Radionuclide venography of lower limbs by subcutaneous injection: A clinical evaluation

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SC-RNV, radionuclide venography by subcutaneous injection of Tc-99m pertechnetate at acupuncture points K-3, a new alternative of lower limb venography, was recently developed in our clinical laboratory. In some of the previous studies, we have proved its superiority to radionuclide venography by intravenous injection. The current investigation was conducted to understand the reliability of SC-RNV in the diagnosis of deep vein thrombosis (DVT). Fifty-seven cases with lower leg edema, from Nov., 1989 through Oct., 1990, received both SC-RNV and duplex US for causative evaluation. As a result of duplex US, 26 were considered normal (non-DVT), 19 were classified as unilateral DVT, and 12 as bilateral DVT. In nineteen cases (61%, 19/31) with DVT also a XCT and/or a CV (contrast venography) was taken, that showed compatible results. All of the non-DVT had a normal pattern of SC-RNV, all of the unilateral DVT had unilateral impairment of deep vein drainage in SC-RNV, and all of the bilateral DVT had impaired deep venous drainage bilaterally in SC-RNV. It is therefore concluded that SC-RNV is one of the best choices among available non-invasive lower-limb venographic methods.

Key words: radionuclide venography, subcutaneous injection, acupuncture points

INTRODUCTION

Radionuclide venography of the lower limbs by subcutaneous injection of Tc-99m pertechnetate at acupuncture points (SC-RNV), an alternative type of lower-limb venography, was recently developed in our clinical laboratory. It was recognized to be easier and simpler, yet more accurate than radionuclide venography by intravenous injection. In the present study, we intend to evaluate the reliability of the SC-RNV in the diagnosis of deep vein thrombosis of the lower limbs (DVT).

MATERIALS AND METHODS

From Nov., 1989 through Oct., 1990, 57 cases with lower leg edema were referred to us for evaluation of the cause of the leg edema. A SC-RNV and a color duplex ultrasound (duplex US) were carried out for every one of them on the same day.

Radionuclide venography by subcutaneous injection (SC-RNV)

A Toshiba GCA-90B Digital Gamma Camera was used for the SC-RNV. The SC-RNV technique has already been described in the literature, and in the present study the procedure was modified. After subcutaneous (SC) injection of 18.5–37 MBq (0.5–1 mCi) Tc-99m pertechnetate at acupuncture point K-3 in each foot, we had the patient lie supinely under the detector of the gamma camera. Lower body scans, from the pelvis to the feet, were then taken with the legs semi-flexed at 6 and 9 minutes after SC injection. At 12 and 15 minutes, another 2 lower body scans were taken with the legs straight.

Interpretation of the SC-RNV was done by both the authors. The intra-observer variation found by the 1st author was 1.1% (2/179, data unpublished), and the inter-observer difference between the 2 authors' findings was 3.4% (6/179, data unpublished). Normally, in SC-RNV there are deep venous images...
Fig. 1 Normal SC-RNV of a 38 year-old woman. A. At 6 minutes after subcutaneous injection of Tc-99m pertechnetate at acupuncture points K-3, there is smooth deep venous drainage in the lower limbs, through the calf deep veins (dv), the popliteal vein (pv), the superficial femoral vein (sv) and common femoral vein (cv). Superficial venous drainage is not visible normally. B. At 9 minutes after injection is almost the same pattern of deep venous drainage. By 12 minutes (C) and 15 minutes (D), the deep venous drainage may persist, or become a little faint.

at 6 minutes and 9 minutes, as shown in Fig. 1. Besides, the soft tissue of the lower extremities is demarcated to tell the anatomical location of the deep veins. At 12 minutes and 15 minutes, the deep venous drainage may become less prominent than the earlier phases, i.e. at 6 minutes and 9 minutes, and occasionally there may be evidence of some superficial venous drainage in the calf or calves. In cases with DVT, the affected limb usually shows impaired deep venous drainage, i.e. poor or non-visualization of the deep vein(s), and prominent collateral superficial venous drainage, as shown in Figs. 4, 6, 7 and 8.

Duplex ultrasound (duplex US)
An ACUSON 128 computed ultrasound scanner was used for duplex US, with an L-538 transducer. When the patient lay supinely, the transducer was put over the common femoral vein (CFV), the superficial femoral vein (SFV) and the popliteal vein (pop. v) to record the duplex US. Normally, by color Doppler US the deep vein is demonstrated as a complete band encoded with blue color (Fig. 2a). With Doppler signal recording, the deep venous flow undergoes normal phasic change with respiration, i.e. increased flow rate in the expiratory phase and decreased flow rate in the inspiratory phase, as shown in Fig. 2b. On proximal compression there is cessation of deep venous flow, and after release of the compression there is transient augmentation of the flow rate (Fig. 2c). And on squeezing, i.e. distal compression, there is initial augmentation followed by cessation of the flow (Fig. 2d). By color Doppler US, in cases with DVT, usually there is low or poor echogenic thrombus instead of color flow in the deep vein(s) (Fig. 3a to 3c); or, in cases of DVT with recanalization there is partial thrombus in the deep vein(s) having segmented color-coded flow that is antegrade or reversed (Fig. 5).

Grouping of patients
Since duplex US is a reliable diagnostic tool for the diagnosis of DVT, the patients were classified into 3 groups according to the findings of duplex US. The diagnostic criteria of duplex US for DVT are: (1) echogenic substance(s) in the deep veins with no Doppler signal, (2) loss of phasic change in deep venous flow with respiration, and (3) loss of normal response of deep venous flow and caliber to compression.

Group 1 (normal): Twenty-six patients were considered normal, since no evidence of DVT was obtained by duplex US. They were 11 males and 15 females, aged 28 to 77 years.

Group 2 (unilateral DVT): Nineteen patients (8 males, 11 females) had unilateral DVT (involving CFV, SFV and/or pop. v) by duplex US, aged 23 to 77 years.

Group 3 (bilateral DVT): Twelve patients (5 males, 7 females, aged 28 to 69 years) had bilateral DVT by duplex US.

Among the patients in groups 2 and 3, 19 (61%,
Fig. 2  Normal duplex US of a 38-year-old female. a. In the normal color flow map, the left common femoral artery (CFA) was encoded with orange-red color, and the left common femoral vein (CFV) was encoded with green-blue color. b. The Doppler signal of the left CFV showed phasic change of blood flow rate with respiration normally, that is lower rate at inspiration and higher rate at expiration. c. With proximal compression of the left CFV, the blood flow was suddenly interrupted (white arrow heads); and after release of the compression, there was normally transient augmentation of blood flow (white arrow). d. On distal compression (squeeze), there was initially augmentation (white arrow) followed by transient decrement/cessation (white arrow heads) of blood flow in a normal deep vein.

Fig. 3  Case 1: A 38-year-old male patient with right lower leg edema. The color Doppler ultrasound (a to c) revealed no visible color flow in the right common femoral vein (CFV) and superficial femoral vein (SFV); instead, there was a low echogenic thrombus (t) in those vessels. By duplex ultrasound (d) was no Doppler signal to be detected.
Fig. 4 Case 1. The SC-RNV revealed non-visualization of the right CFV, SFV, popliteal vein and calf deep veins (arrows) with prominent superficial collateral circulations.

Fig. 5 Case 2. A 67-year-old male patient with right leg edema. a. The color Doppler ultrasound revealed some echogenic thrombi (t, arrows) in the right SFV. b On Valsalva maneuver, there was reversed flow (white arrow heads, encoded with red color) through the SFV (white arrows), indicating partial recanalization. (t=thrombus)

Fig. 6 Case 2. The SC-RNV showed non-visualization of the deep veins of the right lower extremity (arrows) with prominent collateral circulation through the greater saphenous vein.

Fig. 7 Case 3. A 71-year-old female patient with bilateral lower limb edema. The SC-RNV revealed impaired deep venous drainage (arrows) through the right SFV, and left CFV, SFV, popliteal vein and calf deep veins.

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19/31) had also received contrast venography (CV) and/or x-ray computed tomography of the pelvis and thighs (XCT). The results of CV/XCT were compatible with those of duplex US.

RESULTS

Overall
All of the normal subjects (group 1) had a normal SC-RNV pattern of deep venous drainage (as shown in Fig. 1). All of those with unilateral DVT (group 2) had ipsilaterally impaired deep venous drainage (Figs. 4 and 6). All of those with bilateral DVT (group 3) showed bilaterally impaired deep venous drainage (as shown in Figs. 7 and 8).

Case report
Case 1 A 38-year-old man experienced right lower leg edema for 7 months following a traffic accident, and was hospitalized when the swollen leg became painful. A duplex US revealed echogenic thrombi in the right CFV and SFV with no Doppler signal to be detected (Fig. 3). The result of SC-RNV showed non-visualization of deep venous flow through the right deep calf veins, pop. v, SFV and CFV. Instead, there was prominently superficial collateral circulation in the right lower extremity (Fig. 4). The deep venous drainage of the left lower extremity looked normal.

Case 2 A 67-year-old diabetic male patient suffered from right leg edema for 2 weeks before admission. A duplex US demonstrated deep vein thrombi in the right SFV, and on Valsalva maneuver the flow was reversed, indicating partial recanalization (Fig. 5).

On SC-RNV, there was no deep venous flow through the right calf deep veins, pop. v, SFV and CFV. But collateral flow through the right greater saphenous vein was evident. Deep venous drainage was normal on the left side (Fig. 6).

Case 3 A 71-year-old female, having uterine cervical carcinoma noticed for years and undergoing radiotherapy, suffered from edema of the bilateral lower extremities for 4 months intermittently. A duplex US showed DVT in the left CFV/SFV and the right SFV. On SC-RNV, there were bilateral DVTs with prominent superficial collateral circulation (Fig. 7).

Case 4 A 63-year-old male CVA (cerebro vascular accident) patient suffered from painful right leg edema and left calf soreness for a couple of days. A duplex US revealed DVT in the right SFV and the left pop. v. The SC-RNV demonstrated impaired deep venous drainage through the right pop. v and SFV, and through the left calf deep veins and pop. v. (Fig. 8).

DISCUSSION

DVT involving the lower extremity remains a severe clinical problem, as it may predispose to pulmonary embolism. Diagnosis should therefore be done early, before its proximal propagation or even the severe complication, pulmonary embolism, occurs. Contrast venography is currently considered the standard method of diagnosis. However, it is invasive and expensive, often uncomfortable, not uniformly available, and not without morbidity.

In the past decade, some non-invasive assessments
have been developed and are gaining more and more importance.\textsuperscript{1-6,19-26} Among these, SC-RNV is the most recent one, and was developed in our clinical laboratory.\textsuperscript{1,2} Compared with the ascending radio-
nuclide venography (IV-RNV), SC-RNV possesses several advantages. First, in SC-RNV the radio-
nuclide absorbed is finally distributed into the soft tissue of whole body to be an anatomical landmark
telling the deep veins from the superficial veins; thus, a false negative result from misreading a super-
ficial as a deep vein can be prevented. The latter is not uncommon in IV-RNV, since the radionuclide
utilized in IV-RNV is usually Tc-99m MAA (macro-
aggregated albumin) that after injection into a pedal vein accumulates in the lungs, and thereby the soft
tissue of the body cannot be contured as the landmark. Second, since in IV-RNV there is usually a
higher concentration of radionuclide within the deep vein(s), a vessel having a thrombus with recanalization
could be visualized as normal patency. Pre-
nviously such an experience was not unusual in our
routine work. By mean of SC-RNV, it is easy to
prevent false reading in such a case, because the
diseased vein is demonstrated as a faintly or non-
visualized vessel accompanied with prominent col-
lateral superficial venous drainage, as shown in
Fig. 6 (case 2). Finally, on IV-RNV the procedure
may be very time-consuming, esp. in cases of mod-
erate-to-severe foot edema; and, occasionally there
is failure on IV injection.\textsuperscript{1}

In the present study, the clinical availability of
SC-RNV was settled. It was as sensitive and accurate
as duplex US for the diagnosis of DVT. All of those
with US evidence of DVT had a positive SC-RNV
result (group 2 and group 3), and all of those without
US evidence of DVT had a normal result. However,
duplex US is a complicated technique, and is more
technique-dependent and less time-saving than SC-
RNV. By duplex US, the vein(s) could be visualized
only segmentally; and, for most sonographers, it is
difficult to observe the entire course of the deep
veins of the lower limbs. In fact, duplex US is efficaciously in the diagnosis of proximal (i.e. above-knee)
DVT, while its role in detection of distal (i.e. below-
knee) DVT is uncertain.\textsuperscript{24,27} During the last 2 years,
we set up a qualified duplex US technique for the
diagnosis of proximal DVT, but have frequently been
unable to tell a distal DVT by duplex US only. In the
present study, among the 6 cases in which the SC-
RNV revealed evidence of combined proximal and

\textbf{Fig. 9} A case of pyogenic arthritis of knee and calf cellulitis compressing the deep vein
(a 68-year-old man). The SC-RNV on Jun. 6, 1990 revealed abnormally soft tissue accumula-
tion of radioactivity in the left knee (long arrows), that extended into the left upper calf
region (short arrows) on Oct. 30 the same year, after inadequate treatment. However, the deep
venous drainage of the left lower extremity remained normal (small arrows).
distal DVTs, duplex US revealed only proximal DVT in 4. Nevertheless, distal DVT is seldom a critical clinical problem;\textsuperscript{10–12} and in the present study, none of the cases referred to us showed evidence of simple distal DVT in SC-RNV, duplex US, CV and/or XCT.

Another problem is whether SC-RNV could distinguish impaired deep venous drainage due to DVT from venous disturbance due to external compression. Two of our recent cases with lower leg edema may offer an answer; one was a case of left leg edema from pyogenic arthritis and calf cellulitis compressing the left popliteal vein (Figs. 9 and 10), and the other was a case of left common femoral arterial pseudo-aneurysm compressing the left CFV and causing edematous swelling of the left lower

![Fig. 10](image)

**Fig. 10** The same case as in Fig. 9. The real-time ultrasound revealed a focus of low and heterogeneous echogenicity (arrows) in the joint space of the left knee (a: longitudinal scan, b: transverse scan). In the calf region was the other focus of the same echogenic characteristics (c: longitudinal scan), indicating cellulitis.

![Fig. 11](image)

**Fig. 11** A case of pseudo-aneurysm of the left common femoral artery (a 63-year-old woman). The SC-RNV revealed soft tissue swelling, impaired deep venous drainage (arrows), and abnormal accumulation of radioactivity at the groin region (arrow-heads) of the left lower extremity. The abnormal accumulation at the groin region is not a sign of DVT, and is indicative of some vascular mass compressing the left common femoral vein.
Fig. 12 The same case as in Fig. 11. A real-time ultrasound of the left groin region revealed a vascular mass with lumen (I) and circumferential thrombus (II), that by contrast angiography was a pseudo-aneurism of the left common femoral artery.

extremity (Figs. 11 and 12). A SC-RNV of the former one showed increased soft tissue radioactivity at the left knee joint and in the upper left calf region, and there was smooth deep venous drainage both proximal and distal to the left knee (Fig. 9); a real-time US revealed intra-articular fluid collection in the left knee and hypoechoic necrotizing material in the left calf muscle layers (Fig. 10). In the latter case, a SC-RNV demonstrated soft tissue swelling of the left lower extremity, interruption of deep venous drainage through the left popliteal vein and the left SFV with collateral circulation via the greater saphenous vein, and extra-vascular accumulation of radionuclide in the soft tissue of the left groin region (Fig. 11); a duplex US revealed aneurysmal dilatation of the left common femoral artery (CFA) and no evidence of DVT in the left CFV or SFV. Besides, in real-time US there was tortuous and segmentally dilated CFA with a thrombus formation in the latter case (Fig. 12). The major distinguishing point is the extravascular accumulation of radioactivity in the soft tissue around the compressed deep veins in the non-DVT cases. However, there still may be few cases showing impaired deep venous drainage with a cause undetermined by SC-RNV. In such a situation, the duplex US could afford a certain aid for causative diagnosis, although it is undoubtedly more time-consuming.

Our conclusion is that SC-RNV is clinically reliable in the diagnosis of DVT. Since it is simple to perform, less technique-dependent and more time-saving, we recommend it as one of the most available, non-invasive methods for routine use in the diagnosis of DVT.

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REFERENCES


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