

## Background

- Activa® DBS devices (Medtronic Inc, Minneapolis, MN, USA) deliver electrical stimulation in the constant voltage (CV) mode, but allow switching to a constant current (CC) mode.
- No evidence-based guidelines exist to choose one delivery method over another.
- CC stimulation devices regulate voltage to deliver a programmed current, which may be advantageous initially after DBS implantation, when tissue impedance is known to fluctuate.
- Impedance (R) is also expected to change in response to stimulation strength:  $V=IR$  (I – current; V – voltage).
- The extent and clinical significance of impedance changes in vivo over long-term DBS treatment are not well characterized.
- Objectives:
  - 1) To follow therapy impedance changes over time in patients with CV DBS;
  - 2) To determine if a correlation exists between changes in therapy impedance and changes in DBS parameters

## Methods

- A retrospective chart review (1/2010 – 4/2014) identified Parkinson's disease patients with Activa DBS pulse generators (PC, SC) and 3387 leads programmed in CV mode, with documented therapy impedance over at least 3 consecutive visits after the last DBS-related surgery (initial implant, battery exchange).
- Exclusion criteria:
  - Change in electrode polarity
  - Abnormal impedance (therapy or electrode) reading consistent with possible open or short circuit;
- All impedance values were documented at the beginning of each study visit before making any changes in DBS parameters.
- The magnitude of stimulation was calculated as Charge Density (CD):  $CD = (\text{Volts} / \text{Impedance}) * (\text{pulse width}) / \text{surface area}$
- (surface area is assumed unchanged in the same patients / same electrode configuration) [1].
- Changes of CD and impedance between visits, and compared to baseline (first study visit) were calculated in each patient.

## Results

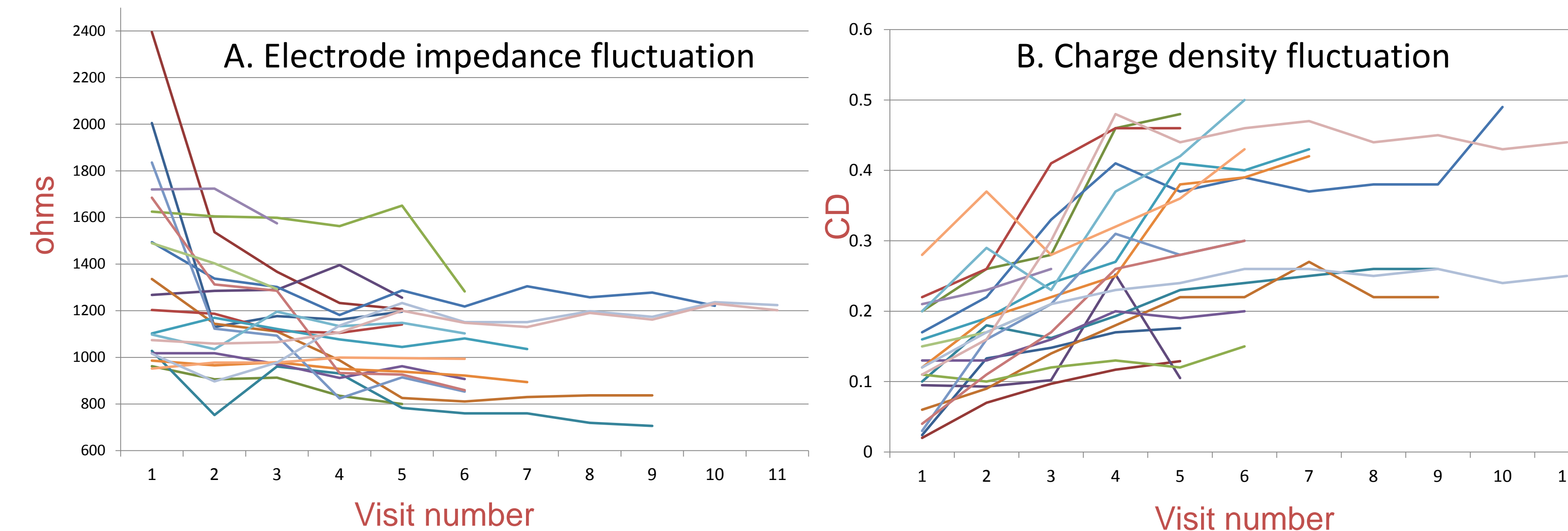
**Table 1. Patient and DBS Characteristics**

|  |   |
|--|---|
| Patients / electrodes, N                                       | 10/20   |
| Gender   | Male (n=6), Female (n=4)                                |
| DBS Device   | Activa PC (n=10)  |
| Target   | b/l STN (N=4), b/l GPi (N=6)                            |
| Mean time to 1 <sup>st</sup> visit with impedance check (days) | 158.8 +/-233.1 (all initial DBS implants)               |
| Mean # follow-up visits/electrode                              | 6.55 +/-2.3   |
| Mean duration of follow-up/electrode                           | 467.65 +/-156.2   |
| Electrode configuration  | Bipolar (n=4), double monopolar (n=1), monopolar (n=15) |

STN – subthalamic nucleus; GPi – globus pallidus interna

## Results, continued

**Figure 1. Individual electrode impedance (A) and charge density (B) fluctuation (absolute value) over time**



\*Each colored line represents an individual electrode.  
Impedance generally varied inversely in relation to CD changes

**Figure 2A,B. Correlation between impedance change, CD change and time since surgery**

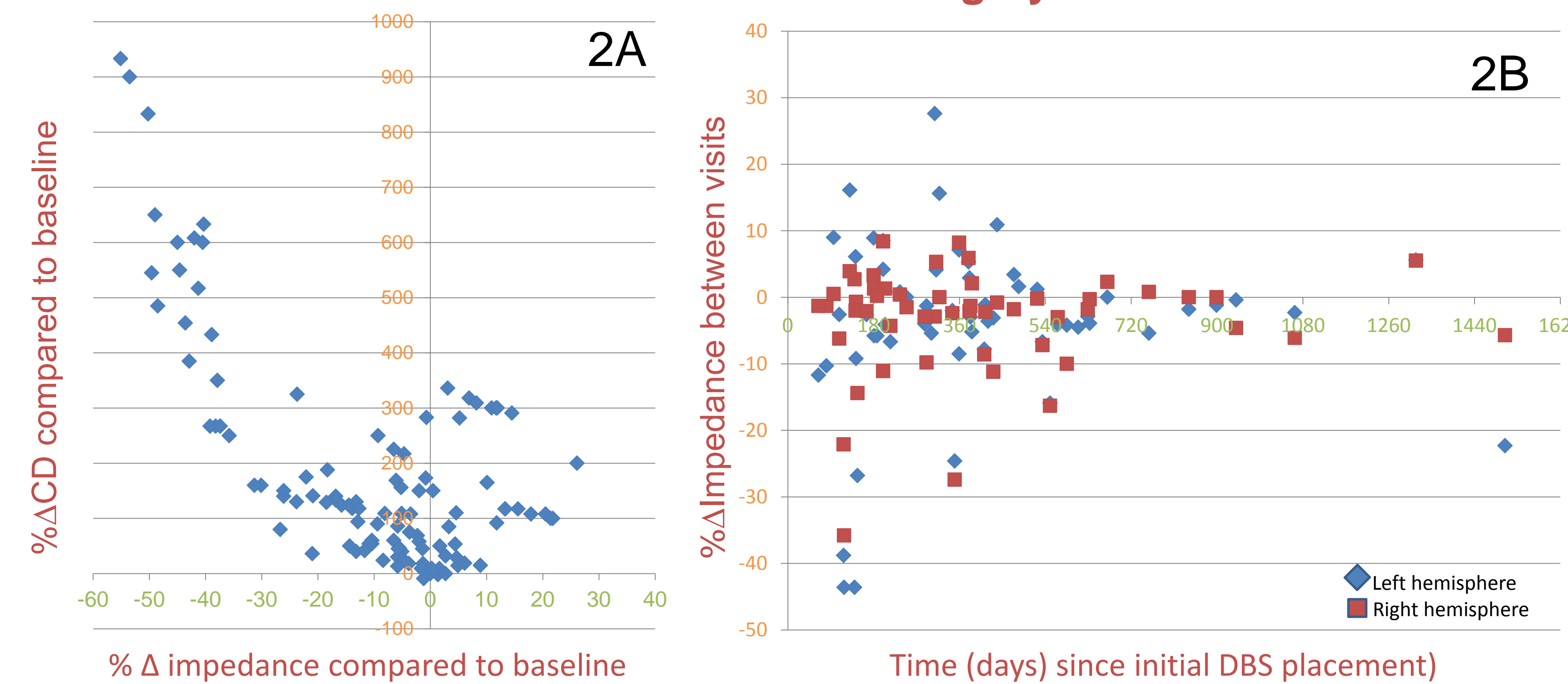


Figure 2A shows change of impedance vs change of CD for all patient-visits (n=111) and all electrodes

Figure 2B shows impedance change between visits vs time since the DBS implantation surgery

**Table 2. Impedance change from baseline vs. change between visits**

|  |               |
|--|---------------|
| Mean %change impedance from baseline in all electrodes over all visits | -11.04 ±19.37 |
| Mean %change impedance between visits                                  | -3.5 ± 10.7   |
| 0-180days  | -8.6 ± 16.7   |
| 181-360days  | -1.1 ± 9.8    |
| >360days   | -2.6 ± 5.8    |

Despite large changes in CD, impedance fluctuations between visits or last visit-vs-baseline were relatively small.

## Discussion

- Stimulation current influences Volume of Tissue Activated (VTA) - activated neuronal elements around active contact(s) producing clinical benefits and adverse effects of DBS treatment.
- In CV stimulation, the actual current (I) delivered to brain tissue may vary if there are significant fluctuations of impedance (R) based on the formula:  $I = \frac{V}{R}$  (V – voltage)
- In CC stimulation, current does not change but the voltage varies in order to maintain the same current in response to impedance fluctuations.
- A previous study of 63 patients with older Medtronic DBS devices (Kinetra and Soletra) did not identify any significant intra-patient impedance fluctuation over 2 visits with unchanged DBS parameters with CV stimulation devices [2].
- On the other hand, impedance can change significantly during first 3 months after electrode implant due to tissue healing [3]. This observation could theoretically justify the advantages of CC over CV stimulation during the early post-implantation period.
- We analyzed serial impedance readings in response to real-world clinical application of DBS (in the absence of changes in polarity) up to 1504 days after electrode implantation.
- Our findings suggest the following about the degree and nature of impedance changes in chronic DBS therapy:
  - Impedance measurements are generally reduced over time, while CD increases with chronic stimulation.
  - Impedance fluctuations are small relative to changes in CD at any stage of DBS treatment, and vary inversely.
  - Impedance fluctuations are greatest in the initial post-implantation period (<6mos):  $-8.6\% \pm 16.7\%$  (range -44 + 16).
  - In the chronic phase of DBS therapy, impedance fluctuations are relatively small, though they may still occur:
    - $-1.1\% \pm 9.8$  (range -27 + 28) for 6-12 months
    - $-2.6 \pm 5.8$  (range -22 to +11) for >12 months

## Conclusions

- Impedance fluctuations in chronic DBS therapy delivered in CV stimulation mode are minimal over time, and generally decrease, in a real world clinical setting.
- In CV mode, impedance appears to decrease proportionately to increases in CD, as can be expected according to  $V=IR$
- The results of our study do not support the use of CC DBS delivery over CV in order to accommodate for fluctuating impedance in chronic DBS therapy.

## References

- [1] Marks W. Deep Brain Stimulation Management. Cambridge University Press 2011, New York; p 37.
- [2] Sillay KA, Chen JC, Montgomery EB. Long-term measurement of therapeutic electrode impedance in deep brain stimulation. Neuromodulation 10;13:195-200.
- [3] Shils JL, Alterman RL, Arle JE. Deep brain stimulation fault testing. In: Deep brain stimulation in neurological and psychiatric disorders. Tarsy et al (Eds) 2008: pp 473-494.