypothesis was accepted for all sets.

<u>Set</u>	<u>_x²</u>	Null Hypothesis
l	38.4	А
2	42.6	А
3	56.6	А
4	65.1	А
5	40.8	A

TABLE A.1

Set	<u>x²</u>	Null Hypothesis
6	48.5	A
7	47.5	A
8	31.7	A
9	37.9	А
10	54.2	A

TABLE A.1 (continued)

Table A.2 shows the results of the application of the χ^2 test to digitized bandlimited white noise generated by a Hewlett Packard Model 3722A noise generator. The A/D conversion procedure is the same described in Chapter II. The Gaussian noise was lowpass filtered with cutoff frequency of 15 Hz and digitized at a rate of 128 samples per second. Each 30 seconds of data was stored as an epoch. In only one epoch was the null hypothesis rejected.

Epoch	<u>_x²</u>	Null Hypothesis
1	53.4	А
2	48.5	A
3	42.5	А
4	53.7	А
5	45.2	А
6	74.7	R
7	55.5	А
8	35.9	А
9	54.4	А
10	40.2	A

In both examples the test's acceptance region is

$$x^2 \le 66.2$$

The execution time for subroutine GOODFT was minimized by using optimization techniques suggested by Dean (1971) and by compiling the subroutine under FORTRAN IV H-Level. For 3840 data points the total execution time was 1.4 seconds. A detailed listing of GOODFT appears in Appendix B. APPENDIX B

COMPUTER PROGRAMS

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1

2:		***************************************	MAIN MAIN MAIN	10 20 30
4; 5:	č	* PRUGRAM CUNVERSIUN *	MAIN	40 50
6: 7:	C C	************************	MAIN	60
8	Ŭ	DATA POT1, POT2, POT3/ PO01', PO02', PO03'/	MAIN	ģŏ
ð:		INTEGER*2 LOCAD1(3841),LOCAD2(3841),DUMNY	MAIN	100
1:		REAL#8 WTMCCW INTEGER#2 IVAL/1000/	MAIN	110
3:		DATA POTI, PUTZ, POT3/ PO01 ', 'PO02', 'PO03'/	MAIN	įžŏ
5:		LOGICAL XRUSNS	MAIN	150
6: 7:		IT=30 N=128*IT	MAIN	$160 \\ 170$
8:	105	WRITE(15,105)	MAIN	180
3 :	103	CALL TYPE(0)	MAIN	200
$\frac{1}{2}$:	108	WRITE(15,108) FORMAT('TURN LOGIC PUSHBUTTON # 1 - ON ALL OTHERS OFF')	MAIN MAIN	210
3:		CALL TYPE(0)	MAIN	230
5:	106	FORMAT(' SET ANALOG IN RMT MODE AND DEPRESS EOB')	MAIN	250
6: 7:		CALL XPOTSS(POT1,IVAL) CALL XPOTSS(POT2,IVAL)	MAIN	260
8		CALL XPOTSS(POT3,IVAL)	MAIN	280
0	10		MAIN	300
2:		$\frac{1}{10R} = 4$	MAIN	310
3:		N2N = N + 2 $CALL CCWCB (TPCCW) (CB, OP, LOCAD) (2) (N2N)$	MAIN	330
5		CALL CCWCB (TPCCW2,CB,OP,LOCAD2(2),N2N)	MAIN	350
6: 7:		CALL READAD (CCWADI)N,2;LUCADI) CALL READAD (CCWAD2;N,2;LUCADI)	MAIN	360
8:		CALL CCHCB (WTMCCW,WTH,OP,DUMMY,N)	MAIN	380
ó:		LOCAD1(1) = 0*256 + NCH - 1	MAIN	400
2:	2000	CONTINUE	MAIN	410
3: 4:		18 = 0 1FT1F = IFT1F + 1	MAIN MAIN	430
5		CALL XRSCTL (IHOLD,0)	MAIN	450
7:	203	CONTINUE	MAIN	400
8:		IF(.NOT.XRDSNS(1)) GO TO 111 CALL XSTCTL([HOLD.])	MAIN MAIN	480
0	С	DELAY TO HOLD LIGHT ON	MAIN	500
2:	20	CONTINUE	MAIN	520
3: 4:		CALL XRSCTL(IHOLD,1) DO 30 [=1.10000	MAIN	530 540
5:	30		MAIN	550
7:	111	CALL XIC	MAIN	570
8: 9:		CALL XSICILIIHULD,3J CALL FSTIO(0,CCWAD1,0)	MAIN	580
0:	1000		MAIN MAIN	600
2		CALL FSILU(0,CCWAD2,0)	MAIN	620
3: 4:		CALL FSTOLDR, IPCCWI, I) CALL FSTCHK(0)	MAIN	640
5: 6:		CALL FSTID(0,CCWAD1,0) CALL FSTID(108,TPCCW2,1)	MAIN	650
7:		IF (.NOT.XRDSNS(1)) GO TO 1000	MAIN	670
8:		CALL FSTCHK(0)	MAIN	680
9: 0:		CALL FSITU (IUR, WIMCLW, I) CALL XRSCTL(IHOLD, 3)	MAIN MAIN	690 700
1:	159	WRITE (6,157) IFILE, IB FORMAT (* EILE NUM = 1.13.1 NO. OF BLOCKS = 1.15	MAIN	710
3	~~/ 777	IF(XRDSNS(4)) GO TO 2000	MAIN	730
5:		CALL_TYPE(0)	MAIN	740 750
6: 7:	115	WRITE(15,110) FORMAT(10FPRESS LOAD/REWIND ON TAPE DRIVE 281 - THEN FORM	MAIN	760
8:		STOP 777	MAIN	780

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1:	C	**************************************	MAIN MAIN	10
3. 4. 5.	č	* PROGRAM DTOA *	MAIN	40 50
6	č	**********	MAIN	60 70
8: 9: 10: 11: 12:		INTEGER*2 LOCAD(3860),LOCA(11581) DIMENSION IE(30) DATA POT4,POT5/*POO4*,*PO05*/ INTEGER*2 IVAL/500/ INTEGER EA,SA LOGICAL XROSNS	MAIN MAIN MAIN MAIN MAIN MAIN	80 90 100 110 120
14:	С	ÎNÎÊĞÊR*2 ÎHÔLD/O/	MAIN	140
15:		IT IS THE TIME DURATION OF THE EPOCH IN SECONDS	MAIN	150
16:		CAL=18.0/11.0	MAIN	160
17:		IBLOCK=61	MAIN	170
18:	100	READ(5,100) KE,(IE(J),J=1,KE)	MAIN	180
19:		FORMAT(20I3)	MAIN	190
20:		NN=11580	MAIN	200
21:		IT=30	MAIN	210
22:		N=128*IT	MAIN	220
23:		N1=N+1	MAIN	230
24:		N3=3*N	MAIN	240
25:		N31=N3+1	MAIN	250
26:		EA=2	MAIN	260
27:		SA=0	MAIN	270
28:		ICON=10	MAIN	280
29:		IS=1	MAIN	290
30:		CALL XPOTSS(POT4,IVAL)	MAIN	300
31:		CALL XPOTSS(POT5,IVAL)	MAIN	310
32:		CALL XIC	MAIN	320
33:		CALL XRSCTL(IHOLD,3)	MAIN	330
34:		IREC=0	MAIN	340
35:		ITEST=3	MAIN	350
36:		IC=ISTART	MAIN	360
37:		CALL_FOPEN	MAIN	370
38:	35	DO 777 IB=1,IBLOCK	MAIN	390
39:		CALL RDTAPE(LOCAD,&9999)	MAIN	390
40:		DO 35 J=1,KE	MAIN	400
41:		IF(IB.EQ.IE(J)) GO TO 36	MAIN	410
42:		CONTINUE	MAIN	420
43:	36	ĜO TO 777 CONTINUE	MA IN MA IN	430 440
45:	200	WRITE(6,200) IB	MA IN	450
46:		Format(20x,13//)	MA IN	460
47:	11	IS=IS+1	MAIN	470
48:		I=0	MAIN	480
49:		DO 72 J=IS,N31,3	MAIN	490
50:		I=I+1	MAIN	500
51:		LOCA(J)=LOCAD(I)/3	MAIN	510
53: 53: 55: 56:	C	IREC=IREC+1 TO TEST IF LOCA IS FILLED WITH THE THREE EPOCHS IF(IREC.NE.ITEST) GO TO 777 IS=1 INFC-0	MAIN MAIN MAIN MAIN MAIN	530 530 550 560
578: 599: 61: 62:	74	L=11521 D0 74 J=1,43,3 L0CA(L+J)=819 L0CA(L+J+1)=1638 L0CA(L+J+2)=2457 CONTINUE	MAIN MAIN MAIN MAIN MAIN MAIN	580 590 600 610 620 630
64: 65:	••	L=L+45 DD 75 J=1.15	MAIN	640 650
66:	75	LOCA(L+J)=0	MAIN	660
67:		CONTINUE	Main	670
68: 69: 70: 71: 72:	85 C 20	DO 85 J=2,NN X=LOCA(J)*CAL LOCA(J)=X CONTINUE PRESS PSB # 5 TO START THE PLOTTING JELYPOSNS(5)) CO TO 31	MAIN MAIN MAIN MAIN MAIN	680 690 700 710 720
74: 75: 76: 77:	50	CALL XSTCTL(IHOLD,5) DO 50 J=1,10000 CONTINUE CALL X85CTL(IHOLD,5)		740 750 760
78:	20	DO 39 K=1,10000	MAIN	780
80:	C 31	GO TO 29	MAIN	800
81:		STARIS THE CLOCK WITH 5 MS OF DELAY	MAIN	810
82:		CALL XSTCTL(THOLD+3)	MAIN	820
83:	C	CALL XWRTDĀ(NN,ĪĊÓN,SA,EA,LOCA)	MAIN	830
84:		Resets the clock at the end of the plotting	MAIN	840
85:	C	CALL XRSCTL(IHOLD,3)	MA ÎN	850
86:		TO REPEAT THE PLOTTING PRESS PSHB #2	MA ÎN	860
87:	с	IF(XRDSNS{2}) GO TO 29	MAIN	870
88:		TO STOP THE ENTIRE RUN PRESS PSHB # 0	MAIN	880
89: 90: 91: 92:	777 9999	IF(XRDSNS(0)) GO TO 9999 CONTINUE STOP END	MAIN MAIN MAIN MAIN	890 900 910 920

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2: C 3: C 4: C 5: C	****************** * * * *	*************** PROGRAM ************	**************************************	**************************************
7: C 3: 0: 1: 2: 3: 4: 5: 5: C	INTEGER*2 LOCA(3860 DIMENSION X(3840),X EQUIVALENCE (LOCA(3 EQUIVALENCE (LOCA(3) ITR=30 N3=128*ITR FN3=N3 KINTER=52 CHISQ=66.3),LOCAD(3860) CHIS(500),XX(1) 841),XX(1)) 3841),YY(1))	D),YY(10)	MAIN 7 MAIN 9 MAIN 9 MAIN 10 MAIN 11 MAIN 12 MAIN 13 MAIN 14 MAIN 15 MAIN 16 MAIN 16
	XMAX=XX(1) XMIN=XX(2) XVARX=XX(3) SKEW=XX(4) XKURT=XX(5) XSQR=XX(5) XBARX=XX(7) CSTDX=XX(8)			MAIN 18 MAIN 19 MAIN 20 MAIN 21 MAIN 22 MAIN 23 MAIN 24 MAIN 25 MAIN 25
7: 3: 9: 0: 100	XX(9)=0.0 XX(10)=0.0 READ(5,100) IBLOCK, FORMAT(14/,14/,14/, ARTF=IARTF/81.92	ISTART,ISKIP,I 14/,F10.0/,F10	ARTF,XSQRL1,XSQRL2 0)	MAIN 27 MAIN 28 MAIN 29 MAIN 30 NAIN 31
2: 199 3: 199 5: 6: 7: 3: 9: 0: 1: 2: 3:	WRIFE(6,199) FORMAI(1H1////) ICDUNT=0 IC=ISTART LT=0 MT=0 IFL=0 SUM=0.0 CALL TOPEN CALL TWOPEN CALL TWOPEN DO 71 IB=1,IBLOCK CALL RDTAPE(LOCA.69	9999)		MAIN 32 MAIN 33 MAIN 35 MAIN 35 MAIN 37 MAIN 38 MAIN 39 MAIN 39 MAIN 40 MAIN 42 MAIN 42 MAIN 42
4: 5: 1000 6: 2000 7: 10 8: 9: 9:	IF(MT) 10,1000,10 IF(LT) 10,200C,10 IF(IB-IC) 71,10,71 CONTINUE DO 11 I=1,N3 LH=LOCA(I) X(I)=LOCA(I)/81.92 IF(IA3S(I)+)-IARIE)	11.20.20		MAIN 44 MAIN 45 MAIN 46 MAIN 47 MAIN 48 MAIN 49 MAIN 50 MAIN 51
2 2.0	II=I	11/20/20		MAIN 52 MAIN 53
5: 5: 6: 7:	CONTINUÉ CALL PARMR(X,N3,XX) CALL GOUDFT(X,N3,KI XSQR=XX(6)	NTER,XX(7),XX(8),CHISQ,NHYPT,XX(6))	MAIN 54 - MAIN 55 MAIN 56 MAIN 57
8: 30 9: 30 0: 2: 2: 3: 5: 6: 7:	IF(XSQR-XSQRL1) 30, IC=IC+ISKIP ICOUNT=ICOUNT+1 LT=0 IFL=0 XCHIS(ICOUNT)=XSQR SUM=SUM+NHYPT CALL TWRITE(LOCA) GO TO 71	. 31, 31		MAIN 58 MAIN 59 MAIN 60 MAIN 61 MAIN 62 MAIN 63 MAIN 64 MAIN 65 MAIN 65 MAIN 67

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				PAGE 2
68: 69: 70: 71:	22 210	IC1=IC+LT+MT WRITE(6,210) IC1,X(II) FORMAT(40X, AMPLITUDE EXCEEDED AT EPOCH N 2/)	NO•'•I4•5X•' X='•F7•2	MAIN 680 MAIN 690 2MAIN 700 MAIN 710
72: 73: 74: 75: 76:	21 41	LT=LT+1 ISL=LT+MT IF(ISL-3) 71,71,21 IF(IFL) 40,41,40 LTI=LT-1 LTI=LT-1		MAIN 720 MAIN 730 MAIN 740 MAIN 750 MAIN 760
78: 79: 80: 81:	250	WRITE(6,250) FORMAT(40X, ** A SET OF EPOCHS WAS SKIPPED DU MI=0 LT=0 IC=IC+ISKIP	UE TO ARTIFACTS'/)	MAIN 770 MAIN 780 MAIN 790 MAIN 800 MAIN 810
82: 83: 84: 85: 86:	31 240	GO TO 71 IC2=IC+MT+LT WRITE(6,240) IC2,XSOR FORMAT(40X, ARTIFACT DETECTED AT EPOCH N 2/)	NO.º,I4,5X,ºXSQR=º,F7.2	MAIN 820 MAIN 830 MAIN 840 MAIN 850 MAIN 860
87: 88: 89: 90:	42	MT=MT+1 ISM=MT+LT IF(XSQR-XSQRL2) 42,42,43 IF(IFL) 44,45,44 DO 47 K-1. N2		MAIN 870 MAIN 880 MAIN 890 MAIN 900
92: 93: 94:	47	LOCAD(K)=LOCA(K) CONTINUE DO 60 J=1,8		MAIN 920 MAIN 930 MAIN 940 MAIN 950
96: 97: 98:	60	CONTINUE XSQRF=XSQR IFL=1 IFLSN=3, 71,71,40		MAIN 960 MAIN 970 MAIN 980
100: 101: 102: 103: 104:	44 43 50 40	IF(ISMR-SSORF)45,43,43 IF(ISM-3) 71,71,50 IF(IFL) 40,32,40 IC=IC+ISKIP ICOUNT=ICOUNT+1		MAIN 990 MAIN1000 MAIN1010 MAIN1020 MAIN1030 MAIN1040
105: 106: 107: 108: 109:		LT=0 MT=0 IFL=0 XCHIS(ICOUNT)=XSQRF DD_61_J=1,8		MAIN1050 MAIN1060 MAIN1070 MAIN1080 MAIN1090
110: 111: 112: 113: 114:	61	XX(J)=YY(J) CONTINUE SUM=SUM+NHYPT CALL TWRITE(LOCAD) CO TO 71		MAINI100 MAIN1110 MAIN1120 MAIN1130 MAIN1140
115: 116: 117: 118: 119:	32	MIT=MT-1 WRITE(6,250) MT=0 IFL=0		MAIN1150 MAIN1160 MAIN1170 MAIN1180 MAIN1190
120: 121: 122: 123:	71 9999	IC=IC+ISKIP CONTINUE CONTINUE IF(IFL) 51,52,51		MAIN1200 MAIN1210 MAIN1220 MAIN1230
124: 125: 126: 127:	51	ICDUNT=ICOUNT+1 XCHIS(ICOUNT)=XSQRF SUM=SUM+NHYPT CALL TWRITE(LOCAD)		MAIN1240 MAIN1250 MAIN1260 MAIN1270
128: 129: 130:	52	CONTINUE STOP END		MAIN1280 MAIN1290 MAIN1300

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RDTP RDTAPE CSECT HYBRID TAPE READ ROUTINE[®] 1234567890123456789012345678901 ******* CALL RDTAPE(IBUF,69999) IBUF = INPUT BUFFER AREA (MAY BE OF ANY MODE) THE SIZE OF THE INCOMING RECORD DETERMINES THE DIMENSIONS OF IBUF. I.E. IF YOU HAD AN A/D TAPE WITH 1024 SAMPLES PER TAPE RECORD YOU WOULD DIMENSION IBUF AS FOLLOWS: INTEGER*2 IBUF(2048) 69999 = END OF FILE ADDRESS THE USER MUST CALL TOPEN ONLY ONCE BEFORE MAKING CALLS ON *** ROTAPE. IN ORDER TO SKIP A FILE MARK ON THE TAPE YOU MAY USE THE FOLLOWING: CALL SKFILE. * * CALL SKFILE. THE ONLY CONTROL CARD NECESSARY IS THE FOLLOWING: //GO.TAPE08 DD UNIT=2400,VOL=SER=ADDATA,DISP=(OLD,PASS), // LABEL=(1,NL),DCB=(RECFM=U,BLKSIZE=B192) SPACE 2 ENTRY TOPEN,SKFILE SPACE 4 RDTAPE ENTRY SAVE (14,12),T,RDTAPE BALR 10,0 USING *,10 ST 13,SAVE+4 LA 5,SAVE ST 5,8(13) LR 13,5 LA 8,EOFADDR1 LA 9,TAPEDCB MVC X'21'(3,9),1(8) LM 2,3,0(1) MVI DECDT,X'00' READ DECBT,SF,TAPEDCB,(2),'S',MF=E CHECK DECBT L 13,SAVE+4 L 13,SAVE+4 RETURN (14,12),T SPACE 2 DCEOF EQU * L 13,SAVE+4 LM 14,12,12(13) * * * * * * 3234567 LOCEOF 44445555555555566666 * 13, SAVE+4 14, 12, 12(13) 12(13),X*FF* 15,4 15,14 L LM MV I LA BCR * EJECT ** CALL TOPEN * * EQU SAVE BALR USING ST LA ST **TOP EN** * (14,12),T,TOPEN 10,0 *,10 13,SAVE+4 5,8(13) 13,5 (TAPEDCB,INPUT) 13,SAVE+4 64: 66: **OPEN**

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			PAGE	2
68:	•	RETURN (14,12),T	MAIN	680
70:	*	EJECT	MAIN	202
72: 73: 74:	* *	CALL SKFILE	MAIN MAIN MAIN MAIN	720 730 740
75: 76:	SKFILE	EQU * SAVE (14,12),T,SKFILE	MAIN MAIN	750 760
77:		BALR 10,0 USING *10	MAIN	770
80:		SI 13,5AVE+4 LA 5,5AVE ST E SIJA	MAIN	800
82:		$\begin{array}{cccc} 131 & 310131 \\ LR & 1335 \\ LA & 5 & 5 \\ LA & 5 $	MAIN	820
84:		LA 9, TAPEDCB	MAIN	840
86:	* *	CNTRL TAPEDCB, FSM	MAIN	860
88:	•	LA 1,TAPEDCB	MAIN	880
90:		BALR 14,15 DC 412(63)	MAIN	900
92: 93:		MVI DECBT,X'OO' READ DECBT.SE.TAPEDCB.DUMARA.'S'.ME≓E	MAIN	920 930
94:	CTRIEDE	CHECK DECBT	MAIN	940 950
96: 97:		L 13, SAVE+4 RETURN (14,12), T	MAIN	960 970
98: 99:	*		MAIN MAIN	980 990
100: 101:	*	EJECT	MAIN1 MAIN1	000
102:	*	DATA CONTROL AREA SPACE 2	MAINI	020
104: 105:	SAVE *	DC 18F'O' SAVE AREA	MAIN1 MAIN1	040
106: 107:		READ DECBT, SF, MF=L SPACE 4	PAIN1 PAIN1	060
108:	EOFADDR1 EOFADDR2	DC AL4(LOCEOF) DC AL4(CTRLEOF)	MAIN1 MAIN1	080
110:	DUMARA	DS CL20 SPACE 4	MAINI	100
112:	TAPEDCB	UCB UDNAME=IAPEO8,DEVD=IA,MACRF=(RC),USURG=PS,EOCAD=(LOCEOF) END	MAINE	130

1234567890112345 1112345	TWR I TE	CSECT ENTRY TWRITE ENTRY TWOPEN USING TWRITE,15 SAVE (14,12),T,* LR 14,13 CNOP 0,4 BAL 13,*+76 USING *,13 DS 18F ST 13,8(14) ST 14,4(13) DROP 15 L 2,0(1) WRITE TODECB,SF,,(2),MF=E	MAIN 10 MAIN 200 MAIN 400 MAIN 500 MAIN 700 MAIN 700 MAIN 700 MAIN 700 MAIN 1000 MAIN 1120 MAIN 1200 MAIN 1200 MAIN 1200 MAIN 1200 MAIN 1200 MAIN 1400
1111122223455578991	TWOPEN	CHECK TODECB L 13,4(13) RETURN (14,12),T,RC=0 SAVE (14,12),T,* USING TWOPEN,15 LR 14,13 CNOP 0,4 BAL 13,*+76 USING *,13 DS 18F ST 13,8(14) ST 14,4(13) DROP 15 OPEN (TAPEO,OUTPUT) L 3,4(13) RETURN (14,12),T_RC=0	AAIN 160 MAIN 170 MAIN 1900 MAIN 2000 MAIN 2200 MAIN 2200 MAIN 2200 MAIN 2400 MAIN 2600 MAIN 2700 MAIN 2900 MAIN 300 MAIN 310
32: 33: 34:	TAPEO	DCB DONAME=TAPEOUT,DEVD=DA,DSORG=PS,MACRF=(W) WRITE TODECB,SF,TAPEO,MF=L END	MAIN 320 MAIN 330 MAIN 340

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	ç	SUBROUTINE GOODFT(XDATA,NSIZE,KINTR,XBARX,CSTDX,CHISQ, 1NHYPT,XSQRT)	6000 6000 6000	10 20 30
	າດດດດດ	THIS SUBROUTINE TESTS THE NORMALITY OF THE ARRAY XDATA BY PERFORMING A CHI-SQUARE GJJDNESS OF FIT TEST USING CLASS INTERVALS WITH EQUAL EXPECTED FREQUENCIES	GOOD GOOD GOOD GOOD GOOD	50 60 70 80
5	č	***********	*čočč	100
3	JUUU Q	XDATA REAL ARRAY CONTAINING THE DATA POINTS NSIZE INTEGER VALUE FOR THE SIZE OF THE ARRAY XDATA KINTR INTEGER VALUE FOR THE NUMBER OF CLASS INTERVALS		120 130 140
	CCCC	XBARX REAL VALUE OF THE SAMPLE MEAN OF THE DATA CSTDX REAL VALUE OF THE SAMPLE STANDARD DEVIATION OF THE DATA ZGAUS ARRAY OF THE ABCISSA VALUES FROM THE NORMAL DENS. FUNCTI CHISO REAL VALUE OBTAINED FROM A CHI-SOUARE TABLE		150 160 170
		*XSORT REAL VALUE OF THE SAMPLE STATISTICS CALCULATED BY GOODF *NHYPT IF IT IS 1 THE NORMALITY HYPOTHESIS IS ACCEPTED AT THE ALPHA LEVEL OF SIGNIFICANCE		190 200 210
	CCCC	IF IT IS O THE NORMALITY HIPOTHESIS IS REJECTED AT ALPHA LEVEL OF SIGNIFICANCE	6000 6000 6000	220 230 240
		THIS SUBROUTINE CALLS NOTRI TO GENERATE THE ZGAUS ARRAY	GOUD GOUD GOUD	260 270 280
	Č	**************************************	*6000 6000 6000	290 300 310
	~	FSIZE=NSIZE FINTR=KINTR KK=KINTR-1	6000 6000 6000	320 330 340
);); 7; ;;	с 30	LUMPOTES THE ZGAUS ARRAT IF(IFLAG) 20,30,20 PROBI=1.0/FINTR DD 19 1=1.KK	6000 6000 6000	360 370 380
		P=PRÚBI+Ĵ CALL NDTRI(P,Z,D,IER) ZGAUS(J)=Z	6000 6000 6000	390 400 410
	19 20	CONTINUE IFLAG=1 CONTINUE CONTINUE	600D 600D 600D	420 430 440
	c c	CALCULATE THE EXPECTED FREQUENCY IN EACH INTERVAL EFREQ=FSIZE/FINTR CONST=1.0/EFREQ CALCULATES THE END POINTS OF FACH INTERVAL	6000 6000 6000	450 460 470 480
):	1	DO 1 I=1,KK X(I)=CSTDX+ZGAUS(I)+XBARX CONTINUE	6000 6000 6000	490 500 510
	с 2	TO FIND THE OBSERVED FREQUENCY AT EACH INTERVAL DO 2 I-1,KINTR OFREQ(I)=0.0	6000 6000 6000	520 530 540
	3	10 - 7 - 1, 1312C IF(XDATA(J)-X(1)) 3,3,4 OFREQ(1)=OFREQ(1)+1.0 CD TO 7	6000 6000 6000	550 560 570 580
	4 5	ĬĔ(XŬATA(J)-X(KK)) 6,6,5 OFREQ(KINTR) =OFREQ(KINTR) +1.0 GO TO 7		590 600 610
3:	6	DD 9 I=2,KK IF(XDATA(J)-X(I)) 8,8,9		620 630
	8 9	DFREQ(I)=OFREQ(I)+1.0 G0 TŪ 7 CONTINUE	GOOD GOOD GOOD	640 650 660
1:	7	CONTINUE	GOOD	670

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			PAGE	2
68: 69: 70: 72: 73: 74: 75:	C 10 C	CALCULATE THE TOTAL DISCREPANCY FOR ALL CLASS INTERVALS XSQRT=0.0 DO 10 K=1,KINTR FDIF=EFREQ-OFREQ(K) XSQRT=XSQRT+OFREQ(K)**2 CONTINUE XSQRT=XSQRT*CONST-FSIZE NUMBER_OF DEGREES OF FREEDOM IS GIVEN BY		680 690 710 720 730 740 750
76: 77: 78: 79: 80: 81:	C 69	N=KINIR=3 TO TEST THE HYPOTHESIS OF NORMALITY NHYPI=1 IF(XSQRT.LE.CHISQ) NHYPT=0 RETURN END		760 770 780 790 800 810

1:	_	SUBROUTINE PARMR(X+N3+XX)	PARM	10
2:	č		PARM	20
3:	č	***************************************	PARM	40
5:	č	THIS SUBROUTINE COMPUTES THE FOLLOWING PARAMETERS FOR THE X	PARM	50
6:	ç	ARRAY: MIN AND MAX VALUES, MEAN, VARIANCE, STD, KURTOSIS AND	PARM	60
	č	SKEWNE SS	PARM	80
۶:	č	******	*PARM	90
10:	C		PARM	100
11:			PARM	120
13:			PARM	1 30
14:		EN3=N3	PARM	140
15:				150
17:			PARM	170
18:		ŠŲMX4=0.0	PARM	180
19:			PARM DADM	190
21:			PARM	210
22:		TERM=TERM*X(I)	PARM	220
23:		SUMX2=SUMX2+TERM	PARM	230
25:		IERM=TERM*A(1) SIMX3=SIMX3+TERM	PARM	250
26:		TERM=TERM*X(I)	PARM	260
27:		SUMX4=SUMX4+TERM	PARM	270
28:		XX(1)=AMAX1(XX(1)+X(1)) XY(2)=AMIN1(YX(2)-X(1))	PARM	290
30:	11	CONTINUE	PARM	ΞÓŎ
31:		XX(7)=\$UMX1/EN3	PARM	310
32:		XX (3)=SUMX2/FN3 VY (3)=SUMX2/FN3	DADM	320
34:		2013 - SUNX 3/FN3-3.0*XX (7) + XX (3) + 2.0*XX (7)	PARM	340
35:		XMT4=ŠUMX4/FN3+4.0*XMT3+6.0*XX(3)*XX(7)**2+5.0*XX(7)**4	PARM	350
36:		XX(4)=XMT3/(XX(3)**1;5)		360
38:		∧∧\)/=\ +/\\^\)/** / RFTURN	PARM	380
39:		END	PARM	390

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123456789012345678901234567890123456789012345678901234567890123456789012345678901234567890123456789 61: 62: 63: 64: 65: 66:

10 20 30 40 NDTR NDTR NDTR SUBROUTINE NDTRI NDTR 5ŏ PURPOSE COMPUTES X = P**(-1)(Y), THE ARGUMENT X SUCH THAT Y= P(X) = THE PROBABILITY THAT THE RANDOM VARIABLE U, DISTRIBUTED NORMALLY(0,1), IS LESS THAN OR EQUAL TO X. F(X), THE ORDINATE OF THE NORMAL DENSITY, AT X, IS ALSO COMPUTED. 60 70 80 90 ND TR 100 110 120 130 USAGE CALL NDTRI(P,X,D,IER) 140 150 160 170 180 DESCRIPTION OF PARAMETERS P - INPUT PROBABILITY. X - OUTPUT ARGUMENT SUCH THAT P = Y = THE PROBABILITY THAT U, THE RANDOM VARIABLE, IS LESS THAN OR EQUAL TO X. D - OUTPUT DENSITY, F(X). IER - OUTPUT EROR CODE = -1 IF P IS NOT IN THE INTERVAL (0,1), INCLUSIVE. X=D=.99999E+74 IN THIS CASE = 0 IF THERE IS NO ERROR. SEE REMARKS, BELOW. REMARKS MAXIMUM ERROR IS 0.00045. IF P = 0, X IS SET TO -(10)**74. IF P = 1, X IS SET TO (10)**74. D IS SET TO O. D IS SET TO O. NO TR NDTR NDTR NDTR SUBROUTINES AND SUBPROGRAMS REQUIRED NONE NDTR NDTR NDTR NDTR NDTR NDTR NDTR ME THOD BASED ON APPROXIMATIONS IN C. HASTINGS, APPROXIMATIONS FOR DIGITAL COMPUTERS, PRINCETON UNIV. PRESS, PRINCETON, N.J., 1955. SEE EQUATION 26.2.23, HANDBOOK OF MATHEMATICAL FUNCTIONS, ABRAMOWITZ AND STEGUN, DOVER PUBLICATIONS, INC., NEW YURK. 340 350 360 370 380 390 **4**00 410 420 430 450 SUBROUTINE NDTRI(P,X,D,IE) С IE=0 X=.999999E+74 D=X IF(P)1,4,2 I IE=-1 GO TO 12 2 IF (P-1.0)7,5,1 4 X=-.9999999E+74 5 D=0.0 GO TO 12 460 480 490 NDTR NDTR NDTR 500 510 520 530 NDTR 540 550 560 570 NOTR C C ND TR 7 D=P IF(D-0.5)9,9,8 8 D=1.0-D 9 T2=AL0G(1.0/(D*D)) T=SQRT(T2) X=T-(2.515517+0.802853*T+0.010328*T2)/(1.0+1.432788*T+0.189269*T2 1 +0.001308*T*T2) IF(P-0.5)10,10,11 10 X=-X NDTR ND TR ND TR 580 590 NDTR NDTR NDTR NDTR NDTR 600 610 620 630 X=-X D=0.3989423*EXP(-X*X/2.0) RETURN NDTR 640 NDTR 650 NDTR 660 NDTR 670 10 11 12 END

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CCCC		* MAIN * MAIN * MAIN * MAIN
č	<pre>INTEGER*2 LOCA(3860) DIMENSION X(4096),PSPEC(2058),PNORM(2048),XX(10),TNORM(90),PARM(9 EQUIVALENCE(TNORM(1),PNORM(1)),(XX(1),PARM(1))</pre>	MAIN MAIN MAIN MAIN
	EQUIVALENCE(PSPEC(1),LUCA(1)) ITR=30 ITZ=32	MAIN MAIN MAIN
	N=128+112 NHALF=N/2 NDEL=NHALF/128	MA IN MA IN MA IN
	N3=128+ITR N1=N3	MAIN MAIN MAIN
	N2=2*N FN=N DELTT=1.0/DFREQ	MAIN MAIN MAIN
	TTIME=FN *DELTT DELTF=1.0/TTIME RFAD(5.100) ISMTH	MAIN MAIN MAIN
100	FORMAT([3) ICOUNT=0 KCOMP=82	MAIN
	UP=81 LL=2048	MAIN
	NLAP=0 L=1920	MAIN
200	FORMAT(3x, 'EPOCH', 3x, 'XMAX', 6X, 'XMIN', 3X, 'VARIANCE', 3X, 'SKEWNESS' 13X, 'KURIOSIS', 3X, 'XSOR', 5X, 'MEAN', 4X, 'PSPECMX', 3X, 'PSPECMN'//)	MAIN MAIN MAIN
	CALL TUPEN DD 71 I8=1,400 CALL RDTAPE(LOCA, δ9999)	MAIN MAIN MAIN
20	DO 20 K=1,10 XX(K)=PSPEC(L+K) CONTINUE	MAIN MAIN MAIN
70	ĎĎ 70 K=N3,N X(K)=0,00 CONTINUE	MAIN MAIN MAIN
	DO 72 J=1,N3 X(J)=LOCA(J)/81.92−XX(7) CONTINUE	MAIN MAIN MAIN
	ČALL TAPER1(X,N1,DFREQ) CALL PWRP(X,PSPEC,N,DELTT) LCDUNT=1COUNTA1	MAIN MAIN MAIN
	CALL SMTH15(PSPEC,N,PNDRM,NLAP) J=0	MAIN
10	J=J+1 PNORM(J)=PSPEC(I)	MAIN MAIN
10	DD 29 I=1,ISMTH CALL_SMODTH(PNORM,KCOMP)	MAIN
ĉ	COMPUTES THE MAX AND MIN VALUES OF PNORM	MAIN
	XX(9)=10 DO 187 KF=1,81 XX(8)=AMAX1(XX(8),PNORM(KF))	MAIN MAIN MAIN
187	XX(9)=AMIN1(XX(9), PNURM(KF)) CONTINUE	MAIN
35	UU JO 1=1,9 PNDRM(JP+I)=XX(I) CDNTINUE	MAIN MAIN MAIN
69	WRI1E(2,69) TNORM FORMAT(9F10.6) WRITE(6,310) IB,PARM	MAIN MAIN MAIN
310 71	FORMAT(3X,13,3X,F7.2,3X,F7.2,3X,F7.2,4X,F6.3,5X,F7.2,2X,F6.2,3X, 1F6.3,3X,F7.3,3X,F7.5) CONTINUE	MAIN MAIN MAIN
9999 747	ČONTINUE WRITE(6,747) ICOUNT ERIMATIZOVIND DE BLOCKS WRITTEN- 1 12//)	MAIN
171	STOP END	HAIN

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1:			SUBROUTINE SMTH15(PSPEC,N,PSPED,NLAP)	SMTH	10
2:	ç			SMTH	20
3:	ç		***************************************	SWIH	30
5:	2000		THIS SUBROUTINE SMOOTS THE SPECTRAL ESTIMATES BY MEANS OF A TRIANGULAR WINDOW WITH A SPAN OF 15	SMTH	50 60
8:	č		******	SMTH	80
9:	č			SMTH	90
10:			DIMENSION PSPEC(1), PSPED(1)	SMTH	100
11:				SMTH	110
12:		1	PSPEU(1)=(8.0#PSPEU(1)+14.0#PSPEU(2)+12.0#PSPEU(3)+10.0#PSPEU(4)+ 18.0#PSPEU(5)+4.0#DSPEC(4)+4.0#PSPEC(7)+2.0#DSPEC(3)+40.0	SMIN	120
14:				SMTH.	140
15:	•		NM=NHALF-I	SMTH	îśŏ
16:			N [= 2	SMTH	160
17:			NS=1	SMTH	170
18:				SMIH CMTU	180
20:		5		SMTH	200
21:		2		SMTH	210
22:			ŇĨ ≠8	SMTH	220
23:			NS=8	SMTH	230
24:		6		SMTH	240
25:			DU I II=NI,NM,NS	SMIH	250
201				CMTH	270
28			PSPED(K)=0_00	SMTH	280
29:			DO 1 I=1.15	SMTH	290
30:				SMTH	300
31:			P=J	SMTH	310
32:			INDX=II+J	SMTH	320
33:			$I \in (I \cap D X \cdot C X \cap D X) = 1 \cap D X$	SMIN	330
35:			$1 \in (1 \text{ ND} X + \mathbb{C} X + \mathbb{C})$ $1 \text{ ND} X = 1 \text{ ND} X - \mathbb{N} H A 1 \in \mathbb{C}$	SMTH	350
35:			PSPED(K) = PSPED(K) + (8.0 - ABS(P)) * PSPEC(INDX)/64.0	SMTH	360
37:	1		CONTINUÉ	SMTH	370
38:			PSPED(K+1)=(8.0*PSPEC(NHALF)+14.0*PSPEC(NHALF-1)+12.0*PSPEC(NHALF-	SMTH	380
39:		1	12)+10.0*PSPEC(NHALF-3)+8.0*PSPEC(NHALF-4)+6.0*PSPEC(NHALF-5)+4.0*	SMTH	390
40:		2	22575266(NHALF-6)+2.0*P5PE6(NHALF-7))/64.0	2WTH	400
413				SMIH	410
43:			DO TO SETTURELINS	SMTH	430
44			$\hat{P}S\hat{P}F\hat{C}(J) = PSPED(J)$	<u>šмrн</u>	440
45:	1	0	CONTINUE	SMTH	450
46:		-	RETURN	SMTH	460
47:			END	SMTH	470

1:2:3:	CCCC	SUBROUTINE PWRP(XDATA, PSPEC, NSIZE, DELTT) ***********************************	PWRP PWRP PWRP PWRP	10 20 30 40
5: 6: 7:		CONTAINED IN THE ARRAY XDATA	PWRP	60 70
8	Č	*******	PWRP	80 90
10: 11: 12: 13:		DIMENSION XDATA(1),PSPEC(1),IWK(12) INTEGER IWK	PWRP	100
		CUMPLEX GAMN FSIZE = NSIZE CALL FOURTAINSIZE		130
15:	C	COMPUTE VALUE OF CORRECTION FACTOR FOR A FINITE FFT.	PWRP	150
17:		J = 0 DO 3 I=1,NSIZE,2	PWRP PWRP	170 180
19:	_	J = J + 1 PSPEC(J)=(XDATA(I)**2*XDATA(I+1)**2)*CORTX	PWRP	190 200
22: 23:	3	CUNTINUE RETURN END	PWRP PWRP PWRP	210 220 230

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1:	~	SUBROUTINE TAPER1(XDATA,NREC,DFREQ)	TAPE	10
2:	č	*****	TAPE	20
4:	č		TAPE	40
5:	Č	THIS SUBROUTINE USES A COSINE TAPER WINDOW TO TAPER XDATA	TAPE	5ŏ
<u>6</u> :	ç		TAPE	60
81	ç	***************************************	TAPE	- 40
9:	č	XDATA REAL ARRAY CONTAINING THE INPUT DATA	TAPE	- 90
10:	Ç	NREC NUMBER OF POINTS IN XDATA	TAPE	100
11:	ç	DFREQ SAMPLING FREQUENCY	TAPE	110
13:	č	****	TAPE	130
14:	č		TAPE	140
15:		DIMENSION XDATA(1), TCDS(400)	TAPE	150
19:		DATA IFLAGION	TAPE	160
18:	1	DE(1) = 1.6 (DEE E)	TAPE	180
ĩš:	•	TREC=DELTT*(NREC-1)	TAPE	190
20:		P1=3.141592654	TAPE	200
21:			TAPE	210
23:				230
24:		00 10 I=1,NTAPR	TAPE	240
25:		XJ=J*DELTT	TAPE	250
29			IAPE	250
28:		$T_{COS}(1) = (1, 0 - COS(ARG))/2, 0$	TAPE	290
29:	10	ĊŎŇŤĬŇŲĖ	TAPE	290
30:		IFLAG=IFLAG+1	TAPE	300
32:			TAPE	310
33:			TAPE	330
34:		XDATA(I) = XDATA(I) * TCOS(I)	TAPE	340
35:	20	XDATA(JC)=XDATA(JC)*TCDS(I)	TAPE	350
37:	2 V	CONTINUE Retirn	TAPE	300
38:		ËND THE	TAPE	380
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0001	FTN,L	
0002		SUBROUTINE GRAF(A, XMAX, XMIN, NUMX, NUMY, XSIZE, YSIZE,
0003		IKEY, ANGLE, YPEAK, YYT, KC)
0004		
0005		TELEGUU
0007		
ŏŏŏå		$DX = 2 \cdot 0 + x \leq 17 \in I \in I \cap AT (NUM X = 1)$
ě 0 00		
0010		T=ÅNGLE*3.14159/180.0
0011		ŔĂŊĠĔ=Ź.Ŏ*ĊŎŚ(Ť)ŦŹŚĬŹĖ/(XMAX-XMIN)
0012		$DY = 2.0 \times SIN(T) \times YSIZE/FLOAT(NUMY-1)$
0013		YPK=YPEAK*RANGE
0014		CALL PLUI(0.0,0.0,-3)
0019	10	DU = 10 = 1 + 2 + NOMA
0018	10	
0019		XP=0.0
ŎŎŹÓ		YP = (A(1) + XMIN) * RANGE + RY
0021		$IF((YP-RY) \cdot GT \cdot YPK) YP = YPK + RY$
0022		OC(1) = YP
0023		CALL_PLOT(XP,YP,3)
0024		DO 20 I=2, NUMX
0025		XF = XP + UX
0020		TP=(A(I)=XMIN)#KANGE + RY
0027		$\Gamma((1) - X) \cdot G(1) \cdot TPK)$ $TP = TPK + RT$
0029	20	C(1) = T
0030	20	
0031		$\hat{Y}\hat{Y}T$ (KC) = DC (NUMX)
0032		RETURN
0033	100	IFLAG=IFLAG+1
0034		IF(IR-EQ-0) GO TO 200
0035		RY=RY+DY
0030		
0037		
0139		IFT(VD=VV).CT.VDEVV
0040		
ŏŏ 4 ĭ		
0042		DC(NUMX) = YP
0043	110	I = I - I
0044		[F(I,EQ,0)] GO TO 160
0045		YO=YP
0049		YP = (A(I) - XMIN) * RANGE + RY
0047		IF((TP-KY).GI.YPK) YP=YPK+KY
0040		$1 \in 1 \times 1 \to 1 \to$
0050		
ŏŏšĭ		
0052		
0053	120	B(=0C(I)
0054		B2=DC(I+1)
0055		XT = (B1 - YP) * DX / (YO - YP + B1 - B2)
00256		Y [= [YU-YP] *XT/OX
0026		入約三人ピキス
0050		CALL DEDT (YM. YM. 2)
0037		CALL FLUI (ANI) INIZI

ZSIZE,

0060 0061 0063 00645 000645 000645 00067 000689 00067 00071 000723 00074	150	$I = I - 1$ $I \in [I - EQ.0] GO TO 160$ $YO = YP$ $YP = (A(I) - XMIN) * RANGE + RY$ $IF ((YP - RY) \cdot GI \cdot YPK) YP = YPK + RY$ $XP = XP - DX$ $IF (YP - LE \cdot DC(I)) GO TO 150$ $B 1 = 0C(I + 1)$ $B 2 = 0C(I + 1)$
0076 0076 0077 0078 0079	160.	GO TO 110 IR=0 KC=KC+1 YYT(KC)=DC(NUMX)
0080 0081 0082 0083 0084	200	RETURN IR=1 XP=0.0 RY=RY+DY I=1
0085 0086 0087 0088 0088 0089 0090	210	<pre>YP=(A(1)-XMIN)*RANGE+RY IF((YP-RY).GT.YPK) YP=YPK+RY IF(YP.LT.DC(1)) GO TO 250 DC(1)=YP CALL PLOT (0.0,YP,3) I=I+1 IF (I.GT.NUMX) GO TO 260</pre>
0092 0093 0094 0095 0095 0096 0097 0098		YO=YP YP=(A(I)-XMIN)*RANGE+RY IF((YP-RY).GT.YPK) YP=YPK+RY XP=XP+DX IF (YP.LT.DC(I)) GO TO 220 DC(I)=YP CALL PLOT(XP,YP,2)
0099 0100 0101 0102 0103 0104 0105	220	GO TO 210 B1=DC(I) B2=DC(I-1) XT=(B1-YP)*DX/(YO-YP+B1-B2) YT=(YO-YP)*XT/DX XM=XP-XT YN=YP+YT YN=YP+YT
0106 0107 0108 0109 0110 0111 0112 0113 0114 0115 0116	250	CALL PLOI (XM,YM,Z) I=I+1 IF (I.GT.NUMX) GO TO 260 YO=YP YP=[A(I)-XMIN)*RANGE+RY IF({YP-RY).GT.YPK) YP=YPK+RY XP=XP+DX IF (YP.LT.DC(I)) GO TO 250 B1=DC(I-1) B2=DC(I) XT=(B1-YQ)*DX/(YP-YO+B1-B2)
0117 0118 0119		$\begin{array}{l} YT = (YP - YU) * XT / UX \\ XM = XP - DX + XT \\ YM = YO + YT \end{array}$
0120 0121 0122 0123 0124 0125 0126 0127	260	CALL PLOI (XP,YP,2) GO TO 210 IR=1 KC=KC+1 YYT(KC)=DC(NUMX) RETURN END END
0128	LIST	ENO ***

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PROGRAM ARA DIMENSION Y(90),YY(250), IARTF(20) IFLAG=0 FAMA(1),20) ISKIP FORMAT(12) IF (15K1P, E0,0) GO TO 7 O 7 L=1,ISK1P CALL EXEC(3,001310B) CANTINUE CALL EXEC(3,001310B) CALL EXEC(3,001310B) CALL EXEC(3,001310B) CALL EXEC(3,001310B) CALL FVG (1010) FORMAT(FETTER NUMY,NUMX,NSKIP IN 213 AND 11 FORMAT") FORMAT(121,110) READ(1,110) NUMY,NUMX,NSKIP IN 213 AND 11 FORMAT") FORMAT(213,111) RITE(6,400) FORMAT(FETTER NUMY,NUMX,NSKIP IN 213 AND 11 FORMAT") READ(1,180) YMAX,YMIN,YPEAK,CAL IN F6.0 FORMAT") READ(1,180) YMAX,YMIN,YPEAK,CAL IN F6.0 FORMAT") READ(1,180) YMAX,YMIN,YPEAK,CAL IN F6.0 FORMAT") READ(1,160) T,(1ARTF(1),1=1,17) RTTE(6,400) HORMAT(FETTER YMAX,YMIN,YPEAK,CAL IN F6.0 FORMAT") READ(1,160) T,(1ARTF(1),1=1,17) RTTE(6,200) HITE(6,200) RTTE(6,200) RTTE(6,200) RTTE(6,200) MATTE(6,200) MATTE(6,300) MATTE(6, 0002 0003 0004 0005 FTN,L 0008 0009 0011 0012 0013 0015 0016 0017 0018 0021 0021 0022 0022 00223 00223 00225 -300 310 320 39 40 0055 0056 0057 0058 0059

0060		K(=(K/2)*2
0001		
0062	10	1 + 1 + 1 + 1 + 1 + 1 + 1 + 1 + 1 + 1 +
2200	17	
0065		
2200		r = r
0067	19	
0024	12	
ñăăă	12	$1 \in (k - 1) \in [1, 1]$
0070	25	
ŏŏżĭ	~ ~ ~	I = (K + E) - I = K = 0
ŎŎŻŹ	•	
0073	561	FORMAT(10X.13)
0074		IF (IFLAG-2) 11,26,11
0075	26	IFLAG=0
0076		GO TO 11
0077	24	CONTINUE
0078		IFLAG=0
0079	13	DO 10 JK=1,NUMX
0080		Y(JK)=SQRT(Y(JK)*CAL)
0081	10	CONTINUE
0082		CALL GRAF(Y, YMAX, YMIN, NUMX, NUMY, XSIZE, YSIZE, ZSIZE, KP, ANG,
0083		IYPEAK, YY, KC)
0084	11	
0085		
0000		CALL IIME(TTTKUTASILETISTUTT
0001		CALL DURDA(ASILE)ISILE,ANG)
0000		STOP END
0000		FN04
****	LIST	

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0001 0002 0003 0004 0005 0006 0007 0008	FTN	L SUBROUTINE EIXO(XSIZE) DIMENSION ITX(10) DATA ITX/2HFR,2HEQ,2HUE,2HNC,2HY,2HIN,2H H,2HER,2HTZ/ DATA IHZ/2HHZ/ X=2.*XSIZE DX=X/8.0 YX=0.1
0009 0010 0011 0012 0013 0014 0015 0016		XP=0.0 CALL PLOT(0.0,-0.1,-3) CALL PLOT(XP,-YX,3) CALL PLOT(XP,0.0,2) DO 10 I=1,8 XP=XP+DX CALL PLOT(XP,0.0,2) CALL PLOT(XP,0.0,2) CALL PLOT(XP,-YX.3)
0017 0018 0019 0020 0021 0022 0023	10	CALL PLOT(XP,0.0,2) CONTINUE CALL PLOT(0.0,0.0,3) HGT=0.14 YP=0.3+HGT/2.0 XP=.02-HGT/2.0 FN=0.0
0024 0025 0026 0027 0028 0029 0030		CALL NUMB(XP,-YP,HGT,FPN,0.0,-1) XP=XP+DX FPN=5.0 CALL NUMB(XP,-YP,HGT,FPN,0.0,-1) XP=DX-HGT D0 20 I=2,8 XP=XP+DX FPN=FL0AT(I*5)
0032 0033 0034 0035 0036 0036 0037 0038	20	CALL NUMB(XP,-YP,HGT,FPN,0.0,-1) CONTINUE XS=X/2.0 -9.0*HGT YS=0.3 + 2.5*HGT CALL SYMB(XS,-YS,HGT,ITX,0.0,18) RETURN END
0039 ****	LIST	END\$ END ****

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00.01	FTN.	.)	
0002	,	SUBROUTINE TIME(YY,KC,XSIZE,TS,DT)	
0003		DIMENSION YY(1), ITM(4), ITN(4)	
0004		DATA IIM/2HII;2HME;2H I;2HN / DATA ITN/2HMI;2HNU;2HTC;2HC /	
0006		X=2,0 + XSI7F + 0,2	
0007		TT=DT/2.5	
0008		XY=0.1	
0010		XI=KC/IT 1 =	
ŏŏīĭ		CALL PLOT(X.0.13)	
0012		CALL PLOT(XY,YY(1),3)	
0013	•	(ALL PLDT(0, 0, YY(1), 2))	
0015		I = IT * J + I	
0016		ČAĽĽ PLOT(0.0,YY(1),2)	
0017		CALL PLOT(XY, YY(I), 3)	
0019	10		
0020		CALL PLOT(0,0, YY(KC),2)	
0021		CALL PLOT(XY, YY(KC), 3)	
0023		CALL PLUI (0,0,14(KC),2)	
0024		HGT=0.14	
0025		XP=0.2 + HGT/2.0	
0027		TP=0.02 - HGT/2.0 EN=TS	
0028		YR = YY(1) + YP	
0029		CALL NUMB(XP,YR,HGT,FN,0.0,-1)	
0031		$N_1 = N_1 - 1$ D0 20 J= 1.4 11	
0032		I = IT * J + I	
0033		YR = YY(I) + YP	
0035		CALL NUMB(XP-XP-HCT-EN.O.O1)	
0036	20	CONTINUE	
0037		I = IT * K I + 1	
0039		TETTET + UI	
0040		YR = YY(I) + YP	
0041	21	CALL NUMB(XP, YR, HGT, FN, 0.0, -1)	
0043	21	YR = YY(KC) + YP	
0044		CALL NUMB (XP, YR, HGT, FN, 0.0, -1)	
0045		XS=2.0*HGT	
0047		13-111NU) + 1+37HUI + U+3 CALL SYMB(+XS+YS+HGT+TTM+0+0+7)	
0348		YS=YS - 1.5*HGT	
0049		CALL SYMB(-XS,YS,HGT,ITN,0.0,7)	
0051		RFTURN	
0052		END	
0053	1 1 6 7		
****	CT31	LND TTTT	

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0001 0003 0004 0005 00008 00010 00113 00113 00114 0015 00117 00113 00117 00118	FTN,	L SUBROUT: ANG=ANG; YT=YSI2; XL=+2.0; YDP=2.0; YDP=2.0; YDP=2.0; CALL PLC CALL PLC CALC PLC CALC PLC CALC PLC CAL	<pre>INE BORDA(XSIZE,YSIZE,ANG) *3.1415/180.0 =*COS(ANG) (XSIZE+0.7) *0.3 *(YT+1.0))*0.5 DT(XR,YDW,3) DT(XR,YUP,2) DT(XL,YUP,2) DT(XL,YUP,2) DT(XL,YDW,2) P(0.0,YNXT,3) DT(0.0,YNXT,2)</pre>
****	LIST	END ****	

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00000000000000000000000000000000000000	* * * XPAND * XPAND I A 8 *	NAM NAM NDDAAZAALPAAALPAAALSIDDALPAAALSIDDALPAAALSIDDBLPAA ILDBBPAAALSIDBBLPAA ILDBBPAAALSIDBBLPAP SSUU D** NOT SSUU D**	XPAND, 7 XPAND, I RTN XPAND, I RTN XPAND A, I E, SLA, ERA *-2 O, I I E = B0000377 XPAND B, I E = SLB, ERB *-2 I, I I = B000377 XPAND B, SLB, ERB *-2 I, I R TN, I 1 1 0 1	FORTRAN SUBROUTINE CALLING SEQUENCE - CALL XPAND(I,II,I2) EXPANDS INTEGER WORD I INTO TWO INTEGER WORDS II AND 12, BITS 15-8 OF I TO 7-0 OF I1, ALL OTHER BITS CLEARED. STORE RETURN ADDRESS GET I CET WORD TO BE EXPANDED CHASE THOSE LITTLE BITS BITS 15-8 MOVE TO 7-0 CLEAR BITS 15-8 GET ADDRESS OF THE SECOND ARGUMENT CHASE THOSE LITTLE BITS SIORE I2 BITS 7-0 MOVE TO 7-0 CLEAR BITS 15-8 GET ADDRESS OF THE SECOND ARGUMENT CHASE THOSE LITTLE BITS SIORE I2 BITS 7-0 MOVE TO 7-0 CLEAR BITS 15-8 GET ADDRESS OF THE SECOND ARGUMENT CHASE THOSE LITTLE BITS STORE I1 RETURN
C001 00003 0005 0006 0008 0008 0010 00112 0013 0014 0016 0016 0017 0016 0017 0014 0016 0017	F (N, 10 100 20 LIST	L SINMACALE UDOEROCKJINOLLARR CREDICKJINOLLARR CREDNU SINDSI	OUTINE CVRT GER EBASC NSION INPUT 0 JJ=1,10 (8) INPUT 0 I=1,45 XPAND(INPU ASC(K) BASC(J) T(I)=256*K T(I)=10R(INI INUE JJ-1)*9 CODE (INPUT,100) MAT(9F10.3) INUE RN **	(Y) (45),Y(90) T(I),K,J) PUT(I),J) (Y(KL+I),I=1,9)

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***** 1234567890112345678901234567890 10 230 40 50 70 * ** * PROGRAM FEATURE \$ ***** 24 THIS PROGRAM DIVIDES THE EEG POWER SPECTRUM IN SEVEN FREQUENCY 80 90 00 THE FREQUENCY BANDS ARE DEFINED AS FOLLOWS, DELTA THETA ALPHA SIGMA BETA1 BETA2 FASTF 0.0 4.0 8.0 11.5 16.0 21.0 30.0 3.5 7.5 11.5 15.5 20.5 29.5 40.0 110 120 130 ΗZ HZHZHZ 140 150 160 170

 DELA1 -- 10.0 - 20.2 HZ
 MAIN

 PASTF -- 30.0 - 20.0 HZ
 MAIN

 PASTF -- 30.0 - 20.0 HZ
 MAIN

 DIMENSION XSGR(400), EXKURT(400), SKEW(400), YSQR(400, 1),
 MAIN

 HYSKEW(400, 1), YEXKUR(200, 1)
 MAIN

 ISSUE SCORE
 YSQR(1, 1), YSQR(1, 1), YSQR(1, 1), YSQR(1, 1), SKEW(1, 1), SKEW(1), YYEXKUR(1, 1), MAIN

 MAIN
 MAIN

 DIMENSION DELTAP(6), THETAP(6), ALPHAP(6), ALPHAP(6), SIGMAP(6), MAIN

 DIMENSION OELTAP(6), THETAF(6), YALPHA(6), YSIGMA(6), YBETA1(6), MAIN

 DIMENSION OELTAF(6), YHETAG(5), YALPHA(6), YSIGMA(6), YBETA1(6), MAIN

 DIMENSION OELTAF(800), THETAF(800), ALPHAF(800), SIGMAF(800), MAIN

 DIMENSION VOELTAF(800), THETAF(800), ALPHAF(800), SIGMAF(800), I, ALF400, I), ALF400, I), MAIN

 DIMENSION VI400, I), YW(400, I), X4001, DI4001400, I), ALF400, I), ALF400, I), MAIN

 DIMENSION VI400, I, SIGMAF(10), I, ALF400, I), ALF400, I), MAIN

 DIMENSION VI400, I, SIGMAF(10), I, THAFF400, SIGMAF(800), FEMEAN(400), I), MAIN

 DIMENSION VI400, I, FSTF(11), YA4001, DI400140, I, ALF400, I), ALF400, I), MAIN

 DIMENSION VI400, I, FSTF(11), YA4001, DI400, I, THAFF400, SIGMAF(800), FMAIN

 MAIN

 DIMENSION VI400, I, FSTF(11), FASTF(11), FASTF(10), FASTF(10), FMAIN

 MAIN

 DIMENSION VI400, I, FSTF(11), FASTF(11), FASTF(10), FASTF(10), FMAIN

 -----ΗŻ 420 430 440 450 560 570 580 590 590 600 610 620 630 640 650 MAIN MAIN 66: c 660 ENTER NSKIP. IF NSKIP=0 ALL EPOCHS ARE PROCESSED, NSKIP=1 ONLY THE 670

295

PAGE

			PAGE 2
68: 69: 70: 71: 72:	C C	ODDS ARE PROCESSED AND NSKIP=2 THE EVEN ONES ARE PROCESSED DELFF=0.5 NC=40./DELFF+1 CHIS01=66.3	MAIN 680 MAIN 690 MAIN 700 MAIN 710 MAIN 720
73: 74: 75: 76: 77:		CMX=200.0 CMN=40.0 EMN=-1.0 EMX=3.0 SMN=-0.5	MAIN 730 MAIN 740 MAIN 750 MAIN 750 MAIN 760 MAIN 770
78: 79: 80: 81: 82:		SMX =0.5 CEM=100.0 FMX=15.0 PHT=0.4	MAIN 780 MAIN 790 MAIN 800 MAIN 810 MAIN 820
83: 84: 85: 86: 87:		PHT=0.15 PLT=0.15 DELTAH=3.5 THETAH=7.5 At PHO1=6.5	MAIN 830 MAIN 840 MAIN 850 MAIN 850 MAIN 860 MAIN 870
88: 89: 90: 91: 92:		ALPHAH=11.5 SIGMAH=15.5 BETA1H=20.5 BETA2H=29.5 K=81	MAIN 880 MAIN 890 MAIN 900 MAIN 910 MAIN 920
93: 94: 95: 96:		TF=(DELTAH+0.5)/DELFF IAF=(THETAH+0.5)/DELFF ISF=(ALPHAH+0.5)/DELFF IOF=(ALPHOL+0.5)/DELFF IBIE=(SIGMAH+0.5)/DELFF IBIE=(SIGMAH+0.5)/DELFF	MAIN 930 MAIN 940 MAIN 950 MAIN 960 MAIN 970
98: 99: 100: 101: 102:	100	182F=(BETA1H+0.5)/DELFF IFTF=(BETA2H+0.5)/DELFF NF=10./DELFF+1 READ(5,100) NSKIP FORMATIII)	MAIN 980 MAIN 990 MAIN1000 MAIN1010 MAIN1020
103: 104: 105: 106: 107:	C C	ISKIP=NSKIP+1 IT IS THE # OF EPOCHS WITH ARTIFACTS AND IARTF IS THE ARRAY CONTAINING THEIR NUMBERS READ(5,110) ITA,(IARTF(I),I=1,ITA) FORMAT(2013)	MAIN1030 MAIN1040 MAIN1050 MAIN1050 MAIN1070
108: 109: 110: 111: 112:	510 C	WRITE(6,510) ITA,(IARTF(I),I=1,ITA) FORMAT(50X,I4) TO READ THE STARING AND ENDING TIMES FOR THE PLOTS READ(5,111) XST,XTIME FORMAT(2F10,0)	MAIN1080 MAIN1090 MAIN1100 MAIN1110 MAIN1120
113: 114: 115: 116: 117:	č 112	TO READ THE EPOCH NO. WHERE THE PLOT IS TO BE PARTITIONED ISS=1 READ(5,112) ITT FORMAT(I3) DO 1000 JJ=1.2	MAIN1130 MAIN1140 MAIN1150 MAIN1160 MAIN1170
118: 119: 120: 121: 122:		DO 300 J=1,800 DELTAF(J)=0.0 THETAF(J)=0.0 ALPHAF(J)=0.0 ALPHOF(J)=0.0	MAIN1180 MAIN1190 MAIN1200 MAIN1210 MAIN1220
123: 124: 125: 126: 127:	300	SIGMAF(J)=0.0 BETA1F(J)=0.0 BETA2F(J)=0.0 FASTFF(J)=0.0 CONTINUE	MAIN1230 MAIN1240 MAIN1250 MAIN1260 MAIN1270
128: 129: 130: 131: 132:	30 200	GO TO (30,31,32),ISKIP WRITE(6,200) FORMAT(20X,*ALL EPOCHS ARE BEING PROCESSED*//) DELT=2.5 GO TO 33	MAIN1280 MAIN1290 MAIN1300 MAIN1310 MAIN1310 MAIN1320
133: 134: 135:	31 210	ŴŔIŦĔ(6,210) Format(20X,"ONLY ODD EPOCHS ARE BEING PROCESSED"//) DELT=5.0	MAIN1330 MAIN1340 MAIN1350

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136:		GO TO 33
1 17:	. 32	WRITE(6,220)
138:	220	FORMAT (20%, ONLY EVEN EPOCHS ARE BEING PROCESSED ///)
139:		DELT=5.0
140:	33	
141:		101=0
1250		ič2=-1
175:		
1 4 3 .		8 Εχή(1,120, ΕΝΠ#99) ΡΝΠRM
144.	120	
1424	120	CO TO (12-19-20) ISKIP
179:	20	
14/1	20	
148:	21	$N = \frac{10}{20} \frac{172}{71}$
149	10	$\frac{1}{1}$
120:	72	IT(IFLA0) IC\$CC\$IC
1211	22	$N_1 = (10/2) + 2 + 1$
1 22		1 + (K) - 1 = 1 = 1 + 1 = 1 = 1 = 1 = 1 = 1 = 1 =
153	12	
124	25	[P[10-14K]P[M]] 27923724
155:	25	
156:		
157:	65 L	
158:		1+(1+LAG-2) /1,20,/1
159:	26	IFLAG=0
160:		<u>GO 10 /1</u>
161:	24	<u>CONTINUE</u>
162:		I FLAG=0
163:		
164:		DELTA(IC)=0.0
165:		$THETA(IC) = Q \cdot Q$
166:		ALPHA(IC)=0.0
167:		ALPHD(IC)=0.0
168:		SIGMA(IC)=0.0
169:		BETA1(IC)=0.0
170:		BETA2(IC)=0.0
171:		FASTF(1C)=0.0
172:		TPOWR=0.0
173:	C	COMPUTE THE TOTAL POWER
174:	Ŭ	DO 1 J=1 + NC
1751		Ť₽OŴR=T₽O₩R+PNORM(J)
176:	1	CONTINUE
177:	-	ŠKĖW(IČ)=PNORM(85)
178:		$E \times K U R T (IC) = P N O R M (86) - 3 \cdot 0$
179:		$\tilde{X}SOR(1C) = PNORM(87)$
1861		10=1
		ÎT=ÎTF
182:		ĨÅ=ĨÅF
193:		Î∩=ÎOF
184		ÍŠ=ÍŠF
185		
186		182=182F
197:		Î Ê Î Ê Î Ê Î Ê
188		ici = ici + 2
180		ič2=ič2+2
100		00 2 I=1.NF
161:		
1 92:		ÎĔ(ÎĎ-ÎTE) 3.3.4
1 62	3	DELTA(IC)=DELTA(IC)+PNORM(ID)
1 04		TĒ DĒ TĀF (ICI) - PNORM (ID)) 50,4,4
165	50	$\vec{D}FITAF(ICI) = PNORM(ID)$
104		DFITAF(IC2) = (ID-1) * DELFF
197	4	
168-	•	ĪĒ(ĪT-ĪAF) 5.5.6
166	5	THE TA(IC) = THE TA(IC) + PNORM(IT)
200	-	IFTTHETAF(ICI) -PNORM(IT)) 51,6,6
201:	51	THETAF(ICI) = PNORM(IT)
202	~	THETAF(IC2) = (IT-1) * DELFF
203:	6	10=10+1

 $\begin{array}{l} \textbf{MAIN13600} \\ \textbf{MAIN13800} \\ \textbf{MAIN13800} \\ \textbf{MAIN13800} \\ \textbf{MAIN13800} \\ \textbf{MAIN14400} \\ \textbf{MAIN14400} \\ \textbf{MAIN144200} \\ \textbf{MAIN144200} \\ \textbf{MAIN144200} \\ \textbf{MAIN144200} \\ \textbf{MAIN144200} \\ \textbf{MAIN144500} \\ \textbf{MAIN1455000} \\ \textbf{MAIN155600} \\ \textbf{MAIN155700} \\ \textbf{MAIN155600} \\ \textbf{MAIN155700} \\ \textbf{MAIN1577700} \\ \textbf{MAIN1577700} \\ \textbf{MAIN1577700} \\ \textbf{MAIN1577700} \\ \textbf{MAIN15777000} \\ \textbf{MAIN15777000} \\ \textbf{MAIN15777000} \\ \textbf{M$

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PAGE

204:		IF(IU-ISF) 60,60,66	MAIN2040
205:	60	ALPHU(IC) = ALPHU(IC) + PNUKM(IU)	MAIN2050
200		IF (ALPHUF (ICI)+PNUKA(IU)) 61,66,66	MAINZUGO
207	01		MAIN2070
208:		ALPHUP(102)=(10+1)*0ELFF	MAINZUBU
2091	00		MA 1 N 2 0 9 0
210:	-	1 + (1A + 1SF) / (1B)	MA 1N2100
211	1	ALPHA(IC) = ALPHA(IC) + PNUKM(IA)	MAIN2110
212:		1F(ALPHAF(ICI) +PNURM(IA)) 52,8,8	MAIN2120
213:	52	ALPHAF(ILI)=PNOKM(IA)	MA IN2130
214		ALPHAF(IC2)=(IA-1)*DELFF	MA1N2140
215	8	15=15+1	MAIN2150
210:	~	1F(15-1B1F) 9,9,10	MA IN 2160
21/1	9	SIGMA(IC) = SIGMA(IC) + PNORM(IS)	MAIN2170
218		IF(SIGMAF(ICI)-PN0RM(IS)) 53,10,10	MAIN2180
219	55	SIGMAF(ICI)=PNORM(IS)	MAIN2190
220:	••	SIGMA+(1,2)=(1,5-1)*DELFF	MA1N2200
221:	10		MAINZZIO
222		1F(181-182F) 40,40,41	MAINZZZU
223	40		MA 1N2230
<u> </u>	F /	IF(BE AIF(ILI)-PNUKM(IBI))24,41,41	MAINZZ40
442	54	BETALF(ICI)=PNURM(IBI)	MAINZZOU
220:			MAINZZOU
221:	41		MAINZZIU
228	10	1F(102-1F1F) 42,43,43	MAINZZSU
229	42		MAINZZAU
230:	e e	1F(BE)A2F(1C1)-PNUKM(1B2)/33,43,43	MAINZ300
231	22		MAINZALO
234	12		MAINZOZU
2331	45		MA 1N2 3 30
2341		1 F (1 F (TN) - 44 + 44 + 2 F (F (TF (TF (TF (TF (TF) + DNOD M / TFT)	MAINZ34U
2321	44	$r_{1} = r_{1} = r_{1$	MAIN232U
230:	E 4		MAINZOOU
221:	20	FASIFF[101]=FN0KM(1F1) FASIFF[101]=FN0KM(1F1)	MAINZOID
230.	2	CONTINUE	MAIN2200
2394	2	CONTINUE C_{1} CONTINUE C_{2} CONTI	MAIN2600
240.			MAIN2410
242.		CALL DIVCKIDNOPMINICHARDELES, INF. ISF. ICZ, DIT. DHT. FRIZZI	NATN2410
243.		CALL DIKCKIDNONMALDHAF OFFF IAF ISF IC2 DIT DHT FRTIAN	MAINZARO
244:		CALL PIKCKIPNOSM'S IGMAF, DELEF, ISF, IBJF, IC2, PLT, PHT, FRT(5))	MATN2440
245		CALL PIKCK (PNORM. BETALF. DELEF. IB) F. TB2F. IC2. PLT. PHT. FRT(6))	MATN2450
246		CALL PIKCK (PNORM BETA2F, DELEF, 182F, LETF, 162, PLT, PHT, FRT (7))	MA 1N2460
247:		CALL PIKCK (PNORM FASTEF DELFF IFTF NC + IC2 + PLT + PHT + FRT(8))	PAIN2670
248:	C		MAINZ480
249:	Č	COMPUTE THE PERCENTAGE OF POWER IN EACH BAND	MA IN 2490
250:	C	· · ·	MA1N2500
251:		DELTA(IC) = DELTA(IC)/TPOWR*100.0	MAIN2510
252:		THETA(IC)=THETA(IC)/TPOWR *100.0	MAIN2520
253:		ALPHA(IC)=ALPHA(IC)/TPOWR *100.0	MAIN2530
254:		ALPHO(IC)=ALPHO(IC)/TPOWR *100.0	MAIN2540
255:		SIGMA(IC)=SIGMA(IC)/TPOWR*100.0	MA IN 2550
256:		BETAL(IC)=BETAL(IC)/TPOWR *100.0	MA1N2560
257:		BEIA2(1C)=BEIA2(1C)/1PUNK *100.0	MAIN2570
258:		FASTF(IC)=FASTF(IC)/TPOWR *100.0	MAIN2580
259		LALL FMED(FRI)FMEAN,8,10)	MA IN2590
260:	/1		MAINZ600
261:	99	CONTINCE	MAINZ610
262:			MAINZ620
203:			MAINZ630
264			MAINZ640
2021	74		MAINZODU
409 ·		101-1410-1400-01 AIIME=AII01	MAINZOOU
2011			MATN26/U
2000			MAIN2000
270.		LEINSORI LACHISOTI RC. 80-81	MA 112090
571	81		MATNOTIO
	.		CONTRACT TO

PAGE 4

		PA	GE
80	CONTINUE RJ=(RJ/FLDAT(IC))*100.0	MA MA	[N272 [N272
	CALL SMOOTH(DELTA,IC)	MA	IN274
	CALL SMOOTH(THETA,IC) CALL SMOOTH(ALPHA,IC)	MA MA	[N276 [N277
	CALL SMOOTH(ALPHO,IC) CALL SMOOTH(SIGMA,IC)	MA MA	IN278
	CALL SMOOTH(BETA1,IC) CALL SMOOTH(BETA2,IC)	MA	IN280
	CALL SMOOTH(FASTF, IC)	MA	IN282
	CALL SMOOTH(XSQR,IC)	MA	IN284
	CALL SMOOTH(EXKURT,IC) CALL SMOOTH(SKEW,IC)	MA	LN285 IN286
5	CONTINUE YMAX=0.0	MA MA	IN287 IN288
	\dot{Y} MIN=10.0 CALL NERVALOEL TAE. (C. YMAY, YMIN)	MA	[N289
	CALL NERKA (THETAF, IC, YMAX, YMIN)	MA	INŽ91
	CALL NERKA(ALPHAF, IC, IMAX, IMIN)	MA	IN292
	CALL NERKA(SIGMAF,IC,YMAX,YMIN) CALL NERKA(BETA1F,IC,YMAX,YMIN)	MA	IN294 [N295
	CALL NERKA(BETA2F,IC,YMAX,YMIN) CALL NERKA(FASTEF.IC,YMAX,YMIN)	MA MA	[N296 (N297
	YMAX=SQRT(YMAX) YMIN=SQRT(YMIN)	MA	N298
	CALL PLOTR2(X, YSQR, YW, IC, 1, 400, 1, XST, XTIME, CMN, CMX, ICHAR, XUNIT2,	MA	INZÓC
	CALL SPARM(XSQR, IC)	MA	N302
33	FORMAT(//59X, PERCENTAGE OF REJECTIONS= +, F5.2)	MA	IN303 IN304
	CALL PLOTR2(X,DD,YW,IC,1,400,1,XST,XTIME,0.0,CEM,ICHAR,XUNIT2, 1YDELTA.0.0.0.0.0)	MA MA	LN305 IN306
	CALE VOST (DELTA, IC) CALE PLOTRZ(X, SL, YW, IC, 1, 400, 1, XST, XTIME, 0, 0, CEM, ICHAR, XUNIT2,	MA	IN307
	1YSIGMA,0.0,0.0,0)	MA	(N309
	CALL PLOTR2(X, FM, YW, IC, 1, 400, 1, XST, XTIME, 0. 0, FMX, ICHAR, XUNIT2,	MA	N311
	CALL FPARM(FMEAN, IC)	MA	INBI
	CALL PLOTR2(X, YSKEW, YW, IC, 1, 400, 1, 0.0, X(IC), SMN, SMX, ICHAP, XUNIT2, 1YV, 0.0, 0.0, 0)	MA MA	EN314 EN315
	CALL SPARM(SKEW,IC) CALL PLOTRZIX, YEXKUR, YW.IC.1.400.1.0.0.X(IC).EMN.EMX.ICHAR.XUNITZ	MA MA	IN318 IN317
	1YK,0.0,0.0,0) CALL SPARMEYKUPT. IC)	MA	
006	CONTINUE	MA	N320
	CALL PLOTR2(X,Y,Y,YW,IC,1,400,1,0.0,X(IC),YMIN,YMAX,ICHAR,XUNIT2,	MA	N322
	CALL WRTE(DELTAF)	MA	IN323 IN324
	CALL ARRAY(DELTAF,Y,IC2) CALL PLOIR2 (X,Y,YW,IC,1,400,1,0,0,X(IC),0,25,4,75,ICHAR,XUNIT2,	MA	[N325 [N326
	1YUNIT3,0.0,0.0,0) CALL WRTE(DELTAE)	MA	N327
	CALL PLOTR2(X, TT, YW, IC, 1, 400, 1, 0.0, X(IC), 0.0, CEM, ICHAR, XUNIT2,	MA	N329
	CALL VOST (THETA, IC)	MA	N331
	CALL RAIZ(THETAF, Y, 102) CALL PLOTR2(X, Y, YW, IC, 1, 400, 1, 0.0, X(IC), YMIN, YMAX, ICHAR, XUNIT2,	MA	N333
	ITHETAP,0.0,0.0,0) CALL WRTE(THETAF)	MA MA	[N334 [N335
	CALL ARRAY (THETAF, Y, IC2) CALL PLOTR2 (X,Y,YW, IC, 1, 400, 1, 0, 0, X(IC), 3, 0, 8, 0, ICHAR, XUNIT2,	MA	IN336
	1YUNIT3,0.0,0.0,0)	MA	IN338

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		PAGE
340:	CALL PLOTR2(X,AL,YW,IC,1,400,1,0.0,X(IC),0.0,CEM,ICHAR,XUNIT2,	MA IN 3400
341:	1YALPHA,0.0,0.0,0)	MA IN 3410
342:	CALL VOST(ALPHA,IC)	MA IN 3420
343:	ČALL RAIŻ(ALPHAF,Y,IC2)	MAIN3430
344:	CALL PLOTR2(X,Y,YW,IC,1,400,1,0.0,X(IC),YMIN,YMAX,ICHAR,XUNIT2,	MAIN3440
345:	1ALPHAP,0.0,0.0,0]	MAIN3450
340:	CALL WRITE(ALPHAF)	MAIN3480
347:	CALL ARRAY(ALPHAF,Y,IC2)	MAIN3470
348:	CALL PLOTR2 (X,Y,YW,IC,1,400,1,0.0,X(IC),7.5,12.5,ICHAR,XUNIT2,	MAIN3480
349:	1YUNIT3.0.0,0.0.0	MAIN3490
350:	CALL WRYE(ALPHAF)	MAIN3500
351:	CALL PLOTR2(X,AO,YW,IC,1,400,1,0.0,X(IC),0.0,CEM,ICHAR,XUNIT2,	MAIN3510
352:	1YALPHO,0.010.020]	MAIN3520
354: 355: 356:	CALL VOSI(ALPHO,IC) CALL RAIZ(ALPHOF,Y,IC2) CALL PLOTR2(X,Y,YW,IC,1,400,1,0.0,X(IC),YMIN,YMAX,ICHAR,XUNIT2, 1ALPHOP.0.0.0.00	MAIN3530 MAIN3540 MAIN3550 MAIN3560
357:	ČÁLĽ WŔŤĚ(ÁĽPHOF)	MAIN3570
358:	CALL ARRAY (ALPHOF, Y, IC2)	MAIN3580
359:	CALL PLOTR2 (X,Y,YW,IC,I,400,1,0.0,X(IC),6.5,12.5,ICHAR,XUNIT2,	MAIN3590
361:	CALL WRTE(ALPHOF)	MAIN3610
362:	CALL RAIZ(SIGMAF,Y,IC2)	MAIN3620
363:	CALL PLOTR2(X,Y,YW,IC,1,400,1,0.0,X(IC),YMIN,YMAX,ICHAR,XUNIT2,	MAIN3630
364:	1SIGMAP,0.0,0.0,0)	MAIN3640
365:	CALL WRTE(SIGMAF)	MAIN3650
366:	CALL ARRAY(SIGMAF,Y,IC2)	MAIN3660
367:	CALL ARRAY(SIGMAF,Y,IC2)	MAIN3670
368:	1YUNIT3,0.0,0.0)	MAIN3680
369:	CALL WRTE(SIGMAF)	MAIN3690
370:	CALL PLOTR2(X,B1,YW,IC,1,400,1,0.0,X(IC),0.0,CEM,ICHAR,XUNIT2,	MAIN3700
371: 372: 373: 374:	1YBETA1,0.0,0.0,0) CALL VOST(BETA1,IC) CALL RAIZ(BETA1F,Y,IC2) CALL RAIZ(SETA1F,Y,IC2) CALL PLOIR2(X,Y,YW,IC,1.400,1.0,0.X(IC),YMIN,YMAX,ICHAR,XUNIT2.	MAIN3710 MAIN3720 MAIN3730 MAIN3740
375: 376: 377:	18ETA1P,0.0,0.0,0) CALL WRTE(BETA1F) CALL ARRAY(BETA1F,Y,IC2) CALL ARRAY(BETA1F,Y,IC2)	MAIN3750 MAIN3760 MAIN3770
379: 380: 381:	CALL PLOTR2 (X, F,	MAIN3790 MAIN3790 MAIN3800 MAIN3810
382:	1YBETA2,0.0,0.0,0)	MAIN3820
383:	CALL VOST(BETA2,IC)	MAIN3830
384:	CALL RAIZ(BETA2F,Y,IC2)	MAIN3840
385:	CALL RAIZ(BETA2F,Y,IC2)	MAIN3850
386:	18ETA2P,0.0,0.0,0)	MAIN3860
387:	CALL WRTE(BETA2F)	MAIN3870
388:	CALL ARRAY(BETA2F,Y,IC2)	MAIN3880
389:	CALL PLUIR2 (X,Y,YW,IC,I,400,I,0.0,X(IC),20.5,30.0,ICHAR,XUN(12,	MAIN3890
390:	IYUNIT3,0.0,0.0,0)	MAIN3900
391:	CALL WRTE(BETA2F)	MAIN3910
392:	CALL PLUIR2(X,FF,YW,IC,1.400,1.0.0,X(IC),0.0,CEM,ICHAR,XUNIT2,	MAIN3920
393: 394: 395:	1YFASTF,0.0,0.0,0) CALL VOST(FASTF,IC) CALL RAIZ(FASTF,Y,IC2) CALL RAIZ(FASTFF,Y,IC2)	MAIN3930 MAIN3940 MAIN3950
397: 398: 399:	1FASTFP,0.0,0.0,0) CALL WRTE(FASTFF) CALL ARRAY(FASTFF,Y,IC2)	MAIN3970 MAIN3970 MAIN3980 MAIN3990
400:	CALL PLOTR2 (X,Y,YW,IC,1,400,1,0.0,X(IC),29.0,40.0,ICHAR,XUNIT2,	MAIN4000
401:	1YUNIT3,0.0,0.0,0)	MAIN4010
402:	CALL WRTE(FASTFF)	MAIN4020
403:	XST=X(IC) + 5.0	MAIN4030
404:	XTIME=0.0	MAIN4040
405:	ISS=ITT+1	MAIN4050
406:	IF(ISS.GT.240) STOP	MAIN4060
407:	ITT=400	MAIN4070
408:	1000 CONTINUE	MAIN4080
409:	STOP	MAIN4090
410:	END	MAIN4100

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2345678901234	11 12 10	SYMEMSION BAND(1) IC2=2+IC SUMPI=0.0 SUMF2=0.0 SUMF2=0.0 PMIN=100.0 PMIN=100.0 PMAX=0.0 TF(BAND(1)]1.12,11 SUMF1=SUMF1+BAND(1) SUMF1=SUMF1+BAND(1) SUMF1=SUMF1+BAND(1)+BAND(1) SUMP1=SUMP1+BAND(1)+BAND(1) SUMP1=SUMP1+BAND(1+1) FMAX=AMAX1(FMAX,BAND(1)) PMAX=AMAX1(FMAX,BAND(1)) PMAX=AMAX1(FMAX,BAND(1)) FMIN=AMIN1(FMIN,BAND(1+1)) G0 TO 10 G0 TO 10 SUMF2=SUMF2/FIC SUMF2=SUMP2 SUMF2=SUMP2/FIC SUMF2=SUMP2/FIC SUMF2=SUMP2/FIC SUMF2=SUMP2/FIC SUMF2=SUMP2/FIC SUMF2=SUMP2/FIC SUMF2=SUMP2/FIC SUMP2=SUMP2/FIC SUMP2=SUMP2/FIC SUMP2=SUMP2/FIC SUMP2=SUMP2/FIC SUMP2=SUMP2/FIC SUMP2=SUMP2/FIC SUMP2=SUMP2/FIC SU	20000000000000000000000000000000000000
123456789012345678901 111111111222	10 200	SUBROUTINE FPARM(BAND,IC) DIMENSION BAND(1) FIC=IC SUM1=0.0 SUM2=0.0 BMAX=0.0 BMIN=10.0 DO 10 I=1,IC SUM1=SUM1+BAND(I) SUM2=SUM2+BAND(I)*BAND(I) BMAX=AMAX1(BMAX,BAND(I)) BMIN=AMIN1(BMIN,BAND(I)) CONTINUE SUM1=SUM1/FIC SUM2=SUM2/FIC STD=SQRT(SUM2-SUM1**2) WRITE(6,200) BMAX,SUM1,BMIN,STD FORMAT(//53X,'MAX FREQ=',F5.2,IOX,'MEAN FREQ=',F5.2/53X, I'MIN FREQ=',F5.2,IOX,'STAND DEV=',F5.2) RETURN END	PAGE 1 FPAR 10 FPAR 20 FPAR 30 FPAR 30 FPAR 40 FPAR 60 FPAR 60 FPAR 60 FPAR 80 FPAR 120 FPAR 120 FPAR 120 FPAR 120 FPAR 150 FPAR 150 FPAR 150 FPAR 180 FPAR 200 FPAR 200

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• •		CURRENT REPORT OF THE THE THE ALL DUT CHR	0.7.4	
1:	c	SUBROUTINE PIKCK(PNORM,BANDF,DELFF,ILF,IHF,IC2,PL1,PH1,CMP)	PIKC	20
3:	č	************	PIRC	÷ 30
4: 5:	C C	THIS SUBROUTINE CHECKS FOR THE FREQUENCY PEAK IN EACH BAND	PIK	50
6:	ç		PIK	<u>60</u>
/ 1	č	***************************************	PIK	; <u>70</u>
9 :	U	COMPLEX CMP	PIRC	5 9ŏ
10:		QIMENSIQN PNORM(1),BANDF(1),PSPEC(200)	PIKC	100
11:			PIKC	110
13:	50	IFREBANDE(IC2)/DELEE+1	PIRC	5 130
14:	20	icc=0	PĪKČ	140
15:		ĮĘ(PNORM(IĘR)-PNORM(IFR+1)) 20,20,1	PIKO	: 150
19:	12	1 + (1 + R - 1) = 35 + 35 + 2	PIK	190
18:	4	IF (PNORM(IFR+1)=PNORM(IFR+2)) 20,20,4	PIK	5 180
19:	4		PÍK	i 190
20:		PH=(PNORM(IFR)-PNORM(IFR+2))/PNORM(IFR)	PIKC	200
21:	-	IF(PNORM(IFR+2)-PNORM(IFR+3))6,6,5	PIK	210
22	2	100=100+1 DH=(DNOPM(TEP)-DNOPM(TEP+3))/DNOPM(TEP)	PIKC	: 230
24:	6	IF(IFR-2) 35.35.7	PIR	240
25:	ž	IF(PNORM(IFR-1)-PNORM(IFR-2))20,20,8	PIK	250
26:	8		B i K	260
2/:		PL=(PNURM(1+R)-PNURM(1+R-2))/PNURM(1+R)	PIKC	2/0
29:	9	1 + (1 + 1 + 1) = 1 + (1 + 1) = 1 = 1 = 1 = 1 = 1 = 1 = 1 = 1 = 1	PIK	290
30:	ío		PIKO	5 3ÓŎ
31:		<u>PL=(PNORM(IFR)-PNORM(IFR-3))/PNORM(IFR)</u>	PIKC	: 310
32:	11	IF(1CC-3)20,29,29,29	PIK	320
34.	29		DIK	240
35	20	IF(IFLAG-1)21,22,22	PIK	5 350
36:	Žľ		PIK	360
37:		PSPEC(I)=PNORM(I)	PIK	370
38:	22		PIK	100
40:			PIK	400
41:		DO 24 J=IL, IHF	PIK	410
42:		IF(PMAX-PSPEC(J))23,24,24	PIK	420
43:	23	BANDF(1C2+1)=PSPEC(J)	PIKC	430
44:		BANDF([02]=(]=1]+DELFF DANY=DSDF([1]	PIK	440
46:	24		PIKO	460
47:	- ·	IFLAG=IFLAG+1	PIK	470
48:	20	IF(IELAG-(IHF-IL))50,50,35	PIKC	480
491	55	BANDF(IC2)=0	PIKU	490
51:	77	CMP = CMP = X (A A N D + (1 C 2) - BAND + (1 C 2 + 1))	PIK	510
52:	••	RETURN	PIK	520
53:		END	PIKC	; 530

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1::	200 210	SUBROUTINE WRTE(BAND) DIMENSION BAND(1) WRITE(6,200) FORMAT(//,58X,*PARAMETERS FOR THE BAND*//) WRITE(6,210) (BAND(1),I=493,500) FORMAT(30X,*MAX FREQ=*,F5.2,5X,*MEAN FREQ=*,F5.2,10X,*MAX PEAK=*, IF6.2,5X,*MEAN PEAK=*,F6.2,/30X,*MIN FREQ=*,F5.2,5X,*STAND DEV=*, 2F5.2,10X,*MIN PEAK=*,F6.4,5X,*STAND DEV=*,F6.2//) RETURN	WRTE WRTEE WRTTEE WRRTEE WRTTEE WRTEE WRTEE WRTEE	10 20 30 50 70 80
9: 10:		RÉTÜRN END	WRTE	90 100

1:	c	SUBROUTINE FMED(FFW,FMEAN,NB,IC)	FMED	10
3:	č	******	FMED	30
4: 5:	ç	THIS SUBROUTINE COMPUTES THE MEAN FREQUENCY IN EACH EPOCH	FMED	40 50
61 7:	C C	*****	FMED	60 70
8: 9:	C	DIMENSION FMEAN(1).FFW(1)	FMED	80 90
10:		FFW(5)=0.0 FFW(6)=0.0	FMED	100
12			FMED	120
14		NBT=2*NB	FMED	140
15:		SUMF=SUMF+FFW(J)*FFW(J+1)	FMED	150
17:	10	SUMW≓SUMW+FFW(J+1) CONTINUE	FMED	170 180
19:		ÉMEAŇ(IČ)=SUMF/SUMW RETURN	FMED	190
žĭ:		END	FMED	210

		PAGE	1
123456789	SUBROUTINE ARRAY(XF,Y,IC2) DIMENSION XF(1),Y(400,1) I=0 DD 10 J=1,IC2,2 I=I+1 Y(I,1)=XF(J) 10 CONTINUE RETURN END	ARRA ARRA ARRA ARRA ARRA ARRA ARRA ARR	10 20 30 40 500 700 80 90

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12345678901123456	10	SUBROUTINE SPARM(BAND,IC) DIMENS[UN BAND(1) FIC=IC SUM1=0.0 SUM2=0.0 BMAX=BAND(1) BMIN=BAND(1) BMIN=BAND(1) SUM2=SUM2+BAND(I) SUM2=SUM2+BAND(I) BMAX=AMAX1(BMAX,BAND(I)) BMIN=AMIN1(BMIN,BAND(I)) BMIN=AMIN1(BMIN,BAND(I)) CONTINUE SUM1=SUM1/FIC SUM2=SUM2/FIC SUM2=SUM2/FIC		SPAR 10 SPAR 20 SPAR 30 SPAR 40 SPAR 60 SPAR 60 SPAR 70 SPAR 80 SPAR 100 SPAR 100 SPAR 120 SPAR 140 SPAR 140 SPAR 160
15: 16: 17:	• • • •	SUM2=SUM2/FIC STD=SQRT(SUM2-SUM1**2) WRITE(6,200) BMAX,SUM1,BMIN,STD		SPAR 140 SPAR 150 SPAR 160 SPAR 170
18: 19: 20: 21:	200	FURMAT(//52X, MAX = ", F6.2, 10X, MEAN 1'MIN = ', F6.2, 10X, 'STAND DEV=', F6.2) RETURN END	=',F6.2/52X,	SPAR 180 SPAR 190 SPAR 200 SPAR 210

1: 2: 3: 4: 5: 6: 10: 11: 12: 10: 14: 15: 16: 17: 200 20: 21: 200	SUBROUFINE VOST(BAND,IC) DIMENSION BAND(1) FIC=IC SUM1=0.0 SUM2=0.0 BMIN=10.0 DO 10 I=1,IC SUM1=SUM1+BAND(I) SUM2=SUM2+BAND(I)*BAND(I) BMAX=AMAX1(BMAX,BAND(I)) BMAX=AMAX1(BMAX,BAND(I)) BMAX=AMAX1(BMAX,BAND(I)) CONTINUE SUM1=SUM1/FIC SUM2=SUM2/FIC STD=SQRT(SUM2-SUM1**2) WRITE(6,200) BMAX,SUM1,BMIN,STD FORMAT(//53X,'MAX PERC=',F5.2,10X,'MEAN PERC=',F5.2/53X, 1'MIN PERC=',F5.2,10X,'STAND DEV=',F5.2) RETURN END	VOST 1 VOST 2 VOST 3 VOST 3 VOST 6 VOST 6 VOST 6 VOST 1 VOST 9 VOST 11 VOST 11 VOST 12 VOST 12 VOST 14 VOST 15 VOST 16 VOST 20 VOST 20
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1: 2: 3: 4: 5: 6: 7: 8: 10 9: 10:	SUBROUTINE RAIZ (XF,Y,IC2) DIMENSION XF(1),Y(400,1) IC=IC2/2 I=0 DO 10 J=2,IC2,2 I=I+1 Y(I,1)=SQRT(XF(J)) CONTINUE RETURN END	RAIZ 10 RAIZ 20 RAIZ 30 RAIZ 40 RAIZ 50 RAIZ 60 RAIZ 70 RAIZ 80 RAIZ 90 RAIZ 100
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