Vascular Patterns in Cutaneous Ulcerated Basal Cell Carcinoma: A Retrospective Blinded Study Including Dermoscopy

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The aim of this retrospective study was to determine the type and prevalence of vascular patterns in the ulcerated and non-ulcerated portions of histologically proven basal cell carcinomas (BCCs) and correlate them with other dermoscopic and clinical features, including the clinically supposed diagnosis. Three authors retrospectively collected 156 clinical and 156 dermoscopic digital images of ulcerated BCCs (histologically confirmed); each image was blindly evaluated by 2 other authors, who did not know the histological diagnosis. Seventeen lesions were completely ulcerated, while 139 lesions presented ulcerated and non-ulcerated portions. Correct clinical diagnosis was associated with the type of lesion, in particular 90.6% of partially ulcerated lesions were correctly diagnosed with clinical-dermoscopic examination, compared with 11.8% of totally ulcerated lesions ($\chi^2 = 64.00$, p = 0.000). Presence of arborizing pattern in the ulcerated portion was associated with a correct diagnosis (Fisher's exact test, p = 0.015). Correct diagnosis was also associated with absence of dotted pattern in the non-ulcerated area ($\chi^2 = 16.18$, p = 0.000); the absence of hairpin ($\chi^2 = 6.08$, p = 0.000) and glomerular patterns were associated with correct diagnosis in the ulcerated areas ($\chi^2 = 18.64$, p = 0.000). In case of completely ulcerated BCC the clinician lacks the means to correctly identify the correct nature of the lesion, and is driven towards an incorrect diagnostic conclusion.

Key words: dermoscopy; basal cell carcinoma; melanoma; vascular pattern; ulcerated basal cell carcinoma.

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Basal cell carcinoma (BCC) is defined as a slowgrowing skin malignancy, predominantly affecting middle-aged and fair-skinned individuals. Clinically BCC can manifest differently, with nodular, superficial, morpheic, and pigmented variants (1).

Dermoscopy is an *in vivo* technique that has proven useful to differentiate BCC from other cutaneous malignancies, such as squamous cell carcinoma (SCC) and melanoma (2–5).

Characteristic dermoscopic features of BCC include large blue-grey ovoid nests, leaf-like areas, multiple blue-grey globules and spoke-wheel areas (6, 7). In addition, specific vascular patterns may aid the dermoscopic diagnosis of BCC, especially when the above-mentioned traditional criteria are lacking (8–11). Arborizing vessels, in particular, are known to be a characteristic and common feature of BCC, with a positive predictive value of 94.1% (9–12). Moreover, short fine telangiectasias are commonly observed in superficial BCCs (13, 14). However, several additional morphological types of vessels have been reported (11–15).

The aim of our study was to determine the type and prevalence of vascular patterns in the ulcerated and non-ulcerated portions of histologically proven BCCs and correlate them with other dermoscopic and clinical features, including the clinically supposed diagnosis.

MATERIALS AND METHODS

Three of the authors (DB, NA, GG) retrospectively collected 156 clinical and 156 dermoscopic images of histologically confirmed ulcerated BCCs, developed by 153 patients treated at the Dermatology Unit of University of Bari Policlinico hospital between January 2011 and December 2013. Anagraphical data (such as age and sex) and lesion-related data (such as presence of pigmentation, lesion anatomical site and diameter) were recorded.

Dermoscopic images had been taken via digital dermoscopy (VideoCAPTM, DS Medica, Milano, Italy) and polarized handheld dermoscopy devices (DermoGenius[®] II, DermoScan GmbH, Regensburg, Germany and DermLite DL3N, 3 Gen LLC, San Juan Capistrano, CA, USA) mounted on a digital camera (10× magnification). Clinical images had been taken via videoCAPTM or digital camera. Skin preparation prior to dermoscopy included hyperkeratotic skin removal by wet gauze and application of petrolatum oil in case of immersion dermoscopy. Each digital image was blindly evaluated by 2 of the authors (AF, MV), who did not know the histological diagnosis, for the presence of the following types of vessels, diversified for the ulcerated and non-ulcerated portions of the lesions: arborizing, dotted, linear-irregular, comma, hairpin, crown, glomerular. In the presence of 3 or more types of vessels, the vascular pattern was defined as polymorph. Diffuse reddish coloration (milky-red globules/areas and erythema) was also considered. Moreover, the prevalent vascular pattern for each lesion was noted. Other dermoscopic features were assessed: large blue-grey ovoid nests, leaf-like areas, multiple blue-grey globules and spoke-wheel areas (classic BCC patterns); brown-to-black dots/globules, pseudopods, blue/white veil, pigment network and peppering (classic melanoma patterns). Pigmented BCCs were defined by the presence of dermoscopic pigmentation on more than 25% of the lesion's surface.

On the basis of the above characteristics a clinical-dermoscopic diagnosis was advanced by the 2 blinded authors.

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Table I. Characteristics of all the 156 considered lesions, developed on 153 patients, assessed retrospectively

| Clinical features | | |
|------------------------------------|-------------|--|
| Sex, n (%) | | |
| Female | 74 (48.40) | |
| Male | 79 (51.60) | |
| Age, years, mean±SD | 73.67±11.3 | |
| Range | 47-98 | |
| Pigmented lesion, n (%) | | |
| Yes | 42 (26.90) | |
| No | 114 (73.10) | |
| Lesion site | | |
| Body | 60 (38.50) | |
| Face | 56 (35.90) | |
| Lower limbs | 11 (7.10) | |
| Upper limbs | 12 (7.70) | |
| Scalp | 17 (10.90) | |
| Lesion diameter, mm, mean \pm SD | 20.16±17.76 | |
| Range | 3-182 | |

SD: standard deviation.

Statistical analysis

To compare the prevalence of each vascular pattern between ulcerated and non-ulcerated portion of the lesions the McNemar test was performed, excluding from the analysis the 17 lesions that were completely ulcerated. Associations between the vascular patterns and other dermoscopic and clinical features were evaluated by χ^2 test. To compare the prevalence of single patterns, as well as dermoscopic and clinical features between correctly diagnosed lesions and not, χ^2 test was also employed. A logistic regression was performed to identify, by a multivariate analysis, which pattern or feature led to a correct diagnosis.

RESULTS

A total of 156 clinical and 156 dermoscopic images of ulcerated BCCs, which had developed on 153 patients, were assessed. Characteristics of lesions and patients are shown in Table I. Seventeen lesions were completely ulcerated, while 139 lesions presented ulcerated and non-ulcerated portions.

Partially ulcerated lesions

The prevalence of the discrete vascular patterns in the 139 lesions with both ulcerated and non-ulcerated areas Table II. Pattern prevalence comparison between ulcerated and non-ulcerated portion in the 139 partially ulcerated lesions

| Vascular pattern | Ulcerated portion % | Non-ulcerated portion % | <i>p</i> -value |
|----------------------------|---------------------------|-------------------------------|-----------------|
| Arborizing | 16.5 | 83.5 | 0.000 |
| Dotted | 82.7 | 8.6 | 0.000 |
| Linear-irregular | 55.4 | 25.9 | 0.000 |
| Comma | 16.5 | 1.4 | 0.000 |
| Diffuse reddish coloration | 0 | 12.9 | - |
| Hairpin | 5.8 | 0.7 | 0.039 |
| Glomerular | 4.3 | 1.4 | 0.219 |
| Crown | 0.7 | 0 | - |
| Polymorph | 28.1 | 2.2 | 0.000 |

Significant values are given in bold.

is shown in Table II and Fig. 1. A statistically significant difference in the prevalence of the arborizing, dotted, linear-irregular, comma, hairpin, and polymorph patterns between the ulcerated and non-ulcerated areas was noted.

In the ulcerated portion the comma pattern was associated with the presence of pigmentation ($\chi^2 = 11.137$, p=0.001) and with the presence of large blue-grey ovoid nests ($\chi^2 = 4.91$, p = 0.027). In the non-ulcerated portion, diffuse reddish coloration was associated with the presence of leaf-like areas ($\chi^2 = 8.29$, p = 0.004); and the presence of blue/white veil was associated with linearirregular pattern (χ^2 =4.41, p=0.036) and with diffuse reddish coloration ($\chi^2 = 8.47$, p = 0.004). The absence of blue/white veil was associated with arborizing pattern $(\chi^2 = 10.6, p = 0.001)$. An inverse association was noted between the presence of hairpin pattern and the presence of at least one of the classic BCC patterns; in fact 36.4% of lesions with hairpin pattern had at least one of the classic BCC patterns vs. 67.6% of lesions without hairpin pattern presenting at least one of these typical patterns $(\chi^2 = 4.4, p = 0.036).$

As expected, the classic dermoscopic criteria of BCC were highly associated with a correct clinical diagnosis in the 139 partially ulcerated lesions: ovoid nest (χ^2 =4.59, p=0.032), blue-gray globules ($\chi^2=4.95, p=0.026$), leaflike ($\chi^2 = 7.83$, p = 0.005) and arborizing vascular pattern $(\chi^2 = 9.1, p = 0.003)$. Ninety-four percent of lesions with at



Fig. 1. Pattern distribution in the 139 lesion with an ulcerated and non-ulcerated portion. *McNemar test.

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least one of these classic criteria of BCC were clinically diagnosed correctly, compared with 61.1% of lesions without these criteria ($\chi^2=24.59$, p=0.000).

Presence of arborizing pattern in the ulcerated portion was associated with a correct diagnosis (Fisher's exact test, p=0.015). One-hundred percent of lesions with arborizing pattern were correctly diagnosed vs. 78.9% of lesions without that pattern. A similar association was evident for the non-ulcerated portion (Fisher's exact test, p=0.008) (Fig. 2).

Correct diagnosis was also associated with absence of the dotted pattern in the non-ulcerated area ($\chi^2=16.18$, p=0.000); and absence of hairpin ($\chi^2=6.08$, p=0.000) and glomerular patterns were associated with correct diagnosis in the ulcerated areas ($\chi^2=18.64$, p=0.000).

Logistic regression analysis, performed on the 139 lesions with ulcerated and non-ulcerated portions, also revealed that the vascular features influencing the correct diagnosis most were: presence of arborizing pattern in the ulcerated portion; absence of dotted pattern in the nonulcerated areas, and absence of hairpin and glomerular patterns in the ulcerated areas, which are those commonly associated with melanoma and/or SCC.

Lesion size was not associated with any of the factors studied, while there was a significant association between the presence of spoke-wheels and the upper limb site ($\chi^2=12.53$, p=0.014). No differences in the lesions' characteristics were observed between males and females.

Totally ulcerated lesion

Concerning the 17 totally ulcerated lesions, the dotted and linear-irregular dermoscopic patterns were the most frequent (88.2% and 70.6%, respectively). The comma and polymorph patterns occurred in \sim 35% of these lesions. Other patterns were rare (hairpin and glomerular) or non existing.

Fig. 3 describes the association between the clinical diagnosis and the prevalence of vascular patterns in the

17 totally ulcerated lesions. Ten out of 17 lesions were blindly diagnosed as malignant melanoma; 5 out of 17 as SCC; and 2 out of 17 as BCCs. In the group misdiagnosed as melanoma, 9 out of 10 lesions presented the dotted pattern, 4 the linear-irregular pattern, and 3 the polymorph pattern.

All lesions

Considering all 156 lesions, the correct clinical diagnosis was associated with the type of lesion (totally or partially ulcerated); in particular, 90.6% of partially ulcerated lesions were correctly diagnosed with clinical-dermoscopic examination, compared with 11.8% of totally ulcerated lesions correctly diagnosed (χ^2 =64.00, *p*=0.000).

DISCUSSION

This study confirms a significant correlation between the correct clinical diagnosis and known classic dermoscopic characteristics of BCC, such as arborizing vessels, large





blue-grev ovoid nests, leaf-like areas, multiple blue-grey globules, and spoke-wheel areas (2-12).

More interestingly, in the 139 lesions presenting both an ulcerated and non-ulcerated area, significant differences in the prevalence of certain vascular patterns were noted between the 2 areas. In particular, the dotted, linear-irregular, hairpin, comma and polymorph patterns were highly represented in the ulcerated areas, whereas the arborizing pattern was prevalent, as expected, in the non-ulcerated areas (Table II, Fig. 1).

Considering exclusively the 17 BCCs that were completely ulcerated, these lesions lacked the classic dermoscopic criteria of BCC (leaf-like, spoke-wheels, ovoid nest, blue-grey globules, arborizing vascular pattern), while showing a distinct prevalence of dotted, linearirregular and polymorph vascular patterns (Figs 3 and 4).

Of the lesions presenting both an ulcerated and a non-ulcerated portion, 90.6% were diagnosed correctly through clinical and dermoscopic examination (Fig. 4); conversely, only 11.8% of completely ulcerated lesions were diagnosed correctly (Fig. 5).

The results of this study raise concern about a high percentage of incorrect diagnosis in those lesions (histologically proven as BCCs) that were, both clinically and dermoscopically, blindly interpreted as different malignant neoplasms, melanoma in particular. In fact, in case of completely ulcerated BCCs lacking specific dermoscopic criteria, and presenting vascular patterns traditionally associated with melanoma and/or SCC (Fig. 5) (16, 17), the clinician simply lacks the means to correctly identify a BCC, and is driven towards an incorrect diagnosis of melanoma or SCC. This results in overdiagnosis of SCC/melanoma and in more urgent and radical surgical removal; this, of course, does not pose



Fig. 4. Handheld dermoscopy of a partially ulcerated basal cell carcinoma of the trunk in a 62-year-old man. Notice the arboriform pattern on the right (non-ulcerated portion) and the polymorph pattern (dotted, linear irregular, comma) on the left (ulcerated portion) (magnification x20).

any particular concern to the patient management (except perhaps increased patient psychological stress and unnecessary increased burden of urgent surgical removals), as the lesion will be ultimately excised. However, from a clinical and scientific aspect, this has long remained a grey area. To the best of our knowledge there are no reported data in the literature describing dermoscopic vascular patterns in ulcerated cutaneous BCC. Of note, our results pertain only to ulcerated histologically proven BCCs; we cannot assume that they apply to other ulcerated lesions of a different nature.

Study limitation

The limitation of this study is its retrospective nature; further controlled studies are warranted to confirm our preliminary results.

The authors declare no conflicts of interest.

REFERENCES

- 1. Roewert-Huber J, Lange-Asschenfeldt B, Stockfleth E, Kerl H. Epidemiology and etiology of basal cell carcinoma. Br J Dermatol 2007; 157: 47-51.
- 2. Altamura D, Menzies SW, Argenziano G, Zalaudek I, Soyer HP, Sera F, et al. Dermatoscopy of basal cell carcinoma: morphologic variability of global and local features and accuracy of diagnosis. J Am Acad Dermatol 2010; 62: 67-75.
- 3. Lallas A, Apalla Z, Argenziano G, Longo C, Moscarella E, Specchio F, et al. The dermatoscopic universe of basal cell carcinoma. Dermatol Pract Concept 2014; 4: 11-24.
- 4. Popadić M. Dermoscopic features in different morphologic types of basal cell carcinoma. Dermatol Surg 2014; 40: 725-732.
- 5. Seidenari S, Bellucci C, Bassoli S, Arginelli F, Magnoni C, Ponti G.High magnification digital dermoscopy of basal cell carcinoma: a single-centre study on 400 cases. Acta Derm Venereol 2014; 94: 677-682.
- 6. Popadić M. Statistical evaluation of dermoscopic features in basal cell carcinomas. Dermatol Surg 2014; 40: 718-724.
- 7. Puig S, Cecilia N, Malvehy J. Dermoscopic criteria and basal cell carcinoma. G Ital Dermatol Venereol 2012; 147: 135 - 140.
- 8. Argenziano G, Zalaudek I, Corona R, Sera F, Cicale L, Petrillo



Fig. 5. Digital dermoscopy image of a completely ulcerated basal cell carcinoma of the left forearm in an 80-year-old woman. Prevalent dotted pattern (magnification x10).

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G, et al. Vascular structures in skin tumors: a dermoscopy study. Arch Dermatol 2004; 140: 1485-1489.

- 9. Kreusch JF. Vascular patterns in skin tumors. Clin Dermatol 2002; 20: 248-254.
- 10. Micantonio T, Gulia A, Altobelli E, Di Cesare A, Fidanza R, Riitano A, et al. Vascular patterns in basal cell carcinoma. J Eur Acad Dermatol Venereol 2011; 25: 358-361.
- 11. Zalaudek I, Kreusch J, Giacomel J, Ferrara G, Catricalà C, Argenziano G. How to diagnose nonpigmented skin tumors: a review of vascular structures seen with dermoscopy: part II. Nonmelanocytic skin tumors. J Am Acad Dermatol 2010; 63: 377-386.
- 12. Menzies SW, Westerhoff K, Rabinovitz H, Kopf AW, McCarthy WH, Katz B. Surface microscopy of pigmented basal cell carcinoma. Arch Dermatol 2000; 136: 1012-1016.
- 13. Giacomel J, Zalaudek I. Dermoscopy of superficial basal cell

carcinoma. Dermatol Surg 2005; 31: 1710-1713.

- 14. Scalvenzi M, Lembo S, Francia MG, Balato A. Dermoscopic patterns of superficial basal cell carcinoma. Int J Dermatol 2008; 47: 1015-1018.
- 15. Sakakibara A, Kamijima M, Shibata S, Yasue S, Kono M, Tomita Y. Dermoscopic evaluation of vascular structures of various skin tumors in Japanese patients. J Dermatol 2010; 37: 316-322.
- 16. Warszawik-Hendzel O, Olszewska M, Maj M, Rakowska A, Czuwara J, Rudnicka L. Non-invasive diagnostic techniques in the diagnosis of squamous cell carcinoma. J Dermatol Case Rep 2015; 9: 89-97.
- 17. Chappuis P, Duru G, Marchal O, Girier P, Dalle S, Thomas L. Dermoscopy: a useful tool for general practitioners in melanoma screening: a nationwide survey. Br J Dermatol 2016; 175: 744-750.