CASE REPORT

Using ultrasonography in evaluating the intramuscular injection techniques used for administering drug treatments to schizophrenic patients in Japan

Yuko Yasuhara¹, Eri Hirai², Sakiko Sakamaki², Tetsuya Tanioka¹, Kazushi Motoki³, Kensaku Takase¹, Rozzano Locsin⁵, Chiemi Kawanishi⁶, Tatsuya Inui⁷, Chie Watari⁸, and Kouichi Makiguchi⁸

¹Department of Nursing Management, Institute of Health Biosciences, the University of Tokushima Graduate School, Tokushima, Japan, ²The Major in Nursing, School of Health Sciences, the University of Tokushima, Japan; ³Department of Clinical Laboratory, and ⁴Department of Cerebral Surgery, Tokushima Prefectural Central Hospital, Tokushima, Japan, ⁵Department of Nursing, Christine E. Lynn College of Nursing, Florida Atlantic University, USA, ⁶Department of Nursing Art and Science, Institute of Health Biosciences, the University of Tokushima Graduate School, Tokushima, Japan, ⁷Department of Psychiatry, Fujii Hospital, Tokushima, Japan, ⁸Department of Psychiatry, Fujishiro Kensei Hospital, Aomori, Japan

Abstract: This study was conducted with six patients with schizophrenia, four of whom received the atypical antipsychotic risperidone long-acting injectable (RLAI), and two patients receiving the typical depot injection (TDI). The purpose of this study was to determine the location (gluteus medius or maximus; deltoid muscles) and diffusion of typical and atypical antipsychotic medications administered intramuscularly using ultrasonography. When using the standardized depth of needle insertion, in some cases, the drug was injected into the gluteus maximus instead of the gluteus medius. Similarly, in some cases the TDI was not visible in the ultrasonographic images until sixteen days after the injection. This verifies how hard the injection site becomes when microspheres of RLAI is injected as compared to other muscle areas. These results confirmed that the gluteus muscle structure was the ideal muscle for depot injection as evidenced by the injection solution being dispersed and rendered not visible immediately after intramuscular injection (IM). With the use of ultrasonography, injection sites and drug dispersions were evaluated under a direct visual guidance, suggesting that ultrasonography is a useful method for establishing evidence for determining correct insertion of IM injection, diffusion of medications, and the effective administration of IM injections. J. Med. Invest. 59: 213-219, February, 2012

Keywords: intramuscular injection, ultrasonography, atypical antipsychotic risperidone long-acting injectable, typical antipsychotic depot intramuscular injection

Received for publication November 30, 2011; accepted December 27, 2011.
Address correspondence and reprint requests to Yuko Yasuhara, Department of Nursing Management, Institute of Health Biosciences, the University of Tokushima Graduate School, Kuramotcho, Tokushima 770-8509, Japan and Fax: +81-88-633-7629.
The medication for patients with schizophrenic can be classified into two: oral drugs and long-acting intramuscular (IM) injections. Oral drugs are easy to take, however patients may easily forget to take them or may choose to discontinue them. In the course of drug treatment, even a short period of partial adherence could increase the risk of the occurrence of relapse.

On the other hand, a long-acting IM injection is accompanied by pain from injection, but the medicinal effects can last for about two weeks (risperidone long-acting injectable) or four weeks (fluphenazine decanoate and haloperidol decanoate). Since with a long-acting injection, definite adherence can be expected as long as an injection is given, it is reported to be more effective in preventing relapses than oral drugs (1, 2).

Typical depot agents are oil preparations and possess the property of being gradually hydrolyzed by esterase in vivo (3). The ester body is administered dissolved in oil. However, these are known to frequently cause injection site reactions. For this reason, a technique to encapsulate the IM drug solution using the Z track method was introduced as a means to prevent injection site reactions (4). However, little evidence for its efficacy has been demonstrated (5, 6). Moreover, there have been concerns about oil preparations that are not properly injected into the right muscles, causing stronger injection site reactions.

The long-acting injectable atypical antipsychotic, risperidone long-acting injectable (RLAI), on the other hand, does not have a hydroxyl group, hence, it could not be made slow-release using long-chain fatty acids. It became possible then to administer RLAI as a soluble suspension after making it slow-release through a technology utilizing microspheres (7, 8). Moreover, it is known to cause little injection site reactions (9). That is, with RLAI, pain or swelling after an injection may be milder than after an injection of a typical depot injection (TDI).

Both TDI, fluphenazine decanoate and haloperidol decanoate, have side effects. Some advantages atypical antipsychotics have over typical antipsychotics are that there are fewer anticholinergic side effects, less parkinsonian and dystonia side effects, and these also suppress negative symptoms including a lower propensity for causing extrapyramidal side effects. The medication in RLAI is enclosed in tiny beads called “microspheres”. After these are injected into the muscle, they slowly dissolve, releasing a constant amount of the risperidone medication. 

RLAI became available in 2009, possessing the advantage of being an atypical antipsychotics and depot agents using, “microspheres.”

IM injections need to be accurately administered. Factors such as the patient’s unique build and sebum thickness need to be carefully considered. Therefore, when an IM injection is given, the depth of needle insertion is left to the assessment of the nurse. Until now, there have been no studies to confirm whether the drug administered was actually injected into the right muscles. Furthermore, no studies have been found that illustrated how the injection solution is absorbed in the muscle and about the differences in absorption between different drugs. By ensuring that drugs are successfully administered through IM injection, the effect of medications can be achieved, and may lead to improvement of patients’ QOL. Moreover, investigating the differences in drug absorption of muscles can also guide in administering IM injection.

Accordingly, in this paper the following ultrasonographic results derived from six cases provided answers to: 1) whether the injection solution was actually infused into the muscle; 2) the pharmacokinetics of the solution after the injection; 3) differences in absorption images between RLAI (water-soluble suspension) and TDI; and 4) hardness of the muscular system using elastography.

METHODS

Participants

Six patients with schizophrenia from two hospitals (in western and east Japan) joined the study. Four of the six schizophrenic patients were treated with RLAI while two were treated with TDI. Five patients had their treatments injected into their gluteus muscle while one patient received the TDI into his deltoid muscle.

Date collection

The study was conducted for a period of eight months (June 2010 to January 2011). Body weight and height were measured and body mass index (BMI) was calculated for all patients. Just before RLAI injection, the distance from epidermis to underfascia (DEUF), distance from epidermis to ilium (DEI) at bilateral gluteal sites, and the most recent RLAI injections were assessed through ultrasonography. Two patients treated with TDI were immediately evaluated after their injection as well as 16 days
thereafter. Additional measurements were taken for (a) the distance from epidermis to upper-arm bone and (b) the distance from epidermis to fascia of deltoid muscle. All ultrasonographic measurements were performed by an experienced sonographer using a 7.5 MHz linear and convex array transducer.

Ultrasonographic images were based on the dorsogluteal injection site. DEUF and DEI measurements were made above and outside a line drawn from the posterior superior iliac spine to the greater trochanter of the femur. Gluteus maximus, medius, and minimus muscles were used as common IM injection regions.

Elastography is a quantitative approach for imaging linear elastic properties of tissues in order to detect suspicious tumors (10). In this study, elastography was used to visualize the tension of muscle tissues after injection (11). When it was difficult to determine the boundaries between the fascia or subcutaneous tissues and muscles, the sonographer applied pressure on the epidermis to identify the boundaries through ultrasonography. In addition, the patients were requested to clench their buttocks for the sonographer to identify measurement sites while observing the movement of the muscles and subcutaneous tissues.

One particular nurse who had 15 years of experience of administering IM injection reported that they identified the measurement site on the gluteal muscles by using the “four-and three-way split” method (12, 13), a method they perceived to be valid and reliable (Fig. 1). Moreover, the nurse noted that the injection site should not be at an uneven surface of the skin. Ultrasonographic evaluation and neurosurgical expertise were also done by two experienced nurses (23 and 30 years, respectively). The results of the measurements were based on the recorded image and after due discussion among all the data gatherers (nurses, physicians, and sonographers).

Date analysis

The ultrasonographic images of the six cases were evaluated for the following: 1) whether the solution was administered into the gluteus medius or the deltoid muscle; 2) absorption of the injection solution; 3) differences in absorption images between RLAI and the TDI; and 4) hardness of the muscular system using elastography. In RLAI injection, inserted needle length was calculated through the total needle length subtracted by the remaining needle length visible on the surface of the buttocks. Further evaluation was then made through ultrasonography.

Ethical considerations

We conducted this study with the approval of the ethics committees of the Tokushima University Hospital, Tokushima Prefectural Central Hospital, and Fujishiro Kensei Hospital. Verbal and written informed consents were given by the six study participants.

RESULTS

Case 1: It was confirmed through ultrasonography that RLAI was injected and diffused into the gluteus medius (Fig. 2). Nurses inserted the needle while checking the needle position through echo images. From the epidermis to the fascia was 13
mm. When the needle passed through the epidermis and the fascia, RLAI was injected while still monitoring the position of the needle tip. The depth of needle insertion was 46 mm.

Case 2: This ultrasonographic image shows that immediately after RLAI had been injected into the gluteus maximus muscle instead of the gluteus medius (Fig. 3). The encircled part shows that RLAI was injected into the gluteus muscle. From the epidermis to the fascia of the gluteus muscle was 11.1 mm.

Case 3: RLAI was administered to the patient two weeks ago. The length of the RLAI needle inserted was 32 mm. The distance from the epidermis to the fascia was 18.19 mm. The penetration depth was 21.66 mm. The encircled part shows that RLAI was injected into the gluteus maximus muscle. The needle was considered to be obliquely-inserted at an angle against the skin surface. In this image, the injected RLAI seemed like an echogenic mass in the gluteus maximus muscle (Fig. 4).

Case 4: With the evaluation immediately after the RLAI injection using elastography, there were differences in the hardness between the areas around the injection site and the other muscle sites. The left part of the elastographic image below illustrates the muscle tissues. The blue portions show the hardening of muscle tissue (Fig. 5).

Case 5: The pharmacokinetic evaluation of the solution two weeks after the injection confirmed the actual infusion of the RLAI into the gluteus medius. (Fig. 6) It was able to confirm that the assessment
made on the depth of needle during insertion by
the nurse who performed the IM injection was ap-
propriate.

Case 6: On the other hand, when the TDI was
taken immediately after the injection, the echogenic
image did not appear in the ultrasonographic image
(Fig. 7). The distance from epidermis to fascia of
muscle was 9.1 mm, and distance from epidermis
to the ilium measured 59.9 mm.

Case 7: The ultrasonographic images also showed
cyst-like (peanut-shaped) images for the TDI
(fluphenazine decanoate) sixteen days after the in-
jection, and little diffusion was observed (Fig. 8).

DISCUSSION

It has been commonly believed that an injection
site located using the upper outer quadrant method
should have the gluteus medius below the subcuta-
neous tissues; however in some cases in this study,
the gluteus maximus overlaid that area. It is said
that in about 30% of autopsied bodies, the gluteus
maximus overlies the gluteus medius as observed
in the cases in the present study (14). From the
viewpoint of IM injection, the same efficacy can be
expected from the injection into the gluteus maxi-
mus, however the gluteus medius is believed to be
thicker than the gluteus maximus, allowing safer
and more accurate absorption of the drug solution.
For this reason, when giving an injection in such
cases, we need to insert the needle into the gluteus
medius while taking into account the thickness of
the gluteus maximus.

In this study, the examiners confirmed the depth
of the needle insertion by putting a mark on the
needle with a pen. In some cases the needle still
did not reach the gluteus medius, even though it
was inserted over 10 mm deeper than the depth to
the fascia. These cases are believed to have been
caused by the position of the nurse upon adminis-
tering the injection. The needle might have been
inserted at an angle other than 90 degrees and did
not reach the gluteus medius. Furthermore, in some
cases, the injection solution was seen flowing be-
tween the gluteus medius and gluteus maximus,
and there might have been many such IM injection
cases in the past.

We gave injections while checking the ultra-
sonographic images. It was considered that there
would be limitations to the assessment of the three-
dimensional IM structure through the two-dimen-
sional images, so recognizing these limitations may
be important for giving IM injections safely.

The elastographic examination indicated differ-
ences in the hardness between the areas around the
injection site and other muscle sites and the injected
muscle was considered harder compared to the rest
of the area. This verifies how hard the injection site
becomes when microspheres of RLAI are injected
into muscle areas. Since we only focused on this
particular aspect, further investigation needs to be
done regarding the effect of other IM injections to
muscle hardness.

Also, two weeks after the injection, it was con-
firmcd that the RLAI remained inside the but-
tock muscles with its microspheres intact and was
diffused across a wide area. From these results it became clear that with RLAI it is easy to check the actual IM injection and diffusion using ultrasonography.

On the other hand, when the image was taken immediately after the TDI the absorption did not appear in the ultrasonographic images, perhaps due to the composition of the drug a difference between water-soluble and oil-based solutions. However, cyst-like (peanut-shaped) ultrasonographic absorption images can be seen 16 days after. The peanut-sized cyst-like image seen was believed to be due to the hardness of the muscles as a resulting effect of the ability of oily TDI medication absorption. This muscle hardening was also confirmed by the patient.

CONCLUSION

This study compared two cohorts of patients with schizophrenia treated with different types of RLAI and TDI injected into the gluteus and deltoid muscles through ultrasonography. When the standardized depth of needle was inserted following the standard procedure, the drug was injected into the gluteus maximus instead of the gluteus medius in some cases. However, the TDI did not appear in the ultrasonographic images immediately after injections, rather sixteen days after. From these results, it was affirmed that the buttock muscle structures are the ideal areas for RLAI injection as evident in the ultrasonographic images indicating appropriate dispersion of medication within this sites.

With the use of ultrasonography, injection sites and drug diffusion can be checked under direct visual guidance. Therefore, it was suggested that ultrasonography could be a very useful tool in establishing evidence for safe administration of IM injections.

This study was supported by a research grant-in-aid from the Ministry of Education, Culture, Sports, Scientific and Technology research of Japan (No. 23660100).

CONFLICT OF INTEREST

None of the authors have any conflicts of interest to declare.

REFERENCES

11. Niitsu M, Michizaki A, Endo A, Takei H, Yanagisawa O: Muscle hardness measurement by using ultrasound elastography, a feasibility...