

1 **Cerebral-localized low density area on post-mortem computed**
2 **tomography images: retrospective analysis of the forensic**
3 **autopsy cases.**

4
5
6 Kanto Ishii^{1*}, Shinobu Yamauchi¹, Akira Kurosu¹.

7
8 ¹Department of Legal Medicine, Dokkyo Medical University, 880 Kitakobayashi, Mibu,
9 Shimotsuga, Tochigi 321-0293, Japan

10
11 Tel: +81-282-87-2135; Fax: +81-282-86-7678; E-mail: kanto915@dokkyomed.ac.jp

12
13 *Corresponding author

1 **SUMMARY**

2

3 Post-mortem computed tomography (PMCT) can noninvasively visualize

4 abnormal organic findings, and is also often applied in the investigation of the cause of

5 death. We have experienced autopsy cases in which we observed abnormal findings on

6 PMCT images unrelated to the direct cause of death such as cerebral-localized

7 low-density area (CLLDA) without having a history of intracranial disease before death.

8 In this study, we investigated the characteristics of CLLDA in deceased cases. We

9 retrospectively reviewed the autopsy records and police reports provided for 1248

10 forensic autopsy cases performed at the Dokkyo Medical University School of Medicine,

11 between 1 February 2011 and 28 February 2014. We found 58 autopsy cases that were

12 over 20 years old and had been performed PMCT. Subjects having a history of

13 intracranial disease before death were excluded.

14 We divided autopsy cases into two groups, one for those with confirmed

15 CLLDA on PMCT images (CLLDA group) and the other for those with no confirmed

16 CLLDA on PMCT images (Control group). The variables obtained from the autopsy

17 records and police reports for each group, and were compared and statistically analyzed.

1 The prevalence of basilar artery sclerosis and coronary artery stenosis was
2 higher in the CLLDA group (p=0.022 and p=0.005 respectively). There was no
3 significant difference in left and right ventricular thickness and cardiac weight between
4 the two groups. It was suggested that the finding of CLLDA indicated the probable
5 presence of vascular abnormalities in forensic autopsy.

6
7

8 Key words: forensic autopsy; post-mortem computed tomography; vascular
9 abnormality.

10
11
12
13
14
15
16
17
18

1 Introduction

2

3 Post-mortem imaging can noninvasively visualize abnormal organic findings,
4 and is also often applied in the investigation of the cause of death. There are several
5 post-mortem imaging techniques currently available, such as post-mortem computed
6 tomography (PMCT), and post-mortem magnetic resonance imaging (PMMRI). PMCT
7 is one of the most prevalent techniques used for several reasons: PMCT was reported to
8 more accurately help identify the cause of death than PMMRI¹⁾; PMCT can be carried
9 out more quickly and cheaply than PMMRI¹⁾. PMCT has attracted attention in the fields
10 of pathology, forensic medicine, clinical medicine, and so far, several reported studies
11 have used this tool to investigate the cause of death. It was found that determining cause
12 of death in certain traumatic cases has been done equally accurately with both PMCT
13 and standard autopsy²⁾. Observations indicating fatal bleeding disorders, such as
14 intracranial vascular lesions and intra-thoracic hemorrhage, have reportedly been
15 confirmed by PMCT³⁾. In addition, PMCT helps ensure the safety of the autopsy
16 operator, since they confirm the presence of internal injury or its by-products, an
17 inferior vena cava filter or weapon debris using PMCT. However, few previous studies
18 focus on abnormal findings on PMCT images unrelated to the direct cause of death.

1 We have observed in autopsy cases a cerebral localized low-density area
2 (CLLDA) on PMCT images without having a history of intracranial disease before
3 death. However, there has been little reported about CLLDA on PMCT images at
4 forensic autopsy. In this study, we investigate the characteristics of CLLDA using the
5 autopsy findings, police records, and PMCT images in forensic autopsy.

6

7 **Materials and Methods**

8

9 The Ethics Committee of the Dokkyo Medical University approved this study
10 (Approval number was dmu27004). All data of the autopsy cases were strictly archived
11 in the department database based on autopsy code number. The autopsy records,
12 provided police reports, and PMCT images were perused. In this study, the provided
13 types of post-mortem images were PMCT and post-mortem X-ray. PMCT was
14 performed in all cases from the head to the lower limbs. In this study, since post-mortem
15 X-ray cannot evaluate the cerebral parenchyma, it was excluded. The necessary data
16 were collected and analyzed anonymously without consent due to the retrospective
17 nature of this study.

1 When the cause of death is unclear from the medical history, circumstances of
2 death, post-mortem imaging or external observations, a forensic autopsy is required to
3 investigate the cause of death. Post-mortem imaging is performed at the neighboring
4 hospital as required by the police or the doctor after confirming death. Because we have
5 not CT instrument, we used PMCT images provided by the police as the reference
6 material.

7 We compared CLLDA on PMCT images with anatomical and pathological
8 findings at the same site. On the other hand, because the imaging conditions such as the
9 slice thickness is different in each autopsy, we analyzed the slice thickness.

10

11 **Study Subjects**

12

13 We retrospectively reviewed the autopsy records and related police reports of
14 1248 forensic autopsy cases from the Dokkyo Medical University School of Medicine,
15 recorded between 1 February 2011 and 28 February 2014. We selected 58 autopsy cases
16 for which the deceased were over 20 years old and performed PMCT. The exclusion
17 criteria were as follows: (1) severe cranial injury; (2) severe cranial air lesions; (3)
18 advanced post-mortem change; (4) neurological abnormalities such as paralysis being

1 subjectively or objectively recognized before death; and (5) history of intracranial
2 disease.

3

4 **Data evaluation**

5

6 PMCT images were interpreted before reviewing the details of the autopsy and
7 police records. Lesions of more than 5 mm in diameter were considered as CLLDA.

8 Detection of CLLDA was performed in the cerebrum, cerebellum and brain stem. We
9 excluded air lesions by the computed tomography (CT) number.

10 We divided autopsy cases into two groups, one for those with confirmed
11 CLLDA on PMCT images (CLLDA group) and the other for those with no confirmed
12 CLLDA on PMCT images (Control group). The following variables were obtained from
13 autopsy records and police reports, and the two groups were compared for each
14 variable.

15 (1) Background factors: We examined the physical features at the time of death,
16 including sex, height, weight, body mass index (BMI), age and resuscitation attempts.
17 We also examined the slice thickness which may affect the detection rate of CLLDA
18 used for PMCT.

1 (2) Autopsy findings: whether the cause of death was classified as a disease or external
2 events. We examined autopsy findings related to the heart and brain as follows:
3 coronary artery stenosis; basilar artery sclerosis; left and right ventricular thickness;
4 cardiac weight; and presence of intramyocardial white scar. Coronary artery stenosis
5 was defined as significant stenosis (50% or more stenosis in actual measurement) if
6 observed in any of the left main trunk, left anterior descending coronary artery, left
7 circumflex coronary artery or right coronary artery. We classified basilar artery sclerosis
8 as no sclerosis, mild sclerosis, moderate sclerosis or severe sclerosis, with the last two
9 categories being classed as significant sclerosis.

10 Imaging conditions for PMCT (slice thickness), cause of death (disease or external
11 events), and resuscitation attempts may affect the interpretation of PMCT. We therefore
12 included a statistical evaluation of the differences in these variables between the two
13 groups in this study.

14

15 **Statistical Analysis**

16

17 Categorical data were analyzed using Fisher's exact test. Continuous data were
18 assessing the equality of variances for a variable using Levene's test. Subsequently, the

1 data found to equal variance were analyzed using Student's t-test, or the data found not
2 to equal variance were analyzed using Welch's t test. Data were expressed as a mean \pm
3 standard deviation (minimum - maximum). A p-value <0.05 was considered statistically
4 significant. All statistical analyses were performed with IBM SPSS software version
5 21.0 for Windows (IBM Japan, Ltd., Tokyo, Japan).

6

7 **Results**

8

9 **General aspects**

10

11 The 58 cases had a mean age of 57.8 ± 18 years. The mean height was $162.8 \pm$
12 10 cm, mean weight was 61.6 ± 16 kg, and mean BMI was 23.0 ± 5 kg/m². Forty-three
13 cases (74.1 %) were male. The mean of the slice thickness of PMCT was 5.4 ± 2
14 (2.0-10.0) mm with the exclusion of one case for which the thickness was not
15 documented. Thirty-four cases (58.6 %) were diagnosed in forensic autopsy as death of
16 disease, and 44 cases (75.9 %) included resuscitation attempts. Twenty-six cases
17 (44.8 %) showed coronary artery stenosis, and six cases (10.3 %) had an
18 intramyocardial white scar. The mean left ventricular thickness was 13.6 ± 2 mm, mean

1 right ventricular thickness was 3.5 ± 1 mm, and mean cardiac weight was 375.5 ± 89 g.
2 Eighteen cases (31.0 %) exhibited significant basilar artery sclerosis. Only one autopsy
3 case of 58 cases was found in which CLLDA on PMCT images was pathologically
4 observed as cavitation and gliosis at the same site.

5 The CLLDA group was made up of 14 cases in total, and the Control group
6 included the other 44 cases. Representative PMCT images from cases in the CLLDA
7 group are shown in Fig 1.

8 **Fig 1.挿入箇所**

9

10 **Detailed comparison of the CLLDA and Control groups**

11

12 We examined whether there was bias in the slice thickness of PMCT, cause of
13 death, and resuscitation attempts found in the two groups (Table 1). Post-mortem
14 changes may have hypostasis effects and resuscitation attempts may cause traumatic
15 changes on the interpretation of PMCT images. Additionally, the slice thickness of
16 PMCT may affect the detection of CLLDA. We found no significant differences for
17 these factors. Therefore, we continued with a detailed comparison of other variables for
18 the two groups.

1 **Table1.挿入箇所**

2 The comparison of physical features and autopsy findings is summarized in
3 Table 2. The prevalence of mean weight and mean BMI were higher in the CLLDA
4 group. There were no significant differences in the mean height for the two groups.

5 **Table2.挿入箇所**

6 Basilar artery sclerosis and coronary artery stenosis were also more common
7 in the CLLDA group than the Control group, but there was no significant difference in
8 the prevalence of intramyocardial white scar but there were no significant differences in
9 the mean right and left ventricular thicknesses and cardiac weights of the two groups.
10 The number and size of CLLDA in each subject were unrelated to basilar artery
11 sclerosis and coronary artery stenosis.

12 Since there was significant difference in the mean age between two groups, we
13 examined basilar artery sclerosis and coronary artery stenosis in elderly subjects (over
14 60 years old) (Table 3). The prevalence of coronary artery stenosis was observed a
15 significant difference. There was no significant difference in the prevalence of basilar
16 artery sclerosis, however, it showed a higher prevalence in the CLLDA group than the
17 Control group (70.0 % vs. 41.2 %).

18 **Table3. 挿入箇所**

1

2 **Discussion**

3

4 In this study, it was suggested that the finding of CLLDA indicated the
5 probable presence of vascular abnormalities such as basilar artery sclerosis and
6 coronary artery stenosis in forensic autopsy, however, relation of the presence of
7 intramyocardial white scar was not admitted. Since the facility is limited to perform
8 PMCT using contrast medium⁴⁾, in cases where contrast medium cannot be used in
9 PMCT, CLLDA may suggest the need for a detailed anatomical investigation about the
10 vascular abnormalities. PMCT is a useful complement to autopsy by the detection of
11 lesions prior to autopsy and lead to improve the anatomical accuracy. Understanding
12 CLLDA on PMCT images and history before death in advance suggests the regions that
13 need forensic anatomically detailed investigation such as coronary artery and basilar
14 artery.

15 PMCT is affected by post-mortem changes from immediately after cardiac
16 arrest. Circulatory arrest, starting from the time of death, results in overall brain
17 ischemia. The contrast between white matter and gray matter decreases over time and
18 gradually obfuscates the visible cerebral sulcus^{5,6)}. The circulation of blood also stops,

1 and the force of gravity causes post-mortem hypostasis throughout the body. This is
2 observed as a high-density area on PMCT images⁷⁾. It was reported that even if the
3 deceased is diagnosed as subarachnoid hemorrhage from the findings on PMCT images,
4 false-positive findings are caused by increasing density from hypostasis in the
5 intracranial sinus and dural hemorrhage caused by hypostasis in the tentorium vessels
6 and subsequent bursting^{8,9)}. The formations of CLLDA exclude air lesions as
7 post-mortem changes. Air lesions have been often observed on PMCT images, caused
8 by resuscitation, putrefaction gas by advanced post-mortem change, and skull base
9 fractures. These lesions can be differentiated from cerebrospinal fluid on PMCT images
10 using the difference in CT number.

11 CLLDA is considered to be the imaging findings, such as expanding
12 perivascular space, brain cyst and cerebral infarction, and magnetic resonance
13 imaging helps differentiate them¹⁰⁾. Liebetrau et al reported that detection rate of
14 CLLDA on ante-mortem CT images was 8.9 % in a study of 239 subjects of the
15 85-year-old living in Sweden¹¹⁾, although there has been little research of CLLDA
16 using PMCT. Shinkawa et al reported that the prevalence of asymptomatic cerebral
17 infarction, which is one of the pathogenesis indicating CLLDA, was 12.9 % in a
18 study of 996 autopsy cases in Japan¹²⁾. Detection rate of CLLDA in our study was

1 24.1 % and was higher than the results of these study, because the slice thickness in
2 the study of using ante-mortem CT was thicker as compared with our study (10 mm
3 vs about 5 mm), and the study of autopsy did not include expanding perivascular
4 space and brain cyst.

5 In this study, only one autopsy case of 58 cases was found in which abnormal
6 macroscopic and pathologic findings consistent with CLLDA was observed. There was
7 a possibility to be overlooked in cases where cross-section of the lesion was not
8 exposed at autopsy. As other factor, there was a possibility that CLLDA had not been
9 recorded as abnormal finding in case where this was confirmed as perivascular space in
10 forensic autopsy. However, if PMCT is performed correctly and interpreted under
11 appropriate conditions for the study, it can be valuable in detecting micro lesions.

12 This study had some limitations. We could not diagnose the cause of CLLDA
13 excluding one subject. Because of this study targeting the deceased cases, detailed
14 living conditions such as arteriosclerosis factor while alive were unknown. Further
15 studies are required to confirm diagnosis of detailed CLLDA by PMCT and anatomical
16 and pathological examinations.

17 In conclusion, this study was the first investigation that suggested an
18 association between CLLDA on PMCT images and coronary artery stenosis, basilar

1 artery sclerosis in forensic autopsy. PMCT is a useful complement to autopsy by the
2 detection of lesions prior to autopsy and lead to improve the anatomical accuracy.

3

4 **Acknowledgements**

5

6 This work was supported by a grant-in-aid from Dokkyo Medical University,
7 Young Investigator Award (Grant No. 2015-12)

8

9 **References**

10

11 1) Roberts IS, Benamore RE, Benbow EW, et al. Post-mortem imaging as an
12 alternative to autopsy in the diagnosis of adult deaths: a validation study. *Lancet*
13 379:136-142, 2012.

14

15 2) Le Blanc-Louvry I, Thureau S, Duval C, et al. Post-mortem computed tomography
16 compared to forensic autopsy findings: a French experience. *Eur Radiol*
17 23:1829-1835, 2013.

18

- 1 3) Takahashi N, Higuchi T, Shiotani M, et al. The effectiveness of
2 post-mortem multidetector computed tomography in the detection of fatal findings
3 related to cause of non-traumatic death in the emergency department. *Eur Radiol*
4 22:152-160, 2012.
- 5
- 6 4) Sakamoto N: *Zouei shigo CT. Autopsy imaging guideline third edition*, Vectorcore,
7 Tokyo, pp147-151, 2015.(in Japanese)
- 8
- 9 5) Ishida M, Gonoï W, Okuma H, et al. Common post-mortem computed tomography
10 findings following atraumatic death: differentiation between normal post-mortem
11 changes and pathologic lesions. *Korean J Radiol* 16:798-809, 2015.
- 12
- 13 6) Shirota G, Gonoï W, Ishida M, et al. Brain swelling and loss of gray and white
14 matter differentiation in human post-mortem cases by computed tomography. *PLoS*
15 *One* 10:e0143848, 2015.
- 16
- 17 7) Shiotani S, Kohno M, Ohashi N, et al. Post-mortem intravascular high-density fluid
18 level (hypostasis): CT findings. *J Comput Assist Tomogr* 26:892-893, 2002.

1

2 8) Kibayashi K, Shojo H, Sumida T. Dural hemorrhage of the tentorium on
3 post-mortem cranial computed tomographic scans in children. *Forensic Sci Int*
4 154:206-209, 2005.

5

6 9) Takahashi N, Satou C, Higuchi T, et al. Quantitative analysis of intracranial
7 hypostasis: comparison of early postmortem and antemortem CT findings. *AJR Am*
8 *J Roentgenol* 195:388-393, 2010.

9

10 10) Kwee RM, Kwee TC. Virchow-Robin spaces at MR imaging. *Radiographics*
11 27:1071-1086, 2007.

12

13 11) Liebetrau M, Steen B, Hamann GF, et al. Silent and symptomatic infarcts on cranial
14 computerized tomography in relation to dementia and mortality: a population-based
15 study in 85-year-old subjects. *Stroke* 35:1816-1820, 2004.

16

1 12) Shinkawa A, Ueda K, Kiyohara Y, et al. Silent cerebral infarction in a
2 community-based autopsy series in Japan. The Hisayama Study. Stroke 26:380-385,
3 1995.