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Introduction

Computed tomography (CT) imaging is a method for visualising the structural organisation of the brain using the attenuation of X-rays to generate image contrast. Tissues in regions of interest are highlighted based on their X-ray absorption properties, as dense tissues attenuate X-rays more than soft tissues, and air attenuates the least. Three-dimensional images are generated from a series of two-dimension X-ray images taken around a single axis of rotation.

Method

We have included only systematic reviews (systematic literature search. detailed methodology with inclusion/exclusion criteria) published in full text, in English, from the year 2000 that report results separately for people with diagnosis of schizophrenia, а schizoaffective disorder, schizophreniform episode disorder or first schizophrenia. Reviews were identified by searching the databases MEDLINE, EMBASE, CINAHL, Current Contents, PsycINFO and the Cochrane library. Hand searching reference lists of identified reviews was also conducted. When multiple copies of reviews were found, only the most recent version was included. Reviews with pooled data are prioritised for inclusion.

Review reporting assessment was guided by the Preferred Reporting Items for Systematic Meta-Analyses Reviews and (PRISMA) checklist, which describes a preferred way to present a meta-analysis¹. Reviews rated as having less than 50% of items checked have been excluded from the library. The PRISMA flow diagram is a suggested way of providing information about studies included and excluded with reasons for exclusion. Where no flow diagram has been presented by individual reviews, but identified studies have been described in the text, reviews have been checked for this item. Note that early reviews may have been guided by less stringent reporting checklists than the PRISMA, and that some reviews may have been limited by journal guidelines.

Evidence was graded using the Grading of Recommendations Assessment, Development and Evaluation (GRADE) Working Group approach where high quality evidence such as that gained from randomised controlled trials (RCTs) may be downgraded to moderate or low if review and study quality is limited, if there is inconsistency in results, indirect comparisons, imprecise or sparse data and high probability of reporting bias. It may also be downgraded if risks associated with the intervention or other matter under review are high. Conversely, low quality evidence such as that gained from observational studies may be upgraded if effect sizes are large, there is a dose dependent response or if results are reasonably precise and direct with low consistent, associated risks (see end of table for an explanation of these terms)². The resulting table represents an objective summary of the available evidence, although the conclusions are solely the opinion of staff of NeuRA (Neuroscience Research Australia).

Results

We found two systematic reviews that met our inclusion criteria^{3, 4}.

- Moderate quality evidence suggests reduced temporal lobe volume in people with schizophrenia.
- Moderate to low quality evidence is unclear as to the utility of structural imaging as a means of identifying individual structural abnormalities in patients following a first episode of psychosis.

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Goulet K, Deschamps B, Evoy F, Trudel JF

Use of brain imaging (computed tomography and magnetic resonance imaging) in first-episode psychosis: review and retrospective study

Canadian Journal of Psychiatry - Revue Canadienne de Psychiatrie 2009; 54(7): 493-501

View review abstract online

Comparison	Classification of structural imaging scans in patients following a first episode of psychosis, into four groups based on the degree of structural abnormality observed: normal; abnormal with no clinical impact; abnormal with impact on management but unlikely to cause psychosis; abnormal with possible causal link to psychotic symptoms.
Summary of evidence	Moderate to low quality evidence (large sample, indirect, unable to assess precision or consistency) is unclear as to the usefulness of structural imaging as a means of identifying individual structural abnormalities in patients following a first episode of psychosis.
Individual structural abnormalities	
384 CT scans from five studies were collated in total.	
78 (20.8%) of these were rated as benign abnormalities with no clinical impact.	
4 (1%) of these were rated as unlikely to have causal links to psychosis.	
5 (1.3%) of these were rated as abnormal with possible causal links to psychotic symptoms including possible infarcts of caudate nucleus and parietal cortex; ischemic changes; and colloid and arachnoid cysts.	
Consistency in results	No measure of consistency reported, although results appear inconsistent.
Precision in results	No confidence intervals are reported.
Directness of results	Indirect assessment of structural abnormality.

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Zakzanis KK, Poulin P, Hansen KT, Jolic D		
Searching the schizophrenic brain for temporal lobe deficits: a systematic review and meta-analysis		
Psychological Medicine 2000; 30(3): 491-504		
View review abstract online		
Comparison	Temporal lobe volume in people with schizophrenia vs. healthy controls.	
Summary of evidence	Moderate to quality evidence (unclear sample size, direct, unable to assess precision and consistency) suggests a temporal lobe volume deficit in people with schizophrenia. Authors suggest the average magnitude of this deficit is not sufficient to attribute any causative role in schizophrenia aetiology.	
Bilateral whole temporal lobe volume		
Small effect of decreased temporal lobe volume in people with schizophrenia;		
2 observational studies, N unclear, $d = 0.49$, SD = 0.55		
Laterality of temporal lobe volume		
Left temporal lobe		
Small effect of decreased left temporal lobe volume in people with schizophrenia;		
3 observational studies, $N = 80$, $d = 0.30$, $SD = 0.21$		
Right temporal lobe		
Small effect of decreased right temporal lobe volume in people with schizophrenia;		
3 observational studies, N = 80, d = 0.26, SD = 0.14		
Consistency in results	No measure of consistency reported, although results appear inconsistent.	
Precision in results	No confidence intervals are reported.	
Directness of results	Direct	

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Explanation of acronyms

CI = confidence interval, CT = computed tomography, d = Cohen's d and g = Hedges' g = standardised mean differences (see below for interpretation of effect sizes), F= ratio of between sample variance and within sample variance, N = number of participants, p = statistical probability of obtaining that result (p < 0.05 generally regarded as significant), SD = standard deviation, vs. = versus

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Explanation of technical terms

- * Bias has the potential to affect reviews of both RCT and observational studies. Forms of bias include; reporting bias - selective reporting of results, publication bias - trials that are not formally published tend to show less effect than published trials, further if there are statistically significant differences between groups in a trial, these trial results tend to get published before those of trials without significant differences; language bias - only including English language reports; funding bias - source of funding for the primary research with selective reporting of results within primary studies; outcome variable selection bias: database bias including reports from some databases and not others; citation bias - preferential citation of authors. Trials can also be subject to bias when evaluators are not blind to treatment condition and selection bias of participants if trial samples are small⁵.
- † Different effect measures are reported by different reviews.

Weighted mean difference scores refer to mean differences between treatment and comparison groups after treatment (or occasionally pre to post treatment) and in a randomised trial there is an assumption that both groups are comparable on this measure prior to treatment. Standardised mean differences are divided by the pooled standard deviation (or the standard deviation of one group when groups are homogenous) which allows results from different scales to be combined and compared. Each study's mean difference is then given a weighting depending on the size of the sample and the variability in the data. 0.2 represents a small effect, 0.5 a medium effect, and 0.8 and over represents a large effect⁵.

Reliability and validity refers to how accurate the instrument is. Sensitivity is the proportion of actual positives that are correctly identified (100% sensitivity = correct identification of all actual positives) and specificity is the proportion of negatives that are correctly identified (100% specificity = not identifying anyone as positive if they are truly not).

Odds ratio (OR) or relative risk (RR) refers to the probability of a reduction (< 1) or an increase (> 1) in a particular outcome in a treatment group, or a group exposed to a risk factor, relative to the comparison group. For example, a RR of 0.75 translates to a reduction in risk of an outcome of 25% relative to those not receiving the treatment or not exposed to the risk factor. Conversely, a RR of 1.25 translates to an increased risk of 25% relative to those not receiving treatment or not having been exposed to a risk factor. A RR or OR of 1.00 means there is no difference between groups. A medium effect is considered if RR > 2 or < 0.5 and a large effect if RR > 5 or < 0.2^{6} . InOR stands for logarithmic OR where a InOR of 0 shows no difference between groups. Hazard ratios measure the effect of an explanatory variable on the hazard or risk of an event.

Correlation coefficients (eg, r) indicate the strength of association or relationship between variables. They are an indication of prediction, but do not confirm causality due to possible and often unforseen confounding variables. An r of 0.10 represents a weak association, 0.25 a medium association and 0.40 and over represents а strong association. Unstandardised (b) regression coefficients indicate the average change in the dependent variable associated with a 1 unit change in the independent variable, statistically controlling for the other independent variables. Standardised regression coefficients represent the change being in units of standard deviations to allow comparison across different scales. Reliability

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and validity refers to how accurate the instrument is. Sensitivity is the proportion of actual positives that are correctly identified (100% sensitivity = correct identification of all actual positives) and specificity is the proportion of negatives that are correctly identified (100% specificity = not identifying anyone as positive if they are truly not).

Prevalence refers to how many existing cases there are at a particular point in time. Incidence refers to how many new cases there are per population in a specified time period. Incidence is usually reported as the number of new cases per 100,000 people per year. Alternatively some studies present the number of new cases that have accumulated over several years against a person-years denominator. This denominator is the sum of individual units of time that the persons in the population are at risk of becoming a case. It takes into account the size of the underlying population sample and its age structure over the duration of observation.

‡ Inconsistency refers to differing estimates of treatment effect across studies (i.e. heterogeneity or variability in results) that is not explained by subgroup analyses and therefore reduces confidence in the effect estimate. I² is the percentage of the variability in effect estimates that is due to heterogeneity rather than sampling error (chance) - 0% to 40%: heterogeneity might not be important, 30% to 60%: may represent moderate heterogeneity, 50% to 90%: may represent substantial heterogeneity and 75% to 100%: considerable heterogeneity.

§ Imprecision refers to wide confidence intervals indicating a lack of confidence in the effect estimate. Based on GRADE



recommendations, a result for continuous data (standardised mean differences, not weighted mean differences) is considered imprecise if the upper or lower confidence limit crosses an effect size of 0.5 in either direction, and for binary and correlation data, an effect size of 0.25. GRADE also recommends downgrading the evidence when sample size is smaller than 300 (for binary data) and 400 (for continuous data), although for some topics, this criteria should be relaxed⁷.

Indirectness of comparison occurs when a comparison of intervention A versus B is not available but A was compared with C and B was compared with C, which allows indirect comparisons of the magnitude of effect of A versus B. Indirectness of population, comparator and or outcome can also occur when the available evidence regarding a particular population. intervention, comparator, or outcome is not available so is inferred from available evidence. These inferred treatment effect sizes are of lower quality than those gained from head-to-head comparisons of A and B.

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