# Treatment protocols at ECT

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# Treatment protocols past present future Scottish ECT Audit Network August 2000

#### This seminar will

- •give an overview of the development of treatment protocols at ECT
- •point to some of the evidence for present day guidelines
- •outline the questions still to be answered

# development of protocols

- ♦ 1939-69: high efficacy high side-effects
- ◆ 1975-85: low efficacy low side-effects
- ◆ 1985-00: search for theraputic supremacy

- •The first thirty years of ECT saw the use of high dose treatment regimes.
- •Ultra low dose treatments followed for the next ten years.
- •Over the last five years attempts have been made to find the formula for maximum efficacy and minimum side-effects.

## development of protocols (2)

#### 1939 - 69:

- sine wave electricity
- little attention to dose
- side-effects accepted or ignored

- •In 1939 ECT had a dramatic effect and was soon adopted as a mainstream treatment for depression.
- •Little attention was paid to the dose of electricity and cognitive side-effects were even thought desirable by some psychiatrists.
- •Anaesthesia for ECT was introduced in the 1950's and this made the whole process of treatment much safer.

#### development of protocols (3)

#### 1969 - 85:

- any seizure activity accepted
- re-introduction of pulsed wave forms
- doses of electricity low
- unilateral ECT
- College statement 1977

- •By 1970 there was a renewal of interest in ECT machines which could deliver brief pulses of electricity.
- •This together with the belief that unilateral ECT was as effective as bilateral ECT saw the introduction of treatment protocols which used very low doses of electricity.
- •A Royal College of Psychiatrist's statement in 1977 ratified the use of ECT as a treatment in psychiatry.

#### development of protocols (4)

#### 1985 - 89:

- importance of dose intensity
- ♦ 25 sec seizure as a guide
- ◆ College guidelines 1989

- •By the early 1980's clinicians had reverted back to using bilateral ECT because the 'new' unilateral treatment appeared ineffective.
- •It had become evident that more than just 'any seizure activity' was required for efficacy.
- •and that something about the *intensity* of the electrical dose was important.
- •The Royal College of Psychiatrist's guidelines of 1989 suggested that, as a guide, a seizure of at least 25 seconds should be observed.
- •Bilateral ECT was recommended as the treatment of choice.

intensity of electrical dose				
◆ Lambourne & Gill	1978	(L & G)*		
cf.				
Royal Edinburgh DB trial	1979	(RE)		
Northwick Park ECT trial	1980	(NWP)		
<ul><li>Sutton Hospital DB trial</li></ul>	1981	(SH)		
<ul><li>Leicestershire trial</li></ul>	1984	(L)		
		*see notes		
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- •Some evidence for the importance of 'intensity' of dose (or energy level) can be found in the series of double blind trials 1978-84.
- •All these studies looked at real vs. 'sham' ECT.
- •The L & G study used *unilateral*, *brief pulsed* ECT(very low energy) and showed *no difference* in outcome.
- •The NWP study used *bilateral*, *brief pulsed* (low energy) ECT and the difference was *equivocal*.
- •The RE, SH and L studies all used *bilateral*, *chopped sine* wave (high energy) ECT and there was a *significant difference* with real ECT.

# intensity of electrical dose (2)

A double blind controlled comparison of the theraputic effects of high and low dose energy electroconvulsive therapies. Robin and deTissera. BJPsych. 1982:141

- 1. low dose
- 2. high dose with pulsed energy
- 3. high dose with chopped sine wave

results:same seizure length

2 and 3 more efficacious; 3 more side-effects than 2

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- •In 1982 Robin and de Tissera looked at outcome using three different ECT machines.
- •There was no difference in terms of patient populations treated
- •There was no difference in terms of the length of seizure produced.
- •But there was a relationship between intensity of electrical dose and outcome ie

the higher the dose the better the antidepressant effect the higher the dose the more chance of side-effects.

#### development of protocols (5)

#### 1989 - 95:

- concept of seizure threshold (ST)
- variability in seizure threshold
- change in ST with treatment
- treatment to suit the individual
- ◆ College guidelines 1995

- •The seizure threshold is the minimum amount of electricity which will trigger a seizure in a given individual.
- •It is the amount by which a dose of electricity exceeds this seizure threshold that determines both the efficacy and side-effects of ECT.
- •Early work from the USA suggested a 40 fold variation in individual seizure thresholds. Research carried out more recently in GB concludes this variation is in the region of 4 10 fold
- •The patient seizure threshold can be
- i) measured or
- ii)estimated according to certain variables eg sex, age
- •The 1995 College guidelines gave examples of both methods.

## millenium treatment protocols

- bilateral ECT as treatment of choice
- ♦ dose 50-100% above seizure threshold (ST)
- measure seizure length as a guide to ST
- restimulate 'missed' seizures, higher dose
- terminate prolonged seizures

- •The College handbook is under review.
- •The present guidelines for the most cost-effective treatment appear in the above slide.
- •The next slide shows some of the evidence for the choice of bilateral ECT at moderately supra threshold doses.

## efficacy data for setting dose

Sackeim et al. (series of studies 1991 - 93, USA)

- ♦ low dose UECT 28% response
- ♦ low dose BECT 70% response
- same seizure length
- cognitive side-effects related to dose above seizure threshold rather than absolute dose

**conclusion**: best outcome when the dose exceeds seizure threshold by 50 - 100% for a given individual

- •In a series of studies Sackeim concluded that:
- •Low dose bilateral ECT was more effective than low dose unilateral ECT.
- •Low dose bilateral ECT was faster acting than high dose unilateral and less likely to end in relapse
- •Cognitive side-effects for bilateral ECT were related to the dose above seizure threshold rather than the absolute dose of electricity.
- •Outcome was best when the dose of electricity was 50 100% above seizure threshold.

#### seizure threshold - first ECT

• measure. pros: specific

theraputic, despite seizure length

decreased risk of overdose

cons: time under anaesthetic

risks of repeated stimulation?

• estimate. pros: quick

cons: variation from the mean in 1 in

20 so need clinical feedback

- •There are two ways to work out the seizure threshold
- i) the first is to measure it by starting off with a low dose of electricity and increase by small increments until a seizure is produced.
- ii) the second is to estimate the seizure threshold by using a table of variables often available from the machine manufacturer, which for example takes into account the patient's sex, age and concurrent medication.
- •The pros and cons of each method are summarised in the above slide.

## variations in seizure threshold

raised by: incr. age lowered by: female sex male sex low CO 2 dehydration some drugs low oxygen caffeine propofol unilateral electrodes propranolol benzodiaz.

bilateral electrodes

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- •This slide shows some of the factors which can affect the patient's seizure threshold.
- •Note that ECT itself is has anti-convulsant properties.

#### subsequent treatments

(seizure threshold measured)

- monitor length of seizure
- increase dose if fit length falls by 30-50%
- re-titrate after 6th ECT

- •The first ECT session measures the seizure threshold (ST).
- •At the second ECT session a theraputic treatment can be given (50-100% above ST).
- •The fact that the electrical dose has been measured means that less emphasis needs to be placed on the seizure length.
- •However the seizure length is still recorded as this gives an indication of any increase in seizure threshold as the course of treatment progresses.
- •Treatment protocols should respond to a significant fall in seizure length by increasing the dose of electricity to remain 50
- 100% above seizure threshold.
- •The seizure threshold can be measured again after say 6 ECT treatments if there is any doubt or concern.
- •Treatment protocols should take account of clinical feedback.

## subsequent treatments

(seizure threshold estimated)

- take account of clinical picture
- reduce dose if any cognitive side-effects
- increase dose if fit length falls by 30-50%
- increase dose if no improvement

- •An estimated treatment dose of electricity is given at the first ECT session.
- •It is vital that the clinical response is monitored because there may be a 1 in 20 variation from the mean in seizure threshold estimation.
- •A dose of electricity too high above seizure threshold may result in cognitive side effects.
- •A dose too low is likely to be ineffective.
- •Treatment protocols must therefore take account of clinical feedback especially if there has been any post treatment confusion.
- •Again the seizure length is recorded to give some indication of a change in seizure threshold as the course of treatment progresses.

initial seizure threshold			
first dose	1997	<u> 1999</u>	
of ECT	(n=36)	(n=35)	
	%	%	
measured	22	26	
estimated	56	63	
fixed	22	11	
(Scottish Audit of ECT 1997-00)  Scottish ECT Audit Network August 2000			

- •The next 3 slides show some results from the national audit of ECT in Scotland from 1997-2000.
- •By 1999 89% of ECT units had adopted the College recommendation that the first dose of ECT should not be 'fixed', ie the same for everyone.

## stimulus dosing protocols

#### **Scotland**

1994: 77%

1997: 89%

1999: 94%

2000: 100%

(E & W in 1996: 34%)

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•Stimulus dosing protocols allow the dose of ECT to be altered according to response, for example:

to take account of missed or partial seizures

to take account of prolonged seizures

to take account of the seizure length as the course of treatment progresses

to take account of clinical feedback.

## outcome in a clinical setting

*definite improvement* (MADRS/CGI)

diagnosis	1997	1999
depressive illness	72%	72%
schizophrenic illn.	66%	61%
manic illness	65%	68%

- •The national audit of ECT in Scotland looked at outcome in a clinical setting.
- •The Montgomery Asberg Depression Rating Scale (MADRS) was used to measure change in those patients being treated for a depressive illness.
- •The Clinical Global Impression (CGI) was used as an outcome measure for other diagnoses.
- •The audit standard was set at:

- •The above slide shows the percentage of patients reaching the audit standard.
- •So, in line with the Welsh audit 1996, this audit confirms that ECT is effective in a clinical setting.

<sup>&#</sup>x27;at least a 50% reduction in the MADRS in 70% or if more relevant a definite improvement in the CGI for the majority of patients'

# questions for 2001

- is seizure threshold measurement worthwhile in a routine clinical setting?
- is high dose unilateral ECT a better option?
- what can we gain from EEG monitoring?

- •Some questions remain unanswered.
- •There is still debate about whether it is worthwhile measurinig seizure threshold if dosing strategies are kept low and the clinical picture is monitored.
- •There is a renewal of interest in unilateral ECT given at very high doses (5 times seizure threshold). This may be as effective as low dose bilateral ECT without the risk of cognitive side-effects but results of recent research needs to be replicated. (see next slide for comparison)
- •And American ECT machines now available in GB offer EEG monitoring.

choice of electrode placement				
type of ECT	outcome side-effects			
1. low dose UECT	22%			
2. high dose UECT (2.5 x ST)	70% (high relapse)			
3. low dose BECT	70% II			
4. high dose BECT Sackeim et al. New Eng J of Medicine, 1993. 32	80% III 28:839-846			
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- •Early work looking at the effects of unilateral vs bilateral ECT used doses of electricity 2.5 times seizure threshold for unilateral ECT.
- •More recent studies from USA have used doses upto 5 times seizure threshold for unilateral ECT and early results are promising; there may be a relation between level above seizure threshold and efficacy.
- •If this is confirmed then very high dose unialeral ECT may become the treatment of choice because even at these levels cognitive side-effects may be less than with bilateral ECT regimes.
- •More recently treatment protocols in GB are becoming more flexible and allow for a choice in treatment modality eg. start with unilateral and switch to bilateral if no improvement or start with bilateral and switch to unilateral if cognitive side-effects are troublesome.

# **EEG** monitoring

- detection of prolonged seizures
- indication of efficacy??

- •Evidence in support of the routine use of EEG monitoring is being considered by the College.
- •Some studies suggest a 2-4% incidence of prolonged cerebral seizure activity not accompanied by prolonged motor activity.
- •Another has shown only a 60% correlation between cerebral seizure and peripheral motor activity.
- •Therefore from a safety angle there may be some justification in monitoring at least the first treatment as prolonged seizures are more likely at this stage.
- •As yet there have been no confirmed studies which have demonstrated a direct relationship between post ECT EEG changes and clinical outcome, despite the claims of some machine manufactures.

#### conclusion

ECT is a safe and effective treatment provided care is taken to fit the treatment to the patient

- •ECT is a safe and effective treatment provided care is taken to fit the treatment to the patient.
- •Much more is now known about the mechanisms of ECT and there is a lot of interesting research emerging which continues to inform the development of treatment protocols.
- •It is the responsibility of the ECT consultant to keep abreast of developments and provide adequate training and supervision for junior staff.